

# Package ‘BioPred’

January 20, 2025

**Type** Package

**Title** An R Package for Biomarkers Analysis in Precision Medicine

**Version** 1.0.2

**Date** 2024-11-03

**Maintainer** Zihuan Liu <zihuan.liu@abbvie.com>

**Description**

Provides functions for training extreme gradient boosting model using propensity score A-learning and weight-learning methods. For further details, see Liu et al. (2024) <[doi:10.1093/bioinformatics/btae592](https://doi.org/10.1093/bioinformatics/btae592)>.

**Encoding** UTF-8

**Language** en

**License** GPL-3

**Imports** xgboost, pROC, ggplot2, PropCIs, survival, survminer, mgcv, onewaytests, car

**RoxygenNote** 7.3.2

**Suggests** knitr, rmarkdown, kableExtra

**VignetteBuilder** knitr

**NeedsCompilation** no

**Depends** R (>= 4.0.0)

**LazyData** true

**Author** Zihuan Liu [aut, cre],

Yan Sun [aut],

Xin Huang [aut]

**Repository** CRAN

**Date/Publication** 2024-11-04 08:30:13 UTC

## Contents

|                       |   |
|-----------------------|---|
| cat_summary . . . . . | 2 |
| cdf_plot . . . . .    | 3 |

|                                    |    |
|------------------------------------|----|
| cut_perf . . . . .                 | 4  |
| eval_metric_bin . . . . .          | 6  |
| eval_metric_con . . . . .          | 7  |
| eval_metric_sur . . . . .          | 7  |
| fixcut_bin . . . . .               | 8  |
| fixcut_con . . . . .               | 10 |
| fixcut_sur . . . . .               | 12 |
| gam_ctr_plot . . . . .             | 13 |
| gam_plot . . . . .                 | 15 |
| get_subgroup_results . . . . .     | 18 |
| predictive_biomarker_imp . . . . . | 19 |
| roc_bin . . . . .                  | 20 |
| roc_bin_plot . . . . .             | 21 |
| scat_cont_plot . . . . .           | 22 |
| subgrp_perf . . . . .              | 24 |
| subgrp_perf_pred . . . . .         | 25 |
| tutorial_data . . . . .            | 27 |
| XGBoostSub_bin . . . . .           | 29 |
| XGBoostSub_con . . . . .           | 31 |
| XGBoostSub_sur . . . . .           | 33 |

|              |           |
|--------------|-----------|
| <b>Index</b> | <b>36</b> |
|--------------|-----------|

---

|             |  |
|-------------|--|
| cat_summary | <i>Summarize Categorical Variables in Subgroup</i> |
|-------------|--|

---

## Description

This function provides a summary of categorical variables in a dataset.

## Usage

```
cat_summary(
  yvar,
  yname,
  xvars,
  xname.list,
  data,
  yvar.display = yvar,
  xvars.display = xvars
)
```

## Arguments

|       |                                      |
|-------|--------------------------------------|
| yvar  | Name of the variable for summary.    |
| yname | A vector of ordered y values.        |
| xvars | Names of the variables for grouping. |

|               |   |
|---------------|---|
| xname.list    | A list (same order as xvars) of ordered x values for each xvar. |
| data          | The dataset.  |
| yvar.display  | Display name for yvar.  |
| xvars.display | Display name for xvars.   |

**Value**

A list containing the contingency table, frequency table, and percentage table.

**Examples**

```
# Load a sample dataset
data <- data.frame(
  outcome = sample(c("A", "B", "C"), 100, replace = TRUE), # categorical outcome
  group1 = sample(c("Male", "Female"), 100, replace = TRUE), # group variable 1
  group2 = sample(c("Young", "Old"), 100, replace = TRUE) # group variable 2
)

# Summarize categorical outcome by two grouping variables
cat_summary(
  yvar = "outcome",
  yname = c("A", "B", "C"), # ordered categories for outcome
  xvars = c("group1", "group2"),
  xname.list = list(c("Male", "Female"), c("Young", "Old")),
  data = data,
  yvar.display = "Outcome Category",
  xvars.display = c("Gender", "Age Group")
)
```

cdf\_plot

*CDF Plot for a biomarker***Description**

Cumulative Distribution Function (CDF) plot for a biomarker.

**Usage**

```
cdf_plot(xvar, data, y.int = 5, xlim = NULL, xvar.display = xvar, group = NULL)
```

**Arguments**

|              |   |
|--------------|---|
| xvar         | The biomarker name.   |
| data         | The dataset.  |
| y.int        | Increase interval on the y.   |
| xlim         | cdf plot range for xvar, when NULL, c(min(x), max(x)) will be used. |
| xvar.display | Display name of the biomarker.                                      |
| group        | A separate CDF line will be plotted for each group.                 |

**Value**

A ggplot object representing the CDF inverse plot.

**Examples**

```
# Load a sample dataset
data <- data.frame(
  biomarker = rnorm(100, mean = 50, sd = 10),
  group = sample(c("Group A", "Group B"), 100, replace = TRUE)
)

# Basic CDF plot for a single biomarker without groups
cdf_plot(
  xvar = "biomarker",
  data = data,
  y.int = 10,
  xlim = c(30, 70),
  xvar.display = "Biomarker Level"
)

# CDF plot for a biomarker with groups
cdf_plot(
  xvar = "biomarker",
  data = data,
  y.int = 10,
  xlim = c(30, 70),
  xvar.display = "Biomarker Level",
  group = "group"
)
```

---

cut\_perf

*Cutoff Performance Evaluation*

---

**Description**

This function evaluates the performance of a predictive model at a selected cutoff point.

**Usage**

```
cut_perf(
  yvar,
  censorvar = NULL,
  xvar,
  cutoff,
  dir,
  xvars.adj = NULL,
  data,
  type,
  yvar.display = yvar,
  xvar.display = xvar
)
```

**Arguments**

|              |   |
|--------------|---|
| yvar         | Response variable name.   |
| sensorvar    | Censoring variable name (0-censored, 1-event).                              |
| xvar         | Biomarker name.   |
| cutoff       | Selected cutoff value.  |
| dir          | Direction for desired subgroup (">", ">=", "<", "<=").                      |
| xvars.adj    | Other covariates to adjust when evaluating the performance.                 |
| data         | Data frame containing the variables.  |
| type         | Type of analysis: "c" for continuous, "s" for survival, and "b" for binary. |
| yvar.display | Display name of response variable.  |
| xvar.display | Display name of biomarker variable.   |

**Value**

A list containing various performance metrics and optionally, plots.

**Examples**

```
# Load a sample dataset
data <- data.frame(
  survival_time = rexp(100, rate = 0.1), # survival time
  status = sample(c(0, 1), 100, replace = TRUE), # censoring status
  biomarker = rnorm(100, mean = 0, sd = 1), # biomarker levels
  covariate1 = rnorm(100, mean = 50, sd = 10) # an additional covariate
)
# Perform cutoff performance evaluation for continuous outcome
data$continuous_outcome <- rnorm(100, mean = 10, sd = 5)
cut_perf(
  yvar = "continuous_outcome",
  xvar = "biomarker",
  cutoff = 0.5,
  dir = ">=",
  data = data,
  type = "c",
  yvar.display = "Continuous Outcome",
  xvar.display = "Biomarker Level"
)

# Perform cutoff performance evaluation for binary outcome
data$binary_outcome <- sample(c(0, 1), 100, replace = TRUE)
cut_perf(
  yvar = "binary_outcome",
  xvar = "biomarker",
  cutoff = 0,
  dir = "<=",
  data = data,
  type = "b",
  yvar.display = "Binary Outcome",
```

```
xvar.display = "Biomarker Level"
)
```

---

eval\_metric\_bin      *Evaluation Metrics for XGBoostSub\_bin Model*

---

### Description

Function for evaluating XGBoostSub\_bin model performance.

### Usage

```
eval_metric_bin(model, X_feature, y_label, pi, trt, Loss_type = "A_learning")
```

### Arguments

|           |  |
|-----------|--|
| model     | The trained XGBoostSub_bin model object.   |
| X_feature | The input features matrix.   |
| y_label   | The input y matrix.  |
| pi        | The propensity scores vector, which should range from 0 to 1, representing the probability of assignment to treatment.                     |
| trt       | The treatment indicator vector. Should take values of 1 or -1, where 1 represents the treatment group and -1 represents the control group. |
| Loss_type | Type of loss function to use: "A_learning" or "Weight_learning".   |

### Details

eval\_metric: Function for Evaluating XGBoostSub\_bin Model Performance

This function evaluates the performance of an XGBoostSub\_bin model using a A-learning or weight-learning function.

### Value

Evaluation result of the XGBoostSub\_bin model.

---

eval\_metric\_con      *Evaluation Metrics for XGBoostSub\_con Model*

---

### Description

Function for evaluating XGBoostSub\_con model performance.

### Usage

```
eval_metric_con(model, X_feature, y_label, pi, trt, Loss_type = "A_learning")
```

### Arguments

|           |  |
|-----------|--|
| model     | The trained XGBoostSub_con model object.   |
| X_feature | The input features matrix.   |
| y_label   | The input y matrix.  |
| pi        | The propensity scores vector, which should range from 0 to 1, representing the probability of assignment to treatment.                     |
| trt       | The treatment indicator vector. Should take values of 1 or -1, where 1 represents the treatment group and -1 represents the control group. |
| Loss_type | Type of loss function to use: "A_learning" or "Weight_learning".   |

### Details

eval\_metric: Function for Evaluating XGBoostSub\_con Model Performance

This function evaluates the performance of an XGBoostSub\_con model using a A-learning or weight-learning function.

### Value

Evaluation result of the XGBoostSub\_con model.

---

eval\_metric\_sur      *Evaluation Metrics for XGBoostSub\_sur Model*

---

### Description

Function for evaluating XGBoostSub\_sur model performance.

**Usage**

```
eval_metric_sur(
  model,
  X_feature,
  y_label,
  pi,
  trt,
  censor,
  Loss_type = "A_learning"
)
```

**Arguments**

|           |  |
|-----------|--|
| model     | The trained XGBoostSub_sur model object.   |
| X_feature | The input features matrix.   |
| y_label   | The input y matrix.  |
| pi        | The propensity scores vector, which should range from 0 to 1, representing the probability of assignment to treatment.                     |
| trt       | The treatment indicator vector. Should take values of 1 or -1, where 1 represents the treatment group and -1 represents the control group. |
| censor    | The censor status vector. Should take values of 1 or 0, where 1 represents censoring and 0 represents an observed event.                   |
| Loss_type | Type of loss function to use: "A_learning" or "Weight_learning".   |

**Details**

eval\_metric: Function for Evaluating XGBoostSub\_con Model Performance

This function evaluates the performance of an XGBoostSub\_con model using a A-learning or weight-learning function.

**Value**

Evaluation result of the XGBoostSub\_sur model.

---

|            |  |
|------------|--|
| fixcut_bin | <i>Fixed Cutoff Analysis for Individual Biomarker Associated with Binary Outcome Variables</i> |
|------------|--|

---

**Description**

This function conducts fixed cutoff analysis for individual biomarker associated with binary outcome variables.



**Usage**

```
fixcut_bin(
  yvar,
  xvar,
  dir,
  cutoffs,
  data,
  method = "Fisher",
  yvar.display = yvar,
  xvar.display = xvar,
  vert.x = FALSE
)
```

**Arguments**

|              |  |
|--------------|--|
| yvar         | Binary response variable name. 0 represents controls and 1 represents cases.   |
| xvar         | Biomarker name.  |
| dir          | Cutoff direction for the desired subgroup. Options are ">", ">=", "<", or "<=".  |
| cutoffs      | A vector of candidate cutoffs.   |
| data         | The dataset containing the variables.  |
| method       | Method for cutoff selection. Options are "Fisher", "Youden", "Conc.Prob", "Accuracy", or "Kappa". - "Fisher": Minimizes the Fisher test p-value. - "Youden": Maximizes the Youden index. - "Conc.Prob": Maximizes sensitivity * specificity. - "Accuracy": Maximizes accuracy. - "Kappa": Maximizes Kappa coefficient. |
| yvar.display | Display name of the response variable.   |
| xvar.display | Display name of the predictor variable.  |
| vert.x       | Whether to display the cutoff in a 90-degree angle when plotting (saves space).  |

**Value**

A list containing statistical summaries, selected cutoff statistics, selected cutoff value, confusion matrix, and a ggplot object for visualization.

**Examples**

```
# Load a sample dataset
data <- data.frame(
  outcome = sample(c(0, 1), 100, replace = TRUE),
  biomarker = rnorm(100, mean = 0, sd = 1)
)

# Perform fixed cutoff analysis using the "Fisher" method for a biomarker
fixcut_bin(
  yvar = "outcome",
  xvar = "biomarker",
  dir = ">",
```

```
    cutoffs = seq(-2, 2, by = 0.5),
    data = data,
    method = "Fisher",
    yvar.display = "Binary Outcome",
    xvar.display = "Biomarker Level",
    vert.x = TRUE
)

# Perform fixed cutoff analysis using the "Youden" method
fixcut_bin(
  yvar = "outcome",
  xvar = "biomarker",
  dir = "<",
  cutoffs = seq(-2, 2, by = 0.5),
  data = data,
  method = "Youden",
  yvar.display = "Binary Outcome",
  xvar.display = "Biomarker Level",
  vert.x = FALSE
)

# Perform fixed cutoff analysis using "Accuracy" method with different direction
fixcut_bin(
  yvar = "outcome",
  xvar = "biomarker",
  dir = ">=",
  cutoffs = c(-1, 0, 1),
  data = data,
  method = "Accuracy",
  yvar.display = "Binary Outcome",
  xvar.display = "Biomarker Level",
  vert.x = TRUE
)
```

---

fixcut\_con

*Fixed Cutoff Analysis for Individual Biomarker Associated with Continuous Outcome*

---

## Description

This function conducts fixed cutoff analysis for individual biomarker associated with continuous outcome variables.

## Usage

```
fixcut_con(
  yvar,
  xvar,
  dir,
  cutoffs,
```

```

    data,
    method = "t.test",
    yvar.display = yvar,
    xvar.display = xvar,
    vert.x = FALSE
  )

```

### Arguments

|              |  |
|--------------|--|
| yvar         | Continuous response variable name.   |
| xvar         | Biomarker name.  |
| dir          | Cutoff direction for the desired subgroup. Options are ">", ">=", "<", or "<=".                          |
| cutoffs      | A vector of candidate cutoffs.   |
| data         | The dataset containing the variables.  |
| method       | Method for cutoff selection. Currently only supports "t.test". - "t.test": Minimizes the t-test p-value. |
| yvar.display | Display name of the response variable.   |
| xvar.display | Display name of the predictor variable.  |
| vert.x       | Whether to display the cutoff in a 90-degree angle when plotting (saves space).                          |

### Value

A list containing statistical summaries, selected cutoff statistics, selected cutoff value, group statistics, and a ggplot object for visualization.

### Examples

```

# Load a sample dataset
data <- data.frame(
  outcome = rnorm(100, mean = 10, sd = 5),
  biomarker = rnorm(100, mean = 0, sd = 1)
)

# Perform fixed cutoff analysis using the "t.test" method with '>' direction
fixcut_con(
  yvar = "outcome",
  xvar = "biomarker",
  dir = ">",
  cutoffs = seq(-2, 2, by = 0.5),
  data = data,
  method = "t.test",
  yvar.display = "Continuous Outcome",
  xvar.display = "Biomarker Level",
  vert.x = TRUE
)

# Perform fixed cutoff analysis with '<=' direction
fixcut_con(
  yvar = "outcome",

```

```

xvar = "biomarker",
dir = "<=",
cutoffs = c(-1, 0, 1),
data = data,
method = "t.test",
yvar.display = "Continuous Outcome",
xvar.display = "Biomarker Level",
vert.x = FALSE
)

```

---

fixcut\_sur

*Fixed Cutoff Analysis for Individual Biomarker Associated with Survival Outcome*


---

### Description

This function conducts fixed cutoff analysis for Individual Biomarker Associated with survival outcome variables.

### Usage

```

fixcut_sur(
  yvar,
  censorvar,
  xvar,
  dir,
  cutoffs,
  data,
  method = "logrank",
  yvar.display = yvar,
  xvar.display = xvar,
  vert.x = FALSE
)

```

### Arguments

|              |  |
|--------------|--|
| yvar         | Survival response variable name.   |
| censorvar    | Censoring variable. 0 indicates censored, 1 indicates an event.  |
| xvar         | Biomarker name.  |
| dir          | Cutoff direction for the desired subgroup. Options are ">", ">=", "<", or "<=".                                  |
| cutoffs      | A vector of candidate cutoffs.   |
| data         | The dataset containing the variables.  |
| method       | Method for cutoff selection. Currently only supports "logrank". - "logrank": Minimizes the logrank test p-value. |
| yvar.display | Display name of the response variable.   |
| xvar.display | Display name of the predictor variable.  |
| vert.x       | Whether to display the cutoff in a 90-degree angle when plotting (saves space).                                  |

**Value**

A list containing statistical summaries, selected cutoff statistics, selected cutoff value, group statistics, and a ggplot object for visualization.

**Examples**

```
# Load a sample dataset
data <- data.frame(
  time = rexp(100, rate = 0.1), # survival time
  status = sample(c(0, 1), 100, replace = TRUE), # censoring status
  biomarker = rnorm(100, mean = 0, sd = 1) # biomarker levels
)

fixcut_sur(
  yvar = "time",
  censorvar = "status",
  xvar = "biomarker",
  dir = "<=",
  cutoffs = c(-1, 0, 1),
  data = data,
  method = "logrank",
  yvar.display = "Survival Time",
  xvar.display = "Biomarker Level",
  vert.x = FALSE
)
```

---

gam\_ctr\_plot

*GAM Contrast Plot*

---

**Description**

Computes and plots the contrasts between treatment and control group based on a GAM for exploring the relationship between treatment benefit and biomarker.

**Usage**

```
gam_ctr_plot(
  yvar,
  censorvar = NULL,
  xvar,
  xvars.adj = NULL,
  sxvars.adj = NULL,
  trtvar = NULL,
  type,
  data,
  k,
  title = "Group Contrast",
  ybreaks = NULL,
```

```

xbreaks = NULL,
rugcol.var = NULL,
link.scale = TRUE,
prt.sum = TRUE,
prt.chk = FALSE,
outlier.rm = FALSE
)

```

### Arguments

|            |   |
|------------|---|
| yvar       | Response variable name.   |
| sensorvar  | Censoring variable name (0-censored, 1-event). Required if type is "s" (survival).  |
| xvar       | Biomarker name.   |
| xvars.adj  | Potential confounding variables to adjust for using linear terms.   |
| sxvars.adj | Potential confounding variables to adjust for using curves.   |
| trtvar     | Treatment variable that the contrast will build upon (treatment-control).   |
| type       | Type of response variable. Options are "c" for continuous, "s" for survival, and "b" for binary response.                     |
| data       | The dataset containing the variables.   |
| k          | Upper limit on the degrees of freedom associated with an s smooth. When this k is too large, program will report error saying |
| title      | Title of the plot.  |
| ybreaks    | Breaks on the y-axis.   |
| xbreaks    | Breaks on the x-axis.   |
| rugcol.var | Variable name that defines the color of the rug.  |
| link.scale | Whether to show the plot (y-axis) in the scale of the link function (linear predictor).                                       |
| prt.sum    | Whether to print summary or not.  |
| prt.chk    | Whether to print model diagnosis.   |
| outlier.rm | Whether to remove outliers based on 1.5IQR.   |

### Value

A list containing the p-value table, summarized p-value table, s-value table, summarized s-value table, and the plot.

### Examples

```

# Load a sample dataset
data <- data.frame(
  response = rnorm(100),
  biomarker = rnorm(100, mean = 50, sd = 10),
  censor = sample(c(0, 1), 100, replace = TRUE),
  treatment = sample(c(0, 1), 100, replace = TRUE),

```

```
age = rnorm(100, mean = 60, sd = 10),
group = sample(c("Group A", "Group B"), 100, replace = TRUE)
)

# Generate a GAM contrast plot for a continuous response variable
gam_ctr_plot(
  yvar = "response",
  xvar = "biomarker",
  trtvar = "treatment",
  type = "c",
  data = data,
  xvars.adj = "age",
  k = 5,
  title = "GAM Contrast Plot for Treatment vs. Control"
)

# Generate a GAM contrast plot for survival analysis
gam_ctr_plot(
  yvar = "response",
  censorvar = "censor",
  xvar = "biomarker",
  trtvar = "treatment",
  type = "s",
  data = data,
  k = 5,
  title = "GAM Contrast Plot for Survival Data"
)

# Generate a GAM contrast plot for a binary response variable
data$binary_response <- as.numeric(data$response > 0)
gam_ctr_plot(
  yvar = "binary_response",
  xvar = "biomarker",
  trtvar = "treatment",
  type = "b",
  data = data,
  k = 5,
  title = "GAM Contrast Plot for Binary Outcome"
)
```

---

gam\_plot

*GAM Plot*

---

### Description

Generates a generalized additive model (GAM) plot for exploring the relationship between a response variable and a biomarker.

**Usage**

```
gam_plot(
  yvar,
  censorvar = NULL,
  xvar,
  xvars.adj = NULL,
  sxvars.adj = NULL,
  type,
  data,
  k,
  pred.type = "iterms",
  link.scale = TRUE,
  title = "Trend Plot",
  ybreaks = NULL,
  xbreaks = NULL,
  rugcol.var = NULL,
  add.points = FALSE,
  prt.sum = TRUE,
  prt.chk = FALSE,
  outlier.rm = FALSE,
  newdat = NULL
)
```

**Arguments**

|            |  |
|------------|--|
| yvar       | Response variable name.  |
| censorvar  | Censoring variable name for survival analysis (0-censored, 1-event). |
| xvar       | Biomarker name.  |
| xvars.adj  | Potential confounding variables to adjust for using linear terms.    |
| sxvars.adj | Potential confounding variables to adjust for using curve terms.     |
| type       | "c" for continuous, "s" for survival, and "b" for binary response.   |
| data       | The dataset containing the variables.                                |
| k          | Upper limit on the degrees of freedom associated with an s smooth.   |
| pred.type  | "iterms" for trend of xvar, "response" for Y at the original scale.  |
| link.scale | Whether to show the plot in the scale of the link function.          |
| title      | Title of the plot.   |
| ybreaks    | Breaks on the y-axis.  |
| xbreaks    | Breaks on the x-axis.  |
| rugcol.var | Variable name defining the color of the rug and points.              |
| add.points | Whether to add data points to the plot.                              |
| prt.sum    | Whether to print summary or not.                                     |
| prt.chk    | Whether to print model diagnosis.                                    |
| outlier.rm | Whether to remove outliers based on 1.5IQR.                          |
| newdat     | User-supplied customized data for prediction and plotting.           |



**Value**

A list containing p-table, s-table, GAM summary, GAM check, and the plot.

**Examples**

```
# Load a sample dataset
data <- data.frame(
  response = rnorm(100),
  biomarker = rnorm(100, mean = 50, sd = 10),
  censor = sample(c(0, 1), 100, replace = TRUE),
  age = rnorm(100, mean = 60, sd = 10),
  group = sample(c("Group A", "Group B"), 100, replace = TRUE)
)

# Generate a GAM plot for a continuous response variable
gam_plot(
  yvar = "response",
  xvar = "biomarker",
  type = "c",
  data = data,
  xvars.adj = "age",
  sxvars.adj = NULL,
  k = 5,
  pred.type = "iterms",
  title = "GAM Plot of Biomarker and Response"
)

# Generate a GAM plot for survival analysis
gam_plot(
  yvar = "response",
  censorvar = "censor",
  xvar = "biomarker",
  type = "s",
  data = data,
  k = 5,
  title = "GAM Survival Plot for Biomarker"
)

# Generate a GAM plot for a binary response variable
data$binary_response <- as.numeric(data$response > 0)
gam_plot(
  yvar = "binary_response",
  xvar = "biomarker",
  type = "b",
  data = data,
  k = 5,
  pred.type = "response",
  title = "GAM Plot for Binary Response"
)
```

---

get\_subgroup\_results *Get Subgroup Results*

---

### Description

This function predicts the treatment assignment for each patient based on a cutoff value.

### Usage

```
get_subgroup_results(model, X_feature, subgroup_label = NULL, cutoff = 0.5)
```

### Arguments

|                |   |
|----------------|---|
| model          | The trained XGBoost-based subgroup model.   |
| X_feature      | The data matrix containing patient features.  |
| subgroup_label | (Optional) The subgroup labels. In real-world data, this information is typically unknown and only available in simulated data. If provided, the prediction accuracy will also be returned. |
| cutoff         | The cutoff value for treatment assignment, defaulted to 0.5.  |

### Value

A data frame containing each subject and assigned treatment (1 for treatment, 0 for control). If subgroup labels are provided, it also returns the prediction accuracy of the subgroup labels.

### Examples

```
X_data <- matrix(rnorm(100 * 10), ncol = 10) # 100 samples with 10 features
y_data <- rnorm(100) # continuous outcome variable
trt <- sample(c(1, -1), 100, replace = TRUE) # treatment indicator (1 or -1)
pi <- runif(100, min = 0.3, max = 0.7) # propensity scores between 0 and 1

# Define XGBoost parameters
params <- list(
  max_depth = 3,
  eta = 0.1,
  subsample = 0.8,
  colsample_bytree = 0.8
)

# Train the model using A-learning loss
model_A <- XGBoostSub_con(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,
  Loss_type = "A_learning",
  params = params,
```

```

nrounds = 5,
disable_default_eval_metric = 1,
verbose = TRUE
)
subgroup_results=get_subgroup_results(model_A, X_data, subgroup_label=NULL, cutoff = 0.5)

```

---

predictive\_biomarker\_imp

*Plot Predictive Biomarker Importance based on XGBoost-based Subgroup Model*

---

### Description

This function calculates and plots the importance of biomarkers in a trained XGBoostSub\_con, XGBoostSub\_bin or XGBoostSub\_sur model.

### Usage

```
predictive_biomarker_imp(model)
```

### Arguments

model            The trained XGBoost-based model.

### Value

A barplot showing the biomarker importance.

### Examples

```

X_data <- matrix(rnorm(100 * 10), ncol = 10) # 100 samples with 10 features
y_data <- rnorm(100) # continuous outcome variable
trt <- sample(c(1, -1), 100, replace = TRUE) # treatment indicator (1 or -1)
pi <- runif(100, min = 0.3, max = 0.7) # propensity scores between 0 and 1

# Define XGBoost parameters
params <- list(
  max_depth = 3,
  eta = 0.1,
  subsample = 0.8,
  colsample_bytree = 0.8
)

# Train the model using A-learning loss
model_A <- XGBoostSub_con(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,

```

```

Loss_type = "A_learning",
params = params,
nrounds = 5,
disable_default_eval_metric = 1,
verbose = TRUE
)
biomarker_imp=predictive_biomarker_imp(model_A)

```

---

roc\_bin

*AUC ROC Table for Biomarkers Associated with Binary Outcomes*


---

### Description

Computes the area under the receiver operating characteristic (ROC) curve for Biomarkers Associated with Binary Outcomes, and returns the results as a table.

### Usage

```
roc_bin(yvar, xvars, dirs, data, yvar.display = yvar, xvars.display = xvars)
```

### Arguments

|               |   |
|---------------|---|
| yvar          | Binary response variable name, where 0 represents controls and 1 represents cases.  |
| xvars         | A vector of biomarker names.  |
| dirs          | A vector of directions for the biomarkers. Options are "auto", ">", or "<". - "auto" (default): automatically determines in which group the median is higher and takes the direction accordingly. - ">": indicates that the biomarkers for the control group are higher than those for the case group (controls > t >= cases). - "<": indicates that the biomarkers for the control group are lower or equal to those for the case group (controls < t <= cases). |
| data          | The dataset containing the variables.   |
| yvar.display  | Display name for the binary response variable.  |
| xvars.display | Display names for the biomarkers.   |

### Value

A table containing the AUC values for each biomarker.

### Examples

```

# Load a sample dataset
data <- data.frame(
  outcome = sample(c(0, 1), 100, replace = TRUE),
  biomarker1 = rnorm(100, mean = 0, sd = 1),
  biomarker2 = rnorm(100, mean = 5, sd = 2)
)

```

```
# Compute AUC for a single biomarker with auto direction
roc_bin(
  yvar = "outcome",
  xvars = "biomarker1",
  dirs = "auto",
  data = data,
  yvar.display = "Binary Outcome",
  xvars.display = "Biomarker 1"
)

# Compute AUC for multiple biomarkers with specified directions
roc_bin(
  yvar = "outcome",
  xvars = c("biomarker1", "biomarker2"),
  dirs = c("auto", "<"),
  data = data,
  yvar.display = "Binary Outcome",
  xvars.display = c("Biomarker 1", "Biomarker 2")
)
```

---

roc\_bin\_plot

*ROC Plot Biomarkers Associated with Binary Outcomes*

---

## Description

Generates ROC plots for different biomarkers associated with binary outcomes.

## Usage

```
roc_bin_plot(
  yvar,
  xvars,
  dirs,
  data,
  yvar.display = yvar,
  xvars.display = xvars
)
```

## Arguments

|       |   |
|-------|---|
| yvar  | Binary response variable name, where 0 represents controls and 1 represents cases.  |
| xvars | A vector of biomarker names.  |
| dirs  | A vector of directions for the biomarkers. Options are "auto", ">", or "<". - "auto" (default): automatically determines in which group the median is higher and takes the direction accordingly. - ">" indicates that the biomarkers for the control group are higher than those for the case group (controls > t >= cases). |

- "<" indicates that the biomarkers for the control group are lower or equal to those for the case group (controls < t <= cases).

data            The dataset containing the variables.

yvar.display    Display name for the binary response variable.

xvars.display   Display names for the biomarkers.

### Value

ROC plots for different biomarkers associated with binary outcomes.

### Examples

```
# Load a sample dataset
data <- data.frame(
  outcome = sample(c(0, 1), 100, replace = TRUE),
  biomarker1 = rnorm(100, mean = 0, sd = 1),
  biomarker2 = rnorm(100, mean = 5, sd = 2)
)

# Generate ROC plot for a single biomarker with auto direction
roc_bin_plot(
  yvar = "outcome",
  xvars = "biomarker1",
  dirs = "auto",
  data = data,
  yvar.display = "Binary Outcome",
  xvars.display = "Biomarker 1"
)

# Generate ROC plots for multiple biomarkers with specified directions
roc_bin_plot(
  yvar = "outcome",
  xvars = c("biomarker1", "biomarker2"),
  dirs = c("auto", "<"),
  data = data,
  yvar.display = "Binary Outcome",
  xvars.display = c("Biomarker 1", "Biomarker 2")
)
```

---

scat\_cont\_plot

*Scatter Plot for a Biomarker Associated with Continuous Outcome*

---

### Description

Generates a scatter plot for exploring the relationship between a continuous response variable and a biomarker variable.

**Usage**

```
scat_cont_plot(  
  yvar,  
  xvar,  
  data,  
  ybreaks = NULL,  
  xbreaks = NULL,  
  yvar.display = yvar,  
  xvar.display = xvar  
)
```

**Arguments**

|              |  |
|--------------|--|
| yvar         | Continuous response variable name.       |
| xvar         | biomarker name.                          |
| data         | The dataset containing the variables.    |
| ybreaks      | Breaks on the y-axis.                    |
| xbreaks      | Breaks on the x-axis.                    |
| yvar.display | Display name for the response variable.  |
| xvar.display | Display name for the biomarker variable. |

**Value**

A list containing correlation coefficients, scatter plot, slope, and intercept.

**Examples**

```
data <- data.frame(  
  outcome = rnorm(100, mean = 10, sd = 2),  
  biomarker = rnorm(100, mean = 0, sd = 1)  
)  
  
# Generate a scatter plot with default axis breaks  
scat_cont_plot(  
  yvar = "outcome",  
  xvar = "biomarker",  
  data = data,  
  yvar.display = "Continuous Outcome",  
  xvar.display = "Biomarker Level"  
)  
  
# Generate a scatter plot with specified axis breaks  
scat_cont_plot(  
  yvar = "outcome",  
  xvar = "biomarker",  
  data = data,  
  ybreaks = seq(5, 15, by = 1),  
  xbreaks = seq(-2, 2, by = 0.5),  
  yvar.display = "Continuous Outcome",
```

```
xvar.display = "Biomarker Level"
)
```

---

subgrp\_perf

*Subgroup Performance Evaluation for Prognostic Cases*


---

### Description

This function evaluates subgroup performance based on different types of response variables.

### Usage

```
subgrp_perf(
  yvar,
  censorvar = NULL,
  grpvar,
  grpname,
  xvars.adj = NULL,
  data,
  type,
  yvar.display = yvar,
  grpvar.display = grpvar
)
```

### Arguments

|                |  |
|----------------|--|
| yvar           | The response variable name.  |
| censorvar      | (Optional) The censoring variable name (0-censored, 1-event).                            |
| grpvar         | The subgroup variable name.  |
| grpname        | A vector of ordered subgroup names (values in the column of grpvar).                     |
| xvars.adj      | (Optional) Other covariates to adjust when evaluating the performance.                   |
| data           | The dataset containing the variables.  |
| type           | The type of response variable: "c" for continuous, "s" for survival, and "b" for binary. |
| yvar.display   | Display name of the response variable.   |
| grpvar.display | Display name of the group variable.  |

### Value

A list containing subgroup performance results including logrank p-value, median and mean survival, Cox model p-value, ANOVA p-value, and more based on the specified response variable type.



**Examples**

```
# Load a sample dataset
data <- data.frame(
  survival_time = rexp(100, rate = 0.1), # survival time
  status = sample(c(0, 1), 100, replace = TRUE), # censoring status
  group = sample(c("Low", "Medium", "High"), 100, replace = TRUE), # subgroup variable
  covariate = rnorm(100, mean = 50, sd = 10) # an additional covariate
)

# Perform subgroup performance evaluation for survival analysis
subgrp_perf(
  yvar = "survival_time",
  censorvar = "status",
  grpvar = "group",
  grpname = c("Low", "Medium", "High"),
  data = data,
  type = "s",
  yvar.display = "Survival Time",
  grpvar.display = "Risk Group"
)

# Perform subgroup performance evaluation for continuous outcome
data$continuous_outcome <- rnorm(100, mean = 10, sd = 5)
subgrp_perf(
  yvar = "continuous_outcome",
  grpvar = "group",
  grpname = c("Low", "Medium", "High"),
  data = data,
  type = "c",
  yvar.display = "Continuous Outcome",
  grpvar.display = "Risk Group"
)

# Perform subgroup performance evaluation for binary outcome
data$binary_outcome <- sample(c(0, 1), 100, replace = TRUE)
subgrp_perf(
  yvar = "binary_outcome",
  grpvar = "group",
  grpname = c("Low", "Medium", "High"),
  data = data,
  type = "b",
  yvar.display = "Binary Outcome",
  grpvar.display = "Risk Group"
)
```

**Description**

This function evaluates the performance of subgroups based on different types of response variables in predictive cases.

**Usage**

```
subgrp_perf_pred(
  yvar,
  censorvar = NULL,
  grpvar,
  grpname,
  trtvar,
  trtname,
  xvars.adj = NULL,
  data,
  type,
  yvar.display = yvar,
  grpvar.display = grpvar,
  trtvar.display = trtvar
)
```

**Arguments**

|                |   |
|----------------|---|
| yvar           | Response variable name.   |
| censorvar      | Censoring variable name (0-censored, 1-event).                        |
| grpvar         | Subgroup variable name.   |
| grpname        | A vector of ordered subgroup names (values in the column of grpvar).  |
| trtvar         | Treatment variable name.  |
| trtname        | A vector of ordered treatment names (values in the column of trtvar). |
| xvars.adj      | Other covariates to adjust when evaluating the performance.           |
| data           | The dataset.  |
| type           | "c" for continuous; "s" for "survival", and "b" for binary.           |
| yvar.display   | Display name of the response variable.                                |
| grpvar.display | Display name of the group variable.                                   |
| trtvar.display | Display name of the treatment variable.                               |

**Value**

A list containing the comparison results, group results, and possibly a plot.

**Examples**

```
# Load a sample dataset
data <- data.frame(
  response = rnorm(100, mean = 10, sd = 5), # continuous response
  survival_time = rexp(100, rate = 0.1), # survival time
```

```
status = sample(c(0, 1), 100, replace = TRUE), # censoring status
group = sample(c("Low", "Medium", "High"), 100, replace = TRUE), # subgroup variable
treatment = sample(c("A", "B"), 100, replace = TRUE) # treatment variable
)

# Subgroup performance evaluation for predictive cases - survival analysis
subgrp_perf_pred(
  yvar = "survival_time",
  censorvar = "status",
  grpvar = "group",
  grpname = c("Low", "Medium", "High"),
  trtvar = "treatment",
  trtname = c("A", "B"),
  data = data,
  type = "s",
  yvar.display = "Survival Time",
  grpvar.display = "Risk Group",
  trtvar.display = "Treatment"
)

# Subgroup performance evaluation for predictive cases - continuous outcome
subgrp_perf_pred(
  yvar = "response",
  grpvar = "group",
  grpname = c("Low", "Medium", "High"),
  trtvar = "treatment",
  trtname = c("A", "B"),
  data = data,
  type = "c",
  yvar.display = "Response",
  grpvar.display = "Risk Group",
  trtvar.display = "Treatment"
)

# Subgroup performance evaluation for predictive cases - binary outcome
data$binary_response <- sample(c(0, 1), 100, replace = TRUE)
subgrp_perf_pred(
  yvar = "binary_response",
  grpvar = "group",
  grpname = c("Low", "Medium", "High"),
  trtvar = "treatment",
  trtname = c("A", "B"),
  data = data,
  type = "b",
  yvar.display = "Binary Response",
  grpvar.display = "Risk Group",
  trtvar.display = "Treatment"
)
```

**Description**

A dataset containing sample data for demonstrating the functionalities of the BioPred package.

**Usage**

```
data(tutorial_data)
```

**Format**

A data frame with the following columns:

**x1** Numeric. A biomarker variable.

**x2** Numeric. A biomarker variable.

**x3** Numeric. A biomarker variable.

**x4** Numeric. A biomarker variable.

**x5** Numeric. A biomarker variable.

**x6** Numeric. A biomarker variable.

**x7** Numeric. A biomarker variable.

**x8** Numeric. A biomarker variable.

**x9** Numeric. A biomarker variable.

**x10** Numeric. A biomarker variable.

**y.con** Numeric. A continuous outcome variable.

**y.bin** Binary. A binary outcome variable, where 0 represents one class and 1 represents another class.

**y.time** Numeric. The time in months, used for survival analysis.

**y.event** Binary. Event indicator variable, where 0 indicates censoring and 1 indicates the event of interest occurred.

**subgroup\_label** Binary. Ground truth of subgroup label. In real-world scenarios, this information is typically unavailable.

**treatment** Binary. Treatment indicator variable, where 0 represents control and 1 represents treatment.

**treatment\_categorical** Factor. A categorical version of the treatment variable, with levels "Placebo" and "Treatment".

**risk\_category** Factor.

**Details**

This dataset is used to illustrate various functions within the BioPred package, including predictive modeling and subgroup analysis. The columns represent different types of data typically encountered in clinical studies.

**Examples**

```
data(tutorial_data)
head(tutorial_data)
```

---

|                |   |
|----------------|---|
| XGBoostSub_bin | <i>XGBoost Model with Modified Loss Function for Subgroup Identification with Binary Outcomes</i> |
|----------------|---|

---

### Description

Function for training XGBoost model with customized loss function for binary outcomes

### Usage

```
XGBoostSub_bin(
  X_data,
  y_data,
  trt,
  pi,
  Loss_type = "A_learning",
  params = list(),
  nrounds = 50,
  disable_default_eval_metric = 1,
  verbose = TRUE
)
```

### Arguments

|                             |  |
|-----------------------------|--|
| X_data                      | The input features matrix.   |
| y_data                      | The input y matrix.  |
| trt                         | The treatment indicator vector. Should take values of 1 or -1, where 1 represents the treatment group and -1 represents the control group. |
| pi                          | The propensity scores vector, which should range from 0 to 1, representing the probability of assignment to treatment.                     |
| Loss_type                   | Type of loss function to use: "A_learning" or "Weight_learning".   |
| params                      | A list of additional parameters for the xgb.train function.  |
| nrounds                     | Number of boosting rounds. Default is 50.  |
| disable_default_eval_metric | If 1, default evaluation metric will be disabled.  |
| verbose                     | Logical. If TRUE, training progress will be printed; if FALSE, no progress will be printed.  |

### Details

XGBoostSub\_bin: Function for Training XGBoost Model with Customized Loss Function for binary outcomes

This function trains an XGBoost model using a customized loss function based on the A-learning and weight-learning.

This function requires the 'xgboost' library. Make sure to install and load the 'xgboost' library before using this function.

After running this function, the returned model can be used like a regular xgboost model.

### Value

Trained XGBoostSub\_bin model.

### Examples

```
X_data <- matrix(rnorm(100 * 10), ncol = 10) # 100 samples with 10 features
y_data <- rbinom(100, 1, 0.5) # binary outcomes (0 or 1)
trt <- sample(c(1, -1), 100, replace = TRUE) # treatment indicator (1 or -1)
pi <- runif(100, min = 0.3, max = 0.7) # propensity scores between 0 and 1

# Define XGBoost parameters
params <- list(
  max_depth = 3,
  eta = 0.1,
  subsample = 0.8,
  colsample_bytree = 0.8
)

# Train the model using A-learning loss
model_A <- XGBoostSub_bin(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,
  Loss_type = "A_learning",
  params = params,
  nrounds = 5,
  disable_default_eval_metric = 1,
  verbose = TRUE
)

# Train the model using Weight-learning loss
model_W <- XGBoostSub_bin(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,
  Loss_type = "Weight_learning",
  params = params,
  nrounds = 5,
  disable_default_eval_metric = 1,
  verbose = TRUE
)
```

---

|                |   |
|----------------|---|
| XGBoostSub_con | <i>XGBoost Model with Modified Loss Function for Subgroup Identification with Continuous Outcomes</i> |
|----------------|---|

---

### Description

Function for training XGBoost model with customized loss function for continuous outcomes

### Usage

```
XGBoostSub_con(
  X_data,
  y_data,
  trt,
  pi,
  Loss_type = "A_learning",
  params = list(),
  nrounds = 50,
  disable_default_eval_metric = 1,
  verbose = TRUE
)
```

### Arguments

|                             |  |
|-----------------------------|--|
| X_data                      | The input features matrix.   |
| y_data                      | The input y matrix.  |
| trt                         | The treatment indicator vector. Should take values of 1 or -1, where 1 represents the treatment group and -1 represents the control group. |
| pi                          | The propensity scores vector, which should range from 0 to 1, representing the probability of assignment to treatment.                     |
| Loss_type                   | Type of loss function to use: "A_learning" or "Weight_learning".   |
| params                      | A list of additional parameters for the xgb.train function.  |
| nrounds                     | Number of boosting rounds. Default is 50.  |
| disable_default_eval_metric | If 1, default evaluation metric will be disabled.  |
| verbose                     | Logical. If TRUE, training progress will be printed; if FALSE, no progress will be printed.  |

### Details

XGBoostSub\_con: Function for Training XGBoost Model with Customized Loss Function for continuous outcomes

This function trains an XGBoost model using a customized loss function based on the A-learning and weight-learning.

This function requires the 'xgboost' library. Make sure to install and load the 'xgboost' library before using this function.

After running this function, the returned model can be used like a regular xgboost model.

### Value

Trained XGBoostSub\_con model.

### Examples

```
X_data <- matrix(rnorm(100 * 10), ncol = 10) # 100 samples with 10 features
y_data <- rnorm(100) # continuous outcome variable
trt <- sample(c(1, -1), 100, replace = TRUE) # treatment indicator (1 or -1)
pi <- runif(100, min = 0.3, max = 0.7) # propensity scores between 0 and 1

# Define XGBoost parameters
params <- list(
  max_depth = 3,
  eta = 0.1,
  subsample = 0.8,
  colsample_bytree = 0.8
)

# Train the model using A-learning loss
model_A <- XGBoostSub_con(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,
  Loss_type = "A_learning",
  params = params,
  nrounds = 5,
  disable_default_eval_metric = 1,
  verbose = TRUE
)

# Train the model using Weight-learning loss
model_W <- XGBoostSub_con(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,
  Loss_type = "Weight_learning",
  params = params,
  nrounds = 5,
  disable_default_eval_metric = 1,
  verbose = TRUE
)
```



---

|                |   |
|----------------|---|
| XGBoostSub_sur | <i>XGBoost Model with Modified Loss Function for Subgroup Identification with Survival Outcomes</i> |
|----------------|---|

---

### Description

Function for training XGBoost model with customized loss function for survival outcomes

### Usage

```
XGBoostSub_sur(
  X_data,
  y_data,
  trt,
  pi,
  censor,
  Loss_type = "Weight_learning",
  params = list(),
  nrounds = 50,
  disable_default_eval_metric = 1,
  verbose = TRUE
)
```

### Arguments

|                             |  |
|-----------------------------|--|
| X_data                      | The input features matrix.   |
| y_data                      | The input y matrix.  |
| trt                         | The treatment indicator vector. Should take values of 1 or -1, where 1 represents the treatment group and -1 represents the control group. |
| pi                          | The propensity scores vector, which should range from 0 to 1, representing the probability of assignment to treatment.                     |
| censor                      | The censor status vector. Should take values of 1 or 0, where 1 represents censoring and 0 represents an observed event.                   |
| Loss_type                   | Type of loss function to use: "A_learning" or "Weight_learning".   |
| params                      | A list of additional parameters for the xgb.train function.  |
| nrounds                     | Number of boosting rounds. Default is 50.  |
| disable_default_eval_metric | If 1, default evaluation metric will be disabled.  |
| verbose                     | Logical. If TRUE, training progress will be printed; if FALSE, no progress will be printed.  |

**Details**

XGBoostSub\_sur: Function for Training XGBoost Model with Customized Loss Function for survival outcomes

This function trains an XGBoost model using a customized loss function based on the A-learning and weight-learning.

This function requires the 'xgboost' library. Make sure to install and load the 'xgboost' library before using this function.

**Value**

Trained XGBoostSub\_sur model.

**Examples**

```
X_data <- matrix(rnorm(100 * 10), ncol = 10) # 100 samples with 10 features
y_data <- rexp(100, rate = 0.1) # survival times, simulated as exponential
trt <- sample(c(1, -1), 100, replace = TRUE) # treatment indicator (1 or -1)
pi <- runif(100, min = 0.3, max = 0.7) # propensity scores between 0 and 1
censor <- rbinom(100, 1, 0.7) # censoring indicator (1 = censored, 0 = observed)

# Define XGBoost parameters
params <- list(
  max_depth = 3,
  eta = 0.1,
  subsample = 0.8,
  colsample_bytree = 0.8
)

# Train the model using A-learning loss
model_A <- XGBoostSub_sur(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,
  censor = censor,
  Loss_type = "A_learning",
  params = params,
  nrounds = 5,
  disable_default_eval_metric = 1,
  verbose = TRUE
)

# Train the model using Weight-learning loss
model_W <- XGBoostSub_sur(
  X_data = X_data,
  y_data = y_data,
  trt = trt,
  pi = pi,
  censor = censor,
  Loss_type = "Weight_learning",
  params = params,
```

```
nrounds = 5,  
disable_default_eval_metric = 1,  
verbose = TRUE  
)
```

# Index

## \* datasets

tutorial\_data, [27](#)

cat\_summary, [2](#)

cdf\_plot, [3](#)

cut\_perf, [4](#)

eval\_metric\_bin, [6](#)

eval\_metric\_con, [7](#)

eval\_metric\_sur, [7](#)

fixcut\_bin, [8](#)

fixcut\_con, [10](#)

fixcut\_sur, [12](#)

gam\_ctr\_plot, [13](#)

gam\_plot, [15](#)

get\_subgroup\_results, [18](#)

predictive\_biomarker\_imp, [19](#)

roc\_bin, [20](#)

roc\_bin\_plot, [21](#)

scat\_cont\_plot, [22](#)

subgrp\_perf, [24](#)

subgrp\_perf\_pred, [25](#)

tutorial\_data, [27](#)

XGBoostSub\_bin, [29](#)

XGBoostSub\_con, [31](#)

XGBoostSub\_sur, [33](#)