# Package 'targeted'

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```
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Title Targeted Inference
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Description Various methods for targeted and semiparametric inference including
        augmented inverse probability weighted (AIPW) estimators for missing data and
        causal inference (Bang and Robins (2005) <doi:10.1111/j.1541-0420.2005.00377.x>),
     variable importance and conditional average treatment effects (CATE)
     (van der Laan (2006) <doi:10.2202/1557-4679.1008>),
        estimators for risk differences and relative risks (Richardson et al. (2017)
        <doi:10.1080/01621459.2016.1192546>), assumption lean inference for generalized
     linear model parameters (Vansteelandt et al. (2022) <doi:10.1111/rssb.12504>).
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```

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## Repository CRAN

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# **R** topics documented:

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	alean		_

## Description

AIPW for the mean (and linear projections of the EIF) with missing observations

alean 3

#### Usage

```
aipw(response_model, data, formula = ~1, missing_model, ...)
```

#### **Arguments**

response\_model Model for the response given covariates (ml\_model or formula)

data data.frame

formula design specifying the OLS estimator with outcome given by the EIF

missing\_model Optional missing\_model (ml\_model or formula). By default will use the same design as the response\_model.

... arguments to cate

### **Examples**

```
m <- lvm(y ~ x+z, r ~ x) distribution(m,~ r) <- binomial.lvm() transform(m, y0~r+y) <- function(x) { x[x[,1]==0,2] <- NA; x[,2] } d <- sim(m,1e3,seed=1) aipw(y0 ~ x, data=d)
```

alean

Assumption Lean inference for generalized linear model parameters

### Description

Assumption lean inference via cross-fitting (Double ML). See <doi:10.1111/rssb.12504

### Usage

```
alean(
  response_model,
  exposure_model,
  data,
  link = "identity",
  g_model,
  nfolds = 1,
  silent = FALSE,
  mc.cores,
  ...
)
```

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### **Arguments**

response_model	formula or ml_model object (formula => glm)
exposure_model	model for the exposure
data	data.frame
link	Link function (g)
g_model	Model for $E[g(Y A,W) W]$
nfolds	Number of folds
silent	supress all messages and progressbars
mc.cores	mc.cores Optional number of cores. parallel::mcmapply used instead of future
• • •	additional arguments to future.apply::future_mapply

#### **Details**

Let Y be the response variable, A the exposure and W covariates. The target parameter is:

$$\Psi(P) = \frac{E(Cov[A,g\{E(Y|A,W)\} \mid W])}{E\{Var(A \mid W)\}}$$

The response\_model is the model for E(Y|A,W), and exposure\_model is the model for E(A|W). link specifies g.

#### Value

alean.targeted object

### Author(s)

Klaus Kähler Holst

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ate

AIPW (doubly-robust) estimator for Average Treatement Effect

### Description

Augmented Inverse Probability Weighting estimator for the Average (Causal) Treatment Effect. All nuisance models are here parametric (glm). For a more general approach see the cate implementation. In this implementation the standard errors are correct even when the nuisance models are misspecified (the influence curve is calculated including the term coming from the parametric nuisance models). The estimate is consistent if either the propensity model or the outcome model / Q-model is correctly specified.

### Usage

```
ate(
  formula,
  data = parent.frame(),
  weights,
  offset,
  family = stats::gaussian(identity),
  nuisance = NULL,
  propensity = nuisance,
  all,
  labels = NULL,
  ...
)
```

### **Arguments**

formula Formula (see details below)

data data.frame

weights optional frequency weights

offset optional offset (character or vector). can also be specified in the formula.

family Exponential family argument for outcome model

nuisance outcome regression formula (Q-model)

propensity propensity model formula

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all If TRUE all standard errors are calculated (default TRUE when exposure only has two levels)

labels Optional treatment labels

Additional arguments to lower level functions

### **Details**

The formula may either be specified as: response ~ treatment | nuisance-formula | propensity-formula

```
For example: ate(y~a | x+z+a | x*z, data=...)

Alternatively, as a list: ate(list(y~a, ~x+z, ~x*z), data=...)

Or using the nuisance (and propensity argument): ate(y~a, nuisance=~x+z, ...)
```

#### Value

An object of class 'ate.targeted' is returned. See targeted-class for more details about this class and its generic functions.

#### Author(s)

Klaus K. Holst

### See Also

cate

```
m <- lvm(y ~ a+x, a~x)
distribution(m, ~y) <- binomial.lvm()
m <- ordinal(m, K=4, ~a)
transform(m, ~a) <- factor
d <- sim(m, 1e3, seed=1)
(a <- ate(y~a|a*x|x, data=d))
## ate(y~a, nuisance=~a*x, propensity=~x, ...)
# Comparison with randomized experiment
m0 <- cancel(m, a~x)
lm(y~a-1, sim(m0,2e4))
# Choosing a different contrast for the association measures
summary(a, contrast=c(2,4))</pre>
```

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calibration

Calibration (training)

### **Description**

Calibration for multiclassication methods

### Usage

```
calibration(
  pr,
  cl,
  weights = NULL,
  threshold = 10,
  method = "bin",
  breaks = nclass.Sturges,
  df = 3,
  ...
)
```

#### **Arguments**

pr matrix with probabilities for each class

cl class variable

weights counts

threshold do not calibrate if less then 'threshold' events

method either 'isotonic' (pava), 'logistic', 'mspline' (monotone spline), 'bin' (local constant)

breaks optional number of bins (only for method 'bin')

df degrees of freedom (only for spline methods)

additional arguments to lower level functions

#### **Details**

•••

#### Value

An object of class 'calibration' is returned. See calibration-class for more details about this class and its generic functions.

### Author(s)

Klaus K. Holst

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```
sim1 <- function(n, beta=c(-3, rep(.5,10)), rho=.5) {
 p <- length(beta)-1</pre>
 xx <- lava::rmvn0(n,sigma=diag(nrow=p)*(1-rho)+rho)</pre>
 y <- rbinom(n, 1, lava::expit(cbind(1,xx)%*%beta))</pre>
 d <- data.frame(y=y, xx)</pre>
 names(d) \leftarrow c("y",paste0("x",1:p))
 return(d)
}
set.seed(1)
beta <- c(-2, rep(1, 10))
d <- sim1(1e4, beta=beta)</pre>
a1 <- NB(y \sim ., data=d)
a2 <- glm(y \sim ., data=d, family=binomial)
## a3 <- randomForest(factor(y) ~ ., data=d, family=binomial)</pre>
d0 < - sim1(1e4, beta=beta)
p1 <- predict(a1, newdata=d0)
p2 <- predict(a2, newdata=d0, type="response")</pre>
## p3 <- predict(a3, newdata=d0, type="prob")</pre>
c2 <- calibration(p2, d0$y, method="isotonic")</pre>
c1 <- calibration(p1, d0$y, breaks=100)</pre>
if (interactive()) {
  plot(c1)
  plot(c2,col="red",add=TRUE)
  abline(a=0,b=1)##'
  with(c1$xy[[1]], points(pred,freq,type="b", col="red"))
}
set.seed(1)
beta <- c(-2, rep(1, 10))
dd <- lava::csplit(sim1(1e4, beta=beta), k=3)</pre>
mod \leftarrow NB(y \sim ., data=dd[[1]])
p1 <- predict(mod, newdata=dd[[2]])</pre>
cal <- calibration(p1, dd[[2]]$y)</pre>
p2 <- predict(mod, newdata=dd[[3]])</pre>
pp <- predict(c1, p2)</pre>
cc <- calibration(pp, dd[[3]]$y)</pre>
if (interactive()) {##'
  plot(cal)
  plot(cc, add=TRUE, col="blue")
```

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#### **Description**

The functions calibration returns an object of the class calibration.

An object of class 'calibration' is a list with at least the following components:

stepfun estimated step-functions (see stepfun) for each class

classes the unique classes

model model/method type (string)

xy list of data.frame's with predictions (pr) and estimated probabilities of success (only for 'bin' method)

#### Value

```
objects of the S3 class 'calibration'
```

### S3 generics

The following S3 generic functions are available for an object of class targeted:

```
predict Apply calibration to new data.

plot Plot the calibration curves (reliability plot).
```

```
calibration, calibrate
```

print Basic print method.

### **Examples**

See Also

```
## See example(calibration) for examples
```

cate

Conditional Average Treatment Effect estimation

#### **Description**

Conditional Average Treatment Effect estimation via Double Machine Learning

### Usage

```
cate(
  treatment,
  response_model,
  propensity_model,
  contrast = c(1, 0),
  data,
  nfolds = 5,
```

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```
type = "dml2",
silent = FALSE,
stratify = FALSE,
mc.cores,
...
)
```

### **Arguments**

formula specifying treatment and variables to condition on treatment response\_model formula or ml\_model object (formula => glm) propensity\_model formula or ml\_model object (formula => glm) treatment contrast (default 1 vs 0) contrast data.frame data nfolds Number of folds 'dml1' or 'dml2' type silent supress all messages and progressbars stratify If TRUE the response\_model will be stratified by treatment mc.cores Optional number of cores. parallel::mcmapply used instead of future mc.cores

additional arguments to future.apply::future\_mapply

#### Value

cate.targeted object

#### Author(s)

Klaus Kähler Holst

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```
e <- cate(a ~ z1+z2+z3, response=u^{-}., data=d) e
```

cate\_link

Conditional Relative Risk estimation

### Description

Conditional average treatment effect estimation via Double Machine Learning

## Usage

```
cate_link(
   treatment,
   link = "identity",
   response_model,
   propensity_model,
   importance_model,
   contrast = c(1, 0),
   data,
   nfolds = 5,
   type = "dml1",
   ...
)
```

### **Arguments**

```
treatment
                 formula specifying treatment and variables to condition on
link
                 Link function
response_model SL object
propensity_model
                  SL object
importance_model
                 SL object
contrast
                 treatment contrast (default 1 vs 0)
data
                  data.frame
nfolds
                 Number of folds
                  'dml1' or 'dml2'
type
                 additional arguments to SuperLearner
```

#### Value

cate.targeted object

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#### Author(s)

Klaus Kähler Holst & Andreas Nordland

### **Examples**

```
# Example 1:
sim1 <- function(n=1e4,</pre>
                  seed=NULL,
                  return_model=FALSE, ...){
suppressPackageStartupMessages(require("lava"))
if (!is.null(seed)) set.seed(seed)
m <- lava::lvm()</pre>
distribution(m, ~x) <- gaussian.lvm()</pre>
distribution(m, ~v) <- gaussian.lvm(mean = 10)</pre>
distribution(m, ~a) <- binomial.lvm("logit")</pre>
regression(m, "a") <- function(v, x)\{.1*v + x\}
distribution(m, "y") <- gaussian.lvm()</pre>
regression(m, "y") <- function(a, v, x){v+x+a*x+a*v*v}
if (return_model) return(m)
lava::sim(m, n = n)
if (require("SuperLearner",quietly=TRUE)) {
  d < - sim1(n = 1e3, seed = 1)
  e <- cate_link(data=d,
           type = "dm12",
            treatment = a \sim v,
           response_model = y^a *(x + v + I(v^2)),
           importance_model = SL(D_ \sim v + I(v^2)),
           nfolds = 10)
  summary(e) # the true parameters are c(1,1)
}
```

cross\_validated-class cross\_validated class object

### Description

The functions cv returns an object of the type cross\_validated.

An object of class 'cross\_validated' is a list with at least the following components:

cv An array with the model score(s) evaluated for each fold, repetition, and model estimates (see estimate.default)

names Names (character vector) of the models

rep number of repetitions of the CV

folds Number of folds of the CV

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### Value

```
objects of the S3 class 'cross_validated'
```

### S3 generics

The following S3 generic functions are available for an object of class cross\_validated:

coef Extract average model scores from the cross-validation procedure.

print Basic print method.

summary Summary of the cross-validation procedure.'

#### See Also

cv

### **Examples**

```
## See example(cv) for examples
```

crr

Conditional Relative Risk estimation

### Description

Conditional Relative Risk estimation via Double Machine Learning

## Usage

```
crr(
   treatment,
   response_model,
   propensity_model,
   importance_model,
   contrast = c(1, 0),
   data,
   nfolds = 5,
   type = "dml1",
   ...
)
```

### **Arguments**

```
treatment formula specifying treatment and variables to condition on response_model SL object propensity_model SL object
```

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```
importance_model
```

SL object

contrast treatment contrast (default 1 vs 0)

data data.frame

nfolds Number of folds type 'dml1' or 'dml2'

... additional arguments to SuperLearner

#### Value

cate.targeted object

#### Author(s)

Klaus Kähler Holst & Andreas Nordland

```
sim1 <- function(n=1e4,</pre>
                  seed=NULL,
                  return_model=FALSE, ...){
suppressPackageStartupMessages(require("lava"))
if (!is.null(seed)) set.seed(seed)
m <- lava::lvm()</pre>
distribution(m, ~x) <- gaussian.lvm()</pre>
distribution(m, ~v) <- gaussian.lvm(mean = 10)</pre>
distribution(m, ~a) <- binomial.lvm("logit")</pre>
regression(m, "a") <- function(v, x)\{.1*v + x\}
distribution(m, "y") <- gaussian.lvm()</pre>
regression(m, "y") <- function(a, v, x)\{v+x+a*x+a*v*v\}
if (return_model) return(m)
lava::sim(m, n = n)
}
d <- sim1(n = 2e3, seed = 1)
if (require("SuperLearner", quietly=TRUE)) {
  e <- crr(data=d,
            type = "dm12",
            treatment = a \sim v,
           response_model = ML(y^a a*(x + v + I(v^2))),
           importance_model = ML(D_ \sim v + I(v^2)),
           propensity_model = ML(a \sim x + v + I(v^2), family=binomial),
           nfolds = 2)
  summary(e) # the true parameters are c(1,1)
}
```

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cv Cross-validation

## Description

Generic cross-validation function

### Usage

```
cv(
  models,
  data,
  response = NULL,
  nfolds = 5,
  rep = 1,
  weights = NULL,
  modelscore,
  seed = NULL,
  shared = NULL,
  args.pred = NULL,
  args.future = list(),
  mc.cores,
  ...
)
```

### **Arguments**

models	List of fitting functions
data	data.frame or matrix
response	Response variable (vector or name of column in data).
nfolds	Number of folds (default 5. K=0 splits in 1:n/2, n/2:n with last part used for testing)
rep	Number of repetitions (default 1)
weights	Optional frequency weights
modelscore	Model scoring metric (default: MSE / Brier score). Must be a function with arguments: response, prediction, weights,
seed	Random seed (argument parsed to future_Apply::future_lapply)
shared	Function applied to each fold with results send to each model
args.pred	Optional arguments to prediction function (see details below)
args.future	Arguments to future.apply::future_mapply
mc.cores	Optional number of cores. parallel::mcmapply used instead of future

Additional arguments parsed to models in models

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#### **Details**

models should be list of objects of class ml\_model. Alternatively, each element of models should be a list with a fitting function and a prediction function.

The response argument can optionally be a named list where the name is then used as the name of the response argument in models. Similarly, if data is a named list with a single data.frame/matrix then this name will be used as the name of the data/design matrix argument in models.

#### Value

An object of class 'cross\_validated' is returned. See cross\_validated-class for more details about this class and its generic functions.

#### Author(s)

Klaus K. Holst

#### **Examples**

```
f0 <- function(data,...) lm(...,data=data)
f1 <- function(data,...) lm(Sepal.Length~Species,data=data)
f2 <- function(data,...) lm(Sepal.Length~Species+Petal.Length,data=data)
x <- cv(list(m0=f0,m1=f1,m2=f2),rep=10, data=iris, formula=Sepal.Length~.)
x</pre>
```

design

Extract design matrix

### **Description**

Extract design matrix from data.frame and formula

#### Usage

```
design(formula, data, intercept = FALSE, rm_envir = FALSE, ...)
```

#### **Arguments**

```
formula formula

data frame

intercept If FALSE (default) an intercept is not included

rm_envir Remove environment

... additional arguments (e.g, specials such weights, offsets, subset)
```

#### Value

An object of class 'design'

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### Author(s)

Klaus Kähler Holst

expand.list

Create a list from all combination of input variables

### **Description**

Similar to expand.grid function, this function creates all combinations of the input arguments but returns the result as a list.

### Usage

```
expand.list(...)
```

### Arguments

... input variables

#### Value

list

### Author(s)

Klaus Kähler Holst

### **Examples**

```
expand.list(x=2:4, z=c("a","b"))
```

ML

ML model

## Description

Wrapper for ml\_model

### Usage

```
ML(formula, model = "glm", ...)
```

### **Arguments**

```
formula formula
```

model (sl, rf, pf, glm, ...)

... additional arguments to model object

ml\_model

#### **Details**

```
model 'sl' (SuperLearner::SuperLearner) args: SL.library, cvControl, f<aamily, method example: model 'grf' (grf::regression_forest) args: num.trees, mtry, sample.weights, sample.fraction, min.node.size, ... example: model 'grf.binary' (grf::probability_forest) args: num.trees, mtry, sample.weights, ... example: model 'glm' args: family, weights, offset, ...
```

ml\_model

R6 class for prediction models

#### **Description**

Provides standardized estimation and prediction methods

#### **Public fields**

info Optional information/name of the model formals List with formal arguments of estimation and prediction functions formula Formula specifying response and design matrix args additional arguments specified during initialization

#### **Active bindings**

fit Active binding returning estimated model object

#### Methods

### **Public methods:**

- ml\_model\$new()
- ml\_model\$estimate()
- ml\_model\$predict()
- ml\_model\$update()
- ml\_model\$print()
- ml\_model\$response()
- ml\_model\$design()
- ml\_model\$opt()
- ml\_model\$clone()

**Method** new(): Create a new prediction model object

Usage:

```
ml_model$new(
    formula = NULL,
    estimate,
    predict = stats::predict,
   predict.args = NULL,
   info = NULL,
   specials,
   response.arg = "y",
   x.arg = "x",
 )
 Arguments:
 formula formula specifying outcome and design matrix
 estimate function for fitting the model (must be a function response, 'y', and design matrix,
     'x'. Alternatively, a function with a single 'formula' argument)
 predict prediction function (must be a function of model object, 'object', and new design
     matrix, 'newdata')
 predict.args optional arguments to prediction function
 info optional description of the model
 specials optional additional terms (weights, offset, id, subset, ...) passed to 'estimate'
 response.arg name of response argument
 x.arg name of design matrix argument
 ... optional arguments to fitting function
Method estimate(): Estimation method
 Usage:
 ml_model$estimate(data, ..., store = TRUE)
 Arguments:
 data data.frame
 ... Additional arguments to estimation method
 store Logical determining if estimated model should be stored inside the class.
Method predict(): Prediction method
 Usage:
 ml_model$predict(newdata, ..., object = NULL)
 Arguments:
 newdata data.frame
 ... Additional arguments to prediction method
 object Optional model fit object
Method update(): Update formula
 ml_model$update(formula, ...)
 Arguments:
```

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```
formula formula or character which defines the new response
 ... Additional arguments to lower level functions
Method print(): Print method
 Usage:
 ml_model$print(...)
 Arguments:
 ... Additional arguments to lower level functions
Method response(): Extract response from data
 Usage:
 ml_model$response(data, ...)
 Arguments:
 data data.frame
 ... additional arguments to 'design'
Method design(): Extract design matrix (features) from data
 Usage:
 ml_model$design(data, ...)
 Arguments:
 data data.frame
 ... additional arguments to 'design'
Method opt(): Get options
 Usage:
 ml_model$opt(arg, ...)
 Arguments:
 arg name of option to get value of
 ... additional arguments to lower level functions
Method clone(): The objects of this class are cloneable with this method.
 Usage:
 ml_model$clone(deep = FALSE)
 Arguments:
 deep Whether to make a deep clone.
```

#### Author(s)

Klaus Kähler Holst

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#### **Examples**

```
data(iris)
rf <- function(formula, ...)</pre>
ml_model$new(formula, info="grf::probability_forest",
  estimate=function(x,y, ...) grf::probability_forest(X=x, Y=y, ...),
  predict=function(object, newdata)
             predict(object, newdata)$predictions, ...)
args <- expand.list(num.trees=c(100,200), mtry=1:3,</pre>
          formula=c(Species ~ ., Species ~ Sepal.Length + Sepal.Width))
models <- lapply(args, function(par) do.call(rf, par))</pre>
x <- models[[1]]$clone()</pre>
x$estimate(iris)
predict(x, newdata=head(iris))
 # Reduce Ex. timing
a <- targeted::cv(models, data=iris)</pre>
cbind(coef(a), attr(args, "table"))
ff <- ml_model$new(estimate=function(y,x) lm.fit(x=x, y=y),</pre>
        predict=function(object, newdata) newdata%*%object$coefficients)
## tmp <- ff$estimate(y, x=x)</pre>
## ff$predict(x)
```

NB

Naive Bayes

### **Description**

Naive Bayes Classifier

#### Usage

```
NB(
  formula,
  data,
  weights = NULL,
  kernel = FALSE,
  laplace.smooth = 0,
  prior = NULL,
  ...
)
```

### **Arguments**

formula

Formula with syntax: response ~ predictors | weights

NB-class

data data.frame

weights optional frequency weights

kernel If TRUE a kernel estimator is used for numeric predictors (otherwise a gaussian

model is used)

laplace.smooth Laplace smoothing

prior optional prior probabilities (default estimated from data)

... additional arguments to lower level functions

#### Value

An object of class 'NB' is returned. See NB-class for more details about this class and its generic functions.

### Author(s)

Klaus K. Holst

### **Examples**

```
data(iris)
m2 <- NB(Species ~ Sepal.Width + Petal.Length, data=iris)
pr2 <- predict(m2, newdata=iris)</pre>
```

NB-class

NB class object

#### **Description**

The functions NB returns an object of the type NB.

An object of class 'NB' is a list with at least the following components:

**prior** Matrix with prior probabilities, i.e. marginal class probabilities Pr(class)

**pcond** list of matrices with conditional probabilities of the features given the classes (one list element per class), Pr(xlclass)

classes Names (character vector) of the classes

xvar number of repetitions of the CV

xmodel Number of folds of the CV

model Number of folds of the CV

#### Value

```
objects of the S3 class 'NB'
```

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### S3 generics

The following S3 generic functions are available for an object of class NB:

```
predict Predict class probabilities for new features data.
print Basic print method.
```

### See Also

NB, NB2

#### **Examples**

```
## See example(NB) for examples
```

nondom

Find non-dominated points of a set

### **Description**

Find the non-dominated point of a set (minima of a point set).

### Usage

```
nondom(x, ...)
```

#### **Arguments**

x matrix

... additional arguments to lower level functions

#### **Details**

A point x dominates y if it is never worse and at least in one case strictly better. Formally, let  $f_i$  denote the ith coordinate of the condition (objective) function, then for all i:  $f_i(x) <= f_i(y)$  and there exists j:  $f_i(x) < f_i(y)$ .

Based on the algorithm of Kung et al. 1975.

#### Value

matrix

### Author(s)

Klaus Kähler Holst

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#### **Examples**

```
rbind(
c(1.0, 0.5),
c(0.0, 1.0),
c(1.0, 0.0),
c(0.5, 1.0),
c(1.0, 1.0),
c(0.8, 0.8)) |> nondom()
```

pava

Pooled Adjacent Violators Algorithm

## Description

Pooled Adjacent Violators Algorithm

### Usage

```
pava(y, x = numeric(0), weights = numeric(0))
```

### Arguments

y response variable

x (optional) predictor vector (otherwise y is assumed to be a priori sorted accord-

ing to relevant predictor)

weights weights (optional) weights

#### Value

List with index (idx) of jump points and values (value) at each jump point.

### Author(s)

Klaus K. Holst

```
x <- runif(5e3, -5, 5)
pr <- lava::expit(-1 + x)
y <- rbinom(length(pr), 1, pr)
pv <- pava(y, x)
plot(pr ~ x, cex=0.3)
with(pv, lines(sort(x)[index], value, col="red", type="s"))</pre>
```

predict.density 25

predict.density

Prediction for kernel density estimates

#### **Description**

Kernel density estimator predictions

#### **Usage**

```
## S3 method for class 'density'
predict(object, xnew, ...)
```

### Arguments

density object object

New data on which to make predictions for xnew additional arguments to lower level functions

#### Author(s)

Klaus K. Holst

predict.NB

Predictions for Naive Bayes Classifier

#### **Description**

Naive Bayes Classifier predictions

#### Usage

```
## S3 method for class 'NB'
predict(object, newdata, expectation = NULL, threshold = c(0.001, 0.001), ...)
```

### **Arguments**

object density object

newdata new data on which to make predictions

expectation Variable to calculate conditional expectation wrt probabilities from NB classifier threshold Threshold parameters. First element defines the threshold on the probabilities

and the second element the value to set those truncated probabilities to.

Additional arguments to lower level functions . . .

### Author(s)

Klaus K. Holst

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**RATE** 

Responder Average Treatment Effect

### Description

Estimation of the Average Treatment Effect among Responders

### Usage

```
RATE(
    response,
    post.treatment,
    treatment,
    data,
    family = gaussian(),
    M = 5,
    pr.treatment,
    treatment.level,
    SL.args.response = list(family = gaussian(), SL.library = c("SL.mean", "SL.glm")),
    SL.args.post.treatment = list(family = binomial(), SL.library = c("SL.mean", "SL.glm")),
    preprocess = NULL,
    efficient = TRUE,
    ...
)
```

## Arguments

Response formula (e.g, Y~D\*A) response post.treatment Post treatment marker formula (e.g., D~W) treatment Treatment formula (e.g, A~1) data data.frame family Exponential family for response (default gaussian) Number of folds in cross-fitting (M=1 is no cross-fitting) (optional) Randomization probability of treatment. pr.treatment treatment.level Treatment level in binary treatment (default 1) SL.args.response Arguments to SuperLearner for the response model SL.args.post.treatment Arguments to SuperLearner for the post treatment indicator (optional) Data preprocessing function preprocess efficient If TRUE, the estimate will be efficient. If FALSE, the estimate will be a simple plug-in estimate. Additional arguments to lower level functions

RATE.surv 27

### Value

estimate object

### Author(s)

Andreas Nordland, Klaus K. Holst

RATE.surv

Responder Average Treatment Effect

#### **Description**

Estimation of the Average Treatment Effect among Responders for Survival Outcomes

### Usage

```
RATE.surv(
  response,
  post.treatment,
  treatment,
  censoring,
  tau,
  data,
 M = 5
  pr.treatment,
  call.response,
  args.response = list(),
 SL.args.post.treatment = list(family = binomial(), SL.library = c("SL.mean", "SL.glm")),
  call.censoring,
  args.censoring = list(),
 preprocess = NULL,
)
```

### **Arguments**

```
response Response formula (e.g., Surv(time, event) \sim D + W).

post.treatment Post treatment marker formula (e.g., D \sim W).

treatment Treatment formula (e.g., A \sim 1).

censoring Censoring formula (e.g., Surv(time, event == 0) \sim D + A + W)).

tau Time-point of interest, see Details.

data data.frame.

M Number of folds in cross-fitting (M=1 is no cross-fitting).

pr.treatment (optional) Randomization probability of treatment.
```

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call.response Model call for the response model (e.g. "mets::phreg").

args.response Additional arguments to the response model.

SL.args.post.treatment

Additional arguments to SuperLearner for the post treatment indicator model.

call.censoring Similar to call.response.

args.censoring Similar to args.response.

preprocess (optional) Data pre-processing function.

... Additional arguments to lower level data pre-processing functions.

#### **Details**

Estimation of

$$\frac{P(T \le \tau | A=1) - P(T \le \tau | A=1)}{E[D|A=1]}$$

under right censoring based on plug-in estimates of  $P(T \le \tau | A = a)$  and E[D|A = 1].

An efficient one-step estimator of  $P(T \leq \tau | A = a)$  is constructed using the efficient influence function

$$\frac{I\{A=a\}}{P(A=a)} \Big( \frac{\Delta}{S_0^c(\tilde{T}|X)} I\{\tilde{T} \leq \tau\} + \int_0^\tau \frac{S_0(u|X) - S_0(\tau|X)}{S_0(u|X) S_0^c(u|X)} dM_0^c(u|X)) \Big) + \Big(1 - \frac{I\{A=a\}}{P(A=a)} \Big) F_0(\tau|A=a,W) - P(T \leq \tau|A) + \frac{I\{A=a\}}{P(A=a)} \Big) F_0(\tau|A=a,W) - \frac{I\{A$$

An efficient one-step estimator of E[D|A=1] is constructed using the efficient influence function

$$\frac{A}{P(A=1)} \left( D - E[D|A=1, W] \right) + E[D|A=1, W] - E[D|A=1].$$

#### Value

estimate object

#### Author(s)

Andreas Nordland, Klaus K. Holst

riskreg

Risk regression

### **Description**

Risk regression with binary exposure and nuisance model for the odds-product.

Let A be the binary exposure, V the set of covariates, and Y the binary response variable, and define  $p_a(v) = P(Y = 1 \mid A = a, V = v), a \in \{0, 1\}.$ 

The **target parameter** is either the *relative risk* 

$$RR(v) = \frac{p_1(v)}{p_0(v)}$$

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or the risk difference

$$RD(v) = p_1(v) - p_0(v)$$

We assume a target parameter model given by either

$$\log\{RR(v)\} = \alpha^t v$$

or

$$\operatorname{arctanh}\{RD(v)\} = \alpha^t v$$

and similarly a working linear nuisance model for the odds-product

$$\phi(v) = \log\left(\frac{p_0(v)p_1(v)}{(1 - p_0(v))(1 - p_1(v))}\right) = \beta^t v$$

A propensity model for E(A=1|V) is also fitted using a logistic regression working model

$$logit{E(A = 1 \mid V = v)} = \gamma^t v.$$

If both the odds-product model and the propensity model are correct the estimator is efficient. Further, the estimator is consistent in the union model, i.e., the estimator is double-robust in the sense that only one of the two models needs to be correctly specified to get a consistent estimate.

### Usage

```
riskreg(
  formula,
  nuisance = ~1,
  propensity = ~1,
  target = ~1,
  data,
  weights,
  type = "rr",
  optimal = TRUE,
  std.err = TRUE,
  start = NULL,
  mle = FALSE,
  ...
)
```

#### **Arguments**

formula formula (see details below)
nuisance nuisance model (formula)
propensity propensity model (formula)
target (optional) target model (formula)
data data.frame
weights optional weights

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type	type of association measure (rd og rr)
optimal	If TRUE optimal weights are calculated
std.err	If TRUE standard errors are calculated
start	optional starting values
mle	Semi-parametric (double-robust) estimate or MLE (TRUE gives MLE)
	additional arguments to unconstrained optimization routine (nlminb)

#### **Details**

```
The 'formula' argument should be given as response ~ exposure | target-formula | nuisance-formula or response ~ exposure | target | nuisance | propensity
```

```
E.g., riskreg(y \sim a | 1 | x+z | x+z, data=...)
```

Alternatively, the model can specifed using the target, nuisance and propensity arguments: riskreg(y ~ a, target=~1, nuisance=~x+z, ...)

The riskreg\_fit function can be used with matrix inputs rather than formulas.

#### Value

An object of class 'riskreg. targeted' is returned. See targeted-class for more details about this class and its generic functions.

#### Author(s)

Klaus K. Holst

#### References

Richardson, T. S., Robins, J. M., & Wang, L. (2017). On modeling and estimation for the relative risk and risk difference. Journal of the American Statistical Association, 112(519), 1121–1130. http://dx.doi.org/10.1080/01621459.2016.1192546

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```
## Model with same design matrix for nuisance and propensity model:
with(d, riskreg_fit(y, a, nuisance=X, type="rr"))

## a <- riskreg(y ~ a, target=~z, nuisance=~x, propensity=~x, data=d, type="rr")
a <- riskreg(y ~ a | z, nuisance=~x, propensity=~x, data=d, type="rr")
a
predict(a, d[1:5,])

riskreg(y ~ a, nuisance=~x, data=d, type="rr", mle=TRUE)</pre>
```

riskreg\_cens

Binary regression models with right censored outcomes

### **Description**

Binary regression models with right censored outcomes

### Usage

```
riskreg_cens(
  response,
  censoring,
  treatment = NULL,
  prediction = NULL,
  data,
  newdata,
  tau,
  type = "risk",
 M = 1,
  call.response = "phreg",
  args.response = list(),
  call.censoring = "phreg",
  args.censoring = list(),
  preprocess = NULL,
  efficient = TRUE,
  control = list(),
)
```

#### **Arguments**

```
\begin{tabular}{lll} response & Response & Response & Formula (e.g., Surv(time, event) $\sim D + W$). \\ & Censoring & Censoring & Formula (e.g., Surv(time, event == 0) $\sim D + A + W$)). \\ & treatment & Optional treatment model (ml_model) \\ & Optional prediction model (ml_model) \\ & data. & frame. \\ \end{tabular}
```

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newdata Optional data.frame. In this case the uncentered influence function evalued in

'newdata' is returned with nuisance parameters obtained from 'data'.

tau Time-point of interest, see Details.
type "risk", "treatment", "rmst", "brier"

M Number of folds in cross-fitting (M=1 is no cross-fitting).

call.response Model call for the response model (e.g. "mets::phreg").

args.response Additional arguments to the response model.

call.censoring Similar to call.response.

args.censoring Similar to args.response.

preprocess (optional) Data pre-processing function.

efficient If FALSE an IPCW estimator is returned

control See details

... Additional arguments to lower level data pre-processing functions.

#### **Details**

The one-step estimator depends on the calculation of an integral wrt. the martingale process corresponding to the counting process  $N(t) = I(C > \min(T, tau))$ . This can be decomposed into an integral wrt the counting process,  $dN_c(t)$  and the compensator  $d\Lambda_c(t)$  where the latter term can be computational intensive to calculate. Rather than calculating this integral in all observed time points, we can make a coarser evaluation which can be controlled by setting control=(sample=N). With N=0 the (computational intensive) standard evaluation is used.##'

### Value

estimate object

#### Author(s)

Klaus K. Holst, Andreas Nordland

scoring Predictive model scoring

#### **Description**

Predictive model scoring

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### Usage

```
scoring(
  response,
  ...,
  type = "quantitative",
  levels = NULL,
  metrics = NULL,
  weights = NULL,
  names = NULL,
  messages = 1
)
```

### Arguments

response	Observed response
	model predictions (continuous predictions or class probabilities (matrices))
type	continuous or categorical response (the latter is automatically chosen if response is a factor, otherwise a continuous response is assumed)
levels	(optional) unique levels in response variable
metrics	which metrics to report
weights	optional frequency weights
names	optional names of models coments (given as, alternatively these can be named arguments)
messages	controls amount of messages/warnings (0: none)

#### Value

Numeric matrix of dimension  $m \times p$ , where m is the number of different models and p is the number of model metrics

```
data(iris)
set.seed(1)
dat <- csplit(iris,2)
g1 <- NB(Species ~ Sepal.Width + Petal.Length, data=dat[[1]])
g2 <- NB(Species ~ Sepal.Width, data=dat[[1]])
pr1 <- predict(g1, newdata=dat[[2]], wide=TRUE)
pr2 <- predict(g2, newdata=dat[[2]], wide=TRUE)
table(colnames(pr1)[apply(pr1,1,which.max)], dat[[2]]$Species)
table(colnames(pr2)[apply(pr2,1,which.max)], dat[[2]]$Species)
scoring(dat[[2]]$Species, pr1=pr1, pr2=pr2)
## quantitative response:
scoring(response=1:10, prediction=rnorm(1:10))</pre>
```

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SL

SuperLearner wrapper for ml\_model

### **Description**

SuperLearner wrapper for ml\_model

### Usage

```
SL(
  formula = ~.,
    ...,
  SL.library = c("SL.mean", "SL.glm"),
  binomial = FALSE,
  data = NULL
)
```

### Arguments

formula Model design

... Additional arguments for SuperLearner::SuperLearner

SL.library character vector of prediction algorithms

binomial boolean specifying binomial or gaussian family (default FALSE)

data Optional data.frame

### Value

ml\_model object

#### Author(s)

Klaus Kähler Holst

softmax

Softmax transformation

### Description

Softmax transformation

### Usage

```
softmax(x, log = FALSE, ref = TRUE, ...)
```

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### **Arguments**

Х	Input matrix (e.g., linear predictors of multinomial logistic model)
log	Return on log-scale (default FALSE)
ref	Add reference level (add 0 column to x)
	Additional arguments to lower level functions

#### Value

Numeric matrix of dimension n x p, where n= nrow(x) and p = ncol(x) + (ref == TRUE)

solve_ode	Solve ODE

### Description

Solve ODE with Runge-Kutta method (RK4)

### Usage

```
solve_ode(ode_ptr, input, init, par = 0)
```

### **Arguments**

ode\_ptr pointer (externalptr) to C++ function or an R function input Input matrix. 1st column specifies the time points init Initial conditions

par Parameters defining the ODE (parsed to ode\_ptr)

### **Details**

The external point should be created with the function targeted::specify\_ode.

### Value

Matrix with solution

### Author(s)

Klaus Kähler Holst

### See Also

specify\_ode

```
example(specify_ode)
```

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specify\_ode

Specify Ordinary Differential Equation (ODE)

#### **Description**

Define compiled code for ordinary differential equation.

### Usage

```
specify_ode(code, fname = NULL, pname = c("dy", "x", "y", "p"))
```

#### **Arguments**

code string with the body of the function definition (see details)

fname Optional name of the exported C++ function

pname Vector of variable names (results, inputs, states, parameters)

#### **Details**

The model (code) should be specified as the body of C++ function. The following variables are defined bye default (see the argument pname)

- dy Vector with derivatives, i.e. the rhs of the ODE (the result).
- x Vector with the first element being the time, and the following elements additional exogenous input variables,
- y Vector with the dependent variable
- p Parameter vector

 $y'(t) = f_p(x(t), y(t))$  All variables are treated as Armadillo (http://arma.sourceforge.net/) vectors/matrices.

As an example consider the Lorenz Equations  $\frac{dx_t}{dt} = \sigma(y_t - x_t) \frac{dy_t}{dt} = x_t(\rho - z_t) - y_t \frac{dz_t}{dt} = x_t y_t - \beta z_t$ 

We can specify this model as ode <- 'dy(0) = p(0)\*(y(1)-y(0)); dy(1) = y(0)\*(p(1)-y(2)); dy(2) = y(0)\*y(1)-p(2)\*y(2); 'dy <- specify\_ode(ode)

As an example of model with exogenous inputs consider the following ODE:  $y'(t) = \beta_0 + \beta_1 y(t) + \beta_2 y(t)x(t) + \beta_3 x(t) \cdot t$  This could be specified as mod <- 'double t = x(0); dy = p(0) + p(1)\*y + p(2)\*x(1)\*y + p(3)\*x(1)\*t; 'dy <- specify\_ode(mod)##'

#### Value

pointer (externalptr) to C++ function

#### Author(s)

Klaus Kähler Holst

targeted-class 37

#### See Also

solve ode

targeted-class

targeted class object

### Description

The functions riskreg and ate returns an object of the type targeted.

An object of class 'targeted' is a list with at least the following components:

estimate An estimate object with the target parameter estimates (see estimate.default)

opt Object returned from the applied optimization routine

**npar** number of parameters of the model (target and nuisance)

type String describing the model

#### Value

objects of the S3 class 'targeted'

### S3 generics

The following S3 generic functions are available for an object of class targeted:

coef Extract target coefficients of the estimated model.

vcov Extract the variance-covariance matrix of the target parameters.

IC Extract the estimated influence function.

print Print estimates of the target parameters.

summary Extract information on both target parameters and estimated nuisance model.'

#### See Also

```
riskreg, ate
```

```
## See example(riskreg) for examples
```

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