

Package ‘PRIMsrc’

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Description Performs a unified treatment of Bump Hunting by Patient Rule Induction Method (PRIM) in Survival, Regression and Classification settings (SRC). The current version is a development release that only implements the case of a survival response. New features will be added soon as they are available.

Depends R (>= 3.0.2), survival, glmnet, superpc, Hmisc, quantreg

Imports parallel

NeedsCompilation yes

URL <https://github.com/jedazard/PRIMsrc>, <http://www.primsr.com>

Repository PRIMsrc, GitHub, Inc.

License GPL (>= 3) | file LICENSE

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LazyData yes

Archs i386, x64

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PRIMsrc-package	<i>Bump Hunting by Patient Rule Induction Method in Survival, Regression and Classification settings</i>
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Description

Performs a unified treatment of Bump Hunting by Patient Rule Induction Method (PRIM) in Survival, Regression and Classification settings (SRC). The method generates decision rules delineating a region in the predictor space, where the response is larger than its average over the entire space. The region is shaped as a hyperdimensional box or hyperrectangle that is not necessarily contiguous. Assumptions are that the multivariate input variables can be discrete or continuous and the univariate response variable can be discrete (Classification), continuous (Regression) or a time-to event, possibly censored (Survival). It is intended to handle low and high-dimensional multivariate datasets, including the situation where the number of covariates exceeds or dominates that of samples ($p > n$ or $p \gg n$ paradigm).

Details

The current version is a development release that only implements the case of a survival response. At this point, survival bump hunting is also restricted to a directed peeling search of the first box covered by the recursive coverage (outer) loop of our Patient Recursive Survival Peeling (PRSP) algorithm. New features will be added soon.

The package relies on an optional variable screening (pre-selection) procedure that is run before the PRSP algorithm and final variable usage (selection) procedure is done. This is done by four possible cross-validated variable screening (pre-selection) procedures offered to the user from the main end-user survival Bump Hunting function [sbh](#) (see below).

The following describes the end-user functions that are needed to run a complete procedure. The other internal subroutines are not documented in the manual and are not to be called by the end-user at any time. For computational efficiency, some end-user functions offer a parallelization option that is done by passing a few parameters needed to configure a cluster. This is indicated by an asterisk (* = optionally involving cluster usage). The R features are categorized as follows:

1. END-USER FUNCTION FOR PACKAGE NEWS
[PRIMsrc.news](#) **Display the PRIMsrc Package News**
 Function to display the log file NEWS of updates of the **PRIMsrc** package.
2. END-USER S3-METHOD FUNCTIONS FOR SUMMARY, DISPLAY, PLOT AND PREDICTION
[summary](#) **Summary Function**
 S3-method summary function to summarize the main parameters used to generate the sbh object.

print Print Function

S3-method print function to display the cross-validated estimated values of the sbh object.

plot 2D Visualization of Data Scatter and Box Vertices

S3-method plotting function for two-dimensional visualization of original or predicted data scatter as well as cross-validated box vertices of a sbh object. The scatter plot is for a given peeling step of the peeling sequence and a given plane, both specified by the user.

predict Predict Function

S3-method predict function to predict the box membership and box vertices on an independent set.

3. END-USER SURVIVAL BUMP HUNTING FUNCTION**sbh (*) Cross-Validated Survival Bump Hunting**

Main end-user function for fitting a cross-validated Survival Bump Hunting (SBH) model. Returns a cross-validated sbh object, as generated by our Patient Recursive Survival Peeling or PRSP algorithm, containing cross-validated estimates of end-points statistics of interest. The function relies on an optional internal variable screening (pre-selection) procedures that is run before the PRSP algorithm and final variable usage (selection) procedure is done. At this point, the user can choose between:

- (a) Univariate Patient Recursive Survival Peeling algorithm (default of package **PRIMsrc**)
- (b) Penalized Censored Quantile Regression (by Semismooth Newton Coordinate Descent algorithm adapted from package **hqreg**)
- (c) Penalized Partial Likelihood (adapted from package **glmnet**)
- (d) Supervised Principal Component Analysis (adapted from package **superpc**)

In this version, the Cross-Validation (CV) procedure and Bump Hunting procedures that control model size (#covariates) and model complexity (#peeling steps), respectively, to fit the Survival Bump Hunting model, are carried out internally by two consecutive tasks within the single main function `sbh()`. The returned S3-class sbh object contains cross-validated estimates of all the decision-rules of used covariates and all other statistical quantities of interest at each iteration of the peeling sequence (inner loop of the PRSP algorithm). This enables the graphical display of results of profiling curves for model selection/tuning, peeling trajectories, covariate traces and survival distributions (see plotting functions for more details). The function offers a number of options for the number of replications of the fitting procedure to be performed: *B*; the type of *K*-fold cross-validation desired: (replicated)-averaged or-combined; as well as the peeling and cross-validation criteria for model selection/tuning, and a few more parameters for the PRSP algorithm. The function takes advantage of the R packages **parallel** and **snow**, which allows users to create a parallel backend within an R session, enabling access to a cluster of compute cores and/or nodes on a local and/or remote machine(s) with either . The **PRIMsrc** supports two types of communication mechanisms between master and worker processes: 'Socket' or 'Message-Passing Interface' ('MPI').

4. END-USER PLOTTING FUNCTIONS FOR MODEL VALIDATION AND VISUALIZATION OF RESULTS**plot_profile Visualization for Model Selection/Validation**

Function for plotting the cross-validated model selection/tuning profiles of a sbh object. It uses the user's choice of cross-validation criterion statistics among the Log Hazard Ratio (LHR), Log-Rank Test (LRT) or Concordance Error Rate (CER). The function plots (as it applies) both profiles of cross-validation criterion as a function of variables screening size (cardinal subset of top-screened variables in the PRSP variable screening procedure), and peeling length (number of peeling steps of the peeling sequence in the inner loop of the PRSP

algorithm).

[plot_boxtraj](#) **Visualization of Peeling Trajectories/Profiles**

Function for plotting the cross-validated peeling trajectories/profiles of a sbh object. Applies to the user-specified covariates among the pre-selected ones and all other statistical quantities of interest at each iteration of the peeling sequence (inner loop of our PRSP algorithm).

[plot_boxtrace](#) **Visualization of Covariates Traces**

Function for plotting the cross-validated covariates traces of a sbh object. Plot the cross-validated modal trace curves of covariate importance and covariate usage of the user-specified covariates among the pre-selected ones at each iteration of the peeling sequence (inner loop of our PRSP algorithm).

[plot_boxkm](#) **Visualization of Survival Distributions**

Function for plotting the cross-validated survival distributions of a sbh object. Plot the cross-validated Kaplan-Meier estimates of survival distributions for the highest risk (inbox) versus lower-risk (outbox) groups of samples at each iteration of the peeling sequence (inner loop of our PRSP algorithm).

5. END-USER DATASETS

[Synthetic.1](#), [Synthetic.1b](#), [Synthetic.2](#), [Synthetic.3](#), [Synthetic.4](#) **Five Datasets From Simulated Regression Survival Models**

Five datasets from simulated regression survival models #1-4 as described in Dazard et al. (2015) representing low- and high-dimensional situations, and where regression parameters represent various types of relationship between survival times and covariates including saturated and noisy situations. In three datasets where non-informative noisy covariates were used, these covariates were not part of the design matrix (models #2-3 and #4). In one dataset, the signal is limited to a box-shaped region R of the predictor space (model #1b). In the last dataset, the signal is limited to 10% of the predictors in a $p > n$ situation (model #4). See each dataset for more details.

[Real.1](#) **Clinical Dataset**

Publicly available HIV clinical data from the Women's Interagency HIV cohort Study (WIHS). The entire study enrolled 1164 women. Inclusion criteria of the study are: women at enrolment must be (i) alive, (ii) HIV-1 infected, and (iii) free of clinical AIDS symptoms. Women were followed until the first of the following occurred: (i) treatment initiation (HAART), (ii) AIDS diagnosis, (iii) death, or administrative censoring. The studied outcomes were the competing risks "AIDS/Death (before HAART)" and "Treatment Initiation (HAART)". However, for simplification purposes, only the first of the two competing events (i.e. the time to AIDS/Death), was used. Likewise, for simplification in this clinical dataset example, only complete cases were used. Variables included history of Injection Drug Use ("IDU") at enrollment, African American ethnicity ('Race'), age ('Age'), and baseline CD4 count ('CD4'). The question in this dataset example was whether it is possible to achieve a prognostication of patients for AIDS and HAART. See Bacon et al. (2005) and the WIHS website for more details.

[Real.2](#) **Genomic Dataset**

Publicly available lung cancer genomic data from the Chemores Cohort Study. This data is part of an integrated study of mRNA, miRNA and clinical variables to characterize the molecular distinctions between squamous cell carcinoma (SCC) and adenocarcinoma (AC) in Non Small Cell Lung Cancer (NSCLC) aside large cell lung carcinoma (LCC). Tissue samples were analysed from a cohort of 123 patients, who underwent complete surgical resection at the Institut Mutualiste Montsouris (Paris, France) between 30 January 2002 and 26 June 2006. The studied outcome was the "Disease-Free Survival Time". Patients were followed until the first relapse occurred or administrative censoring. In this genomic dataset,

the expression levels of Agilent miRNA probes ($p = 939$) were included from the $n = 123$ cohort samples. In addition to the genomic data, five clinical variables, also evaluated on the cohort samples, are included as continuous variable ('Age') and nominal variables ('Type', 'KRAS.status', 'EGFR.status', 'P53.status'). This dataset represents a situation where the number of covariates dominates the number of complete observations, or $p \gg n$ case. See dataset for more details.

Known Bugs/Problems : None at this time.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

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References

- Dazard J-E. and Rao J.S. (2017). "Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods." (in prep).
- Yi C. and Huang J. (2016). "Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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- Dazard J-E., Choe M., LeBlanc M. and Rao J.S. (2014). "Cross-Validation of Survival Bump Hunting by Recursive Peeling Methods." In JSM Proceedings, Survival Methods for Risk Estimation/Prediction Section. Boston, MA, USA. American Statistical Association IMS - JSM, p. 3366-3380.
- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

See Also

- makeCluster (R package **parallel**)
- plot.survfit (R package **survival**)
- glmnet (R package **glmnet**)

Description

S3-method plot function for two-dimensional visualization of original data as well as predicted data scatter with cross-validated box vertices of a sbh object. The scatter plot is for a given peeling step of the peeling sequence and in a given plane of the used covariates of the sbh object, both specified by the user.

Usage

```
## S3 method for class 'sbh'
plot(x,
      main = NULL,
      proj = c(1,2),
      splom = TRUE,
      boxes = FALSE,
      steps = x$cvfit$cv.nsteps,
      pch = 16,
      cex = 0.5,
      col = 2:(length(steps)+1),
      col.box = 2:(length(steps)+1),
      lty.box = rep(2,length(steps)),
      lwd.box = rep(1,length(steps)),
      add.legend = TRUE,
      device = NULL,
      file = "Scatter Plot",
      path=getwd(),
      horizontal = FALSE,
      width = 5,
      height = 5, ...)
```

Arguments

x	Object of class sbh as generated by the main function sbh .
main	Character vector. Main Title. Defaults to NULL.
proj	Integer vector of length two, specifying the two dimensions of the projection plane of of the used covariates of the sbh object. Defaults to first two dimensions: {1,2}.
splom	Logical scalar. Shall the scatter plot of points inside the box(es) be plotted? Default to TRUE.
boxes	Logical scalar. Shall the box vertices be plotted or just the scatter of points? Default to FALSE.
steps	Integer vector. Vector of peeling steps at which to plot the in-box samples and box vertices. Defaults to the last peeling step of sbh object.
pch	Integer scalar of symbol number for the scatter plot. Defaults to 16.
cex	Integer scalar of symbol expansion. Defaults to 0.5.

col	Integer vector specifying the symbol color for each step. Defaults to vector of colors of length the number of steps. The vector is reused cyclically if it is shorter than the number of steps.
col.box	Integer vector of line color of box vertices for each step. Defaults to vector of colors of length the number of steps. The vector is reused cyclically if it is shorter than the number of steps.
lty.box	Integer vector of line type of box vertices for each step. Defaults to vector of 2's of length the number of steps. The vector is reused cyclically if it is shorter than the number of steps.
lwd.box	Integer vector of line width of box vertices for each step. Defaults to vector of 1's of length the number of steps. The vector is reused cyclically if it is shorter than the number of steps.
add.legend	Logical scalar. Shall the legend of steps numbers be plotted? Defaults to TRUE.
device	Graphic display device in {NULL, "PS", "PDF"}. Defaults to NULL (standard output screen). Currently implemented graphic display devices are "PS" (Postscript) or "PDF" (Portable Document Format).
file	File name for output graphic. Defaults to "Scatter Plot".
path	Absolute path (without final (back)slash separator). Defaults to working directory path.
horizontal	Logical scalar. Orientation of the printed image. Defaults to FALSE, that is potrait orientation.
width	Numeric scalar. Width of the graphics region in inches. Defaults to 5.
height	Numeric scalar. Height of the graphics region in inches. Defaults to 5.
...	Generic arguments passed to other plotting functions.

Details

The scatterplot is drawn on a graphical device with geometrically equal scales on the x and y axes.

Value

Invisible. None. Displays the plot(s) on the specified device.

Acknowledgments

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Note

End-user plotting function.

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- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

plot_boxkm

Visualization of Survival Distributions

Description

Function for plotting the cross-validated survival distributions of a sbh object. Plot the cross-validated Kaplan-Meier estimates of survival distributions for the highest risk (inbox) versus lower-risk (outbox) groups of samples at each iteration of the peeling sequence (inner loop of our PRSP algorithm).

Usage

```
plot_boxkm(object,
            main = NULL,
            xlab = "Time",
            ylab = "Probability",
            precision = 1e-3,
            mark = 3,
            col = 2,
            lty = 1,
            lwd = 0.5,
            cex = 0.5,
            steps = 1:object$cvfit$cv.nsteps,
            nr = 3,
            nc = 4,
            device = NULL,
            file = "Survival Plots",
            path=getwd(),
            horizontal = TRUE,
            width = 11,
            height = 8.5, ...)
```


Arguments

object	Object of class sbh as generated by the main function sbh .
main	Character vector. Main Title. Defaults to NULL.
xlab	Character vector. x-axis label. Defaults to "Time".
ylab	Character vector. y-axis label. Defaults to "Probability".
precision	Precision of cross-validated log-rank p-values of separation between two survival curves. Defaults to 1e-3.
mark	Integer scalar of mark parameter, which will be used to label the inbox and out-of-box curves. Defaults to 3.
col	Integer scalar specifying the color of the inbox curve. Defaults to 2.
lty	Integer scalar. Line type for the survival curve. Defaults to 1.
lwd	Numeric scalar. Line width for the survival curve. Defaults to 0.5.
cex	Numeric scalar specifying the size of the marks, symbol expansion used for titles, legends, and axis labels. Defaults to 0.5.
steps	Integer vector. Vector of peeling steps at which to plot the survival curves. Defaults to all the peeling steps of sbh object object.
nr	Integer scalar of the number of rows in the plot. Defaults to 3.
nc	Integer scalar of the number of columns in the plot. Defaults to 4.
device	Graphic display device in {NULL, "PS", "PDF"}. Defaults to NULL (standard output screen). Currently implemented graphic display devices are "PS" (Postscript) or "PDF" (Portable Document Format).
file	File name for output graphic. Defaults to "Survival Plots".
path	Absolute path (without final (back)slash separator). Defaults to the working directory path.
horizontal	Logical scalar. Orientation of the printed image. Defaults to TRUE, that is potrait orientation.
width	Numeric scalar. Width of the graphics region in inches. Defaults to 11.
height	Numeric scalar. Height of the graphics region in inches. Defaults to 8.5.
...	Generic arguments passed to other plotting functions, including <code>plot.survfit</code> (R package survival).

Details

Some of the plotting parameters are further defined in the function `plot.survfit` (R package **survival**). Step #0 always corresponds to the situation where the starting box covers the entire test-set data before peeling. Cross-validated LRT, LHR of inbox samples and log-rank p-values of separation are shown at the bottom of the plot with the corresponding peeling step. P-values are lower-bounded by the precision limit given by $1/A$, where A is the number of permutations.

Value

Invisible. None. Displays the plot(s) on the specified device.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

Note

End-user plotting function.

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- Dazard J-E., Choe M., LeBlanc M. and Rao J.S. (2016). "*Cross-validation and Peeling Strategies for Survival Bump Hunting using Recursive Peeling Methods*." Statistical Analysis and Data Mining, 9(1):12-42.
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- Dazard J-E. and J.S. Rao (2010). "*Local Sparse Bump Hunting*." J. Comp Graph. Statistics, 19(4):900-92.

See Also

- plot.survfit (R package **survival**)

plot_boxtrace

Visualization of Covariates Traces

Description

Function for plotting the cross-validated covariates traces of a sbh object. Plot the cross-validated modal trace curves of covariate importance and covariate usage of the pre-selected covariates specified by user at each iteration of the peeling sequence (inner loop of our PRSP algorithm).

Usage

```
plot_boxtrace(object,
              main = NULL,
              xlab = "Box Mass",
              ylab = "Covariate Range (centered)",
              topplot = object$cvfit$cv.used,
              center = TRUE,
              scale = FALSE,
              col.cov,
              lty.cov,
              lwd.cov,
              col = 1,
              lty = 1,
              lwd = 0.5,
              cex = 0.5,
              add.legend = FALSE,
              text.legend = NULL,
              device = NULL,
              file = "Covariate Trace Plots",
              path=getwd(),
              horizontal = FALSE,
              width = 8.5,
              height = 8.5, ...)
```

Arguments

object	Object of class sbh as generated by the main function sbh .
main	Character vector. Main Title. Defaults to.
xlab	Character vector. x-axis label. Defaults to "Box Mass". NULL
ylab	Character vector. y-axis label. Defaults to "Covariate Range (centered)".
topplot	Numeric vector. Which of the pre-selected covariates to plot (in reference to the original index of covariates). Defaults to covariates used for peeling.
center	Logical scalar. Shall the data be centered?. Defaults to TRUE.
scale	Logical scalar. Shall the data be scaled? Defaults to FALSE.
col.cov	Integer vector. Line color for the covariate importance curve of each selected covariate. Defaults to vector of colors of length the number of selected covariates. The vector is reused cyclically if it is shorter than the number of selected covariates.
lty.cov	Integer vector. Line type for the covariate importance curve of each selected covariate. Defaults to vector of 1's of length the number of selected covariates. The vector is reused cyclically if it is shorter than the number of selected covariates.
lwd.cov	Integer vector. Line width for the covariate importance curve of each selected covariate. Defaults to vector of 1's of length the number of selected covariates. The vector is reused cyclically if it is shorter than the number of selected covariates.
col	Integer scalar. Line color for the covariate trace curve. Defaults to 1.
lty	Integer scalar. Line type for the covariate trace curve. Defaults to 1.
lwd	Numeric scalar. Line width for the covariate trace curve. Defaults to 0.5.

cex	Numeric scalar. Symbol expansion used for titles, legends, and axis labels. Defaults to 0.5.
add.legend	Logical scalar. Should the legend be added to the current open graphics device?. Defaults to FALSE.
text.legend	Character vector of legend content. Defaults to NULL.
device	Graphic display device in {NULL, "PS", "PDF"}. Defaults to NULL (standard output screen). Currently implemented graphic display devices are "PS" (Postscript) or "PDF" (Portable Document Format).
file	File name for output graphic. Defaults to "Covariate Trace Plots".
path	Absolute path (without final (back)slash separator). Defaults to working directory path.
horizontal	Logical scalar. Orientation of the printed image. Defaults to FALSE, that is potrait orientation.
width	Numeric scalar. Width of the graphics region in inches. Defaults to 8.5.
height	Numeric scalar. Height of the graphics region in inches. Defaults to 8.5.
...	Generic arguments passed to other plotting functions.

Details

The trace plots limit the display of traces to those only covariates that are used for peeling. If centered, an horizontal black dotted line about 0 is added to the plot.

Due to the variability induced by cross-validation and replication, it is possible that more than one covariate be used for peeling at a given step. So, for simplicity of the trace plots, only the modal or majority vote trace value (over the folds and replications of the cross-validation) is plotted.

The top plot shows the overlay of covariate importance curves for each covariate. The bottom plot shows the overlay of covariate usage curves for each covariate. It is a discretized view of covariate importance.

Both point to the magnitude and order with which covariates are used along the peeling sequence.

Value

Invisible. None. Displays the plot(s) on the specified device.

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- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

plot_boxtraj

Visualization of Peeling Trajectories/Profiles

Description

Function for plotting the cross-validated peeling trajectories/profiles of a sbh object. Applies to the pre-selected covariates specified by user and all other statistical quantities of interest at each iteration of the peeling sequence (inner loop of our PRSP algorithm).

Usage

```
plot_boxtraj(object,
  main = NULL,
  topplot = object$cvfit$cv.used,
  col.cov,
  lty.cov,
  lwd.cov,
  col = 1,
  lty = 1,
  lwd = 0.5,
  cex = 0.5,
  add.legend = FALSE,
  text.legend = NULL,
  nr = NULL,
  nc = NULL,
  device = NULL,
  file = "Trajectory Plots",
  path=getwd(),
  horizontal = FALSE,
  width = 8.5,
  height = 11, ...)
```

Arguments

object	Object of class sbh as generated by the main function sbh .
main	Character vector. Main Title. Defaults to NULL.
toplot	Numeric vector. Which of the pre-selected covariates to plot (in reference to the original index of covariates). Defaults to covariates used for peeling.
col.cov	Integer vector. Line color for the covariate trajectory curve of each selected covariate. Defaults to vector of colors of length the number of selected covariates. The vector is reused cyclically if it is shorter than the number of selected covariates.
lty.cov	Integer vector. Line type for the covariate trajectory curve of each selected covariate. Defaults to vector of 1's of length the number of selected covariates. The vector is reused cyclically if it is shorter than the number of selected covariates.
lwd.cov	Integer vector. Line width for the covariate trajectory curve of each selected covariate. Defaults to vector of 1's of length the number of selected covariates. The vector is reused cyclically if it is shorter than the number of selected covariates.
col	Integer scalar. Line color for the trajectory curve of each statistical quantity of interest. Defaults to 1.
lty	Integer scalar. Line type for the trajectory curve of each statistical quantity of interest. Defaults to 1.
lwd	Numeric scalar. Line width for the trajectory curve of each statistical quantity of interest. Defaults to 0.5.
cex	Numeric scalar. Symbol expansion used for titles, legends, and axis labels. Defaults to 0.5.
add.legend	Logical scalar. Should the legend be added to the current open graphics device? Defaults to FALSE.
text.legend	Character vector of legend content. Defaults to NULL.
nr	Integer scalar of the number of rows in the plot. If NULL, defaults to 3.
nc	Integer scalar of the number of columns in the plot. If NULL, defaults to 3.
device	Graphic display device in {NULL, "PS", "PDF"}. Defaults to NULL (standard output screen). Currently implemented graphic display devices are "PS" (Postscript) or "PDF" (Portable Document Format).
file	File name for output graphic. Defaults to "Trajectory Plots".
path	Absolute path (without final (back)slash separator). Defaults to working directory path.
horizontal	Logical scalar. Orientation of the printed image. Defaults to FALSE, that is potrait orientation.
width	Numeric scalar. Width of the graphics region in inches. Defaults to 8.5.
height	Numeric scalar. Height of the graphics region in inches. Defaults to 11.
...	Generic arguments passed to other plotting functions.

Details

The plot limits the display of trajectories to those only covariates that are used for peeling.

The plot includes box descriptive statistics (such as support), survival endpoint statistics (such as Maximum Event-Free Time (MEFT), Minimum Event-Free Probability (MEVP), LHR, LRT) and prediction performance (such as CER).

Value

Invisible. None. Displays the plot(s) on the specified device.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

Note

End-user plotting function.

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References

- Dazard J-E. and Rao J.S. (2017). "*Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods*." (in prep).
- Yi C. and Huang J. (2016). "*Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression*." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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- Dazard J-E. and J.S. Rao (2010). "*Local Sparse Bump Hunting*." J. Comp Graph. Statistics, 19(4):900-92.

Description

Function for plotting the cross-validated model selection/tuning profiles of a sbh object. It uses the user's choice of cross-validation criterion statistics among the Log Hazard Ratio (LHR), Log-Rank Test (LRT) or Concordance Error Rate (CER). The function plots (as it applies) both profiles of cross-validation criterion as a function of variables screening size (cardinal subset of top-screened variables in the PRSP variable screening procedure), and peeling length (number of peeling steps of the peeling sequence in the inner loop of the PRSP algorithm).

Usage

```
plot_profile(object,
             main = NULL,
             xlim = NULL,
             ylim = NULL,
             add.sd = TRUE,
             add.legend = TRUE,
             add.profiles = TRUE,
             pch = 20,
             col = 1,
             lty = 1,
             lwd = 0.5,
             cex = 0.5,
             device = NULL,
             file = "Profile Plots",
             path=getwd(),
             horizontal = FALSE,
             width = 8.5,
             height = 5.0, ...)
```

Arguments

object	Object of class sbh as generated by the main function sbh .
main	Character vector. Main Title. Defaults to NULL.
xlim	Numeric vector of length 2. The x limits [x1, x2] of the plot. Defaults to NULL.
ylim	Numeric vector of length 2. The y limits [y1, y2] of the plot. Defaults to NULL.
add.sd	Logical scalar. Shall the standard error bars be plotted? Defaults to TRUE.
add.legend	Logical scalar. Shall the legend be plotted? Defaults to TRUE.
add.profiles	Logical scalar. Shall the individual profiles (for all replicates) be plotted? Defaults to TRUE.
pch	Integer scalar of symbol number for all the profiles. Defaults to 20.
col	Integer scalar of line color of the mean profile. Defaults to 1.
lty	Integer scalar of line type of the mean profile. Defaults to 1.
lwd	Numeric scalar of line width of the mean profile. Defaults to 0.5.
cex	Numeric scalar of symbol expansion for all the profiles. Defaults to 0.5.

device	Graphic display device in {NULL, "PS", "PDF"}. Defaults to NULL (standard output screen). Currently implemented graphic display devices are "PS" (Postscript) or "PDF" (Portable Document Format).
file	File name for output graphic. Defaults to "Profile Plot".
path	Absolute path (without final (back)slash separator). Defaults to working directory path.
horizontal	Logical scalar. Orientation of the printed image. Defaults to FALSE, that is potrait orientation.
width	Numeric scalar. Width of the graphics region in inches. Defaults to 8.5.
height	Numeric scalar. Height of the graphics region in inches. Defaults to 5.0.
...	Generic arguments passed to other plotting functions.

Details

Model tuning is done by applying the cross-validation criterion defined by the user's choice of specific statistic. The goal is to find the optimal value of model parameters by maximization of LHR or LRT, or minimization of CER. The parameters to optimize are (i) the cardinal of top-ranked variables subsets (if the "prsp" variable screening is chosen), and (ii) the number of peeling steps of the peeling sequence (inner loop of our PRSP algorithm) in any case of variable screening method.

Currently, this is done internally for visualization purposes, but it will ultimately offer the option to be done interactively with the end-user as well for parameter choosing/model selection.

Value

Invisible. None. Displays the plot(s) on the specified device.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

Note

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References

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- Dazard J-E., Choe M., LeBlanc M. and Rao J.S. (2016). "Cross-validation and Peeling Strategies for Survival Bump Hunting using Recursive Peeling Methods." Statistical Analysis and Data Mining, 9(1):12-42.
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- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

predict.sbh

Predict Function

Description

S3-method predict function to predict the box membership and box vertices on an independent set, using a cross-validated sbh fitted object.

Usage

```
## S3 method for class 'sbh'
predict(object,
        newdata,
        steps,
        na.action = na.omit, ...)
```

Arguments

object	Object of class sbh as generated by the main function sbh .
newdata	Either a numeric matrix or numeric vector containing the new input data of same dimensionality as the final sbh object of used covariates. A vector will be transformed to a (#sample x 1) matrix.
steps	Integer vector. Vector of peeling steps at which to predict the box memberships and box vertices. Defaults to the last peeling step only.
na.action	A function to specify the action to be taken if NAs are found. The default action is na.omit, which leads to rejection of incomplete cases.
...	Further generic arguments passed to the predict function.

Value

List containing the following 2 fields:

boxind	Logical matrix of predicted box membership indicator (columns) by peeling steps (rows). TRUE = in-box, FALSE = out-of-box.
vertices	List of size the number of chosen peeling steps where each entry is a numeric matrix of predicted box vertices: lower and upper bounds (rows) by covariate (columns).

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Note

End-user predict function.

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- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

PRIMsrc.news

Display the **PRIMsrc** Package News

Description

Function to display the log file NEWS of updates of the **PRIMsrc** package.

Usage

```
PRIMsrc.news(...)
```

Arguments

... Further arguments passed to or from other methods.

Value

None.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

Note

End-user function.

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print.sbh

Print Function

Description

S3-method print function to display the cross-validated estimated values of the sbh object.

Usage

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

Arguments

x Object of class sbh as generated by the main function [sbh](#).
 ... Further generic arguments passed to the print function.

Value

Display of the cross-validated fitted values of its argument.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

Note

End-user print function.

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Real.1

Real Dataset #1: Clinical Dataset ($p < n$ case)

Description

Publicly available HIV clinical data from the Women's Interagency HIV cohort Study (WIHS). The entire study enrolled 1164 women. Inclusion criteria of the study are: women at enrolment must be (i) alive, (ii) HIV-1 infected, and (iii) free of clinical AIDS symptoms. Women were followed until the first of the following occurred: (i) treatment initiation (HAART), (ii) AIDS diagnosis, (iii) death, or administrative censoring. The studied outcomes were the competing risks "AIDS/Death (before HAART)" and "Treatment Initiation (HAART)". However, for simplification purposes, only the first of the two competing events (i.e. the time to AIDS/Death), was used. Likewise, for simplification in this clinical dataset example, only complete cases were used. Variables included history of Injection Drug Use ("IDU") at enrollment, African American ethnicity ('Race'), age ('Age'), and baseline CD4 count ('CD4'). The question in this dataset example was whether it is possible to achieve a prognostication of patients for AIDS and HAART. See Bacon et al. (2005) and the WIHS website for more details.

Usage

Real.1

Format

Dataset consists of a numeric data.frame containing $n = 485$ complete observations (samples) by rows and $p = 4$ clinical covariates by columns, not including the censoring indicator and (censored) time-to-event variables. It comes as a compressed Rda data file.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

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Source

See real data application in Dazard et al., 2015.

References

- Dazard J-E. and Rao J.S. (2017). "*Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods*." (in prep).
- Yi C. and Huang J. (2016). "*Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression*." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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- Dazard J-E. and J.S. Rao (2010). "*Local Sparse Bump Hunting*." J. Comp Graph. Statistics, 19(4):900-92.

See Also

statepiaps.jhsph.edu/wihs/

Real.2

*Real Dataset #2: Genomic Dataset ($p \gg n$ case)***Description**

Publicly available lung cancer genomic data from the Chemores Cohort Study. This data is part of an integrated study of mRNA, miRNA and clinical variables to characterize the molecular distinctions between squamous cell carcinoma (SCC) and adenocarcinoma (AC) in Non Small Cell Lung Cancer (NSCLC) aside large cell lung carcinoma (LCC). Tissue samples were analysed from a cohort of 123 patients, who underwent complete surgical resection at the Institut Mutualiste Montsouris (Paris, France) between 30 January 2002 and 26 June 2006. The studied outcome was the "Disease-Free Survival Time". Patients were followed until the first relapse occurred or administrative censoring. In this genomic dataset, the expression levels of Agilent miRNA probes ($p = 939$) were included from the $n = 123$ cohort samples. The miRNA data contains normalized expression levels. See below the paper by Lazar et al. (2013) and Array Express data repository for complete description of the samples, tissue preparation, Agilent array technology, and data normalization. In addition to the genomic data, five clinical variables, also evaluated on the cohort samples, are included as continuous variable ('Age') and nominal variables ('Type', 'KRAS.status', 'EGFR.status', 'P53.status'). This dataset represents a situation where the number of covariates dominates the number of complete observations, or $p \gg n$ case.

Usage

Real.2

Format

Dataset consists of a numeric data.frame containing $n = 123$ complete observations (samples) by rows and $p = 939$ genomic covariates by columns, not including the censoring indicator and (censored) time-to-event variables. It comes as a compressed Rda data file.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

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Source

See real data application in Dazard et al., 2015.

References

- Dazard J-E. and Rao J.S. (2017). "Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods." (in prep).
- Yi C. and Huang J. (2016). "Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

See Also

Array Express data repository at the European Bioinformatics Institute. Accession number: #E-MTAB-1134 (MIR). www.ebi.ac.uk/arrayexpress/
 CHEMORES Consortium and website. www.chemores.ki.se/index.html

 sbh

Cross-Validated Survival Bump Hunting

Description

Main end-user function for fitting a cross-validated Survival Bump Hunting (SBH) model. Returns a cross-validated sbh object, as generated by our Patient Recursive Survival Peeling (PRSP) algorithm, containing cross-validated estimates of end-points statistics of interest. Generates an object of class sbh.

Usage

```
sbh(X,
    y,
    delta,
    B = 30,
    K = 5,
    A = 1000,
    vs = TRUE,
    vstype = "ppl",
    vsarg = "alpha=1,
          nalphabet=1,
          nlambd=100,
```

```

        vscons="0.5",
cv = TRUE,
cvtype = "combined",
cvarg = "alpha=0.01,
        beta=0.05,
        minn=5,
        L=NULL,
        peelcriterion="lrt",
        cvcriterion="cer",
pv = FALSE,
decimals = 2,
onese = FALSE,
probval = NULL,
timeval = NULL,
parallel.vs = FALSE,
parallel.rep = FALSE,
parallel.pv = FALSE,
conf = NULL,
verbose = TRUE,
seed = NULL)

```

Arguments

X	($n \times p$) data.frame or numeric matrix of n observations and p input covariates. If a data.frame is provided, it will be coerced to a numeric matrix. Discrete nominal covariates will be treated as ordinal variables.
y	n -numeric vector of observed times to event.
delta	n -numeric vector of observed status (censoring) indicator variable.
B	Positive integer of the number of replications of the cross-validation procedure. Defaults to 30.
K	Positive integer of the number of cross-validation folds (partitions) into which the observations (n) should be randomly split. K must be bigger than 2 for a regular K -fold cross-validation procedure to work and should be greater than 3 for a regular procedure to make sense; $K \in \{5, \dots, 10\}$ is recommended; defaults to $K=5$. Setting K also specifies the type of cross-validation to be done: <ul style="list-style-type: none"> • $K = 1$ carries no cross-validation out, or set-value when $cv = FALSE$ (see below). • $K \in \{2, \dots, n - 1\}$ carries out K-fold cross-validation. • $K = n$ carries out leave-one-out cross-validation.
A	Positive integer of the number of permutations for the computation of cross-validated p-values. Defaults to 1000. Ignored if pv is FALSE.
vs	logical scalar. Flag for optional variable (covariate) screening (pre-selection). Defaults to TRUE.
vstype	character vector in {"prsp", "pcqr", "ppl", "spca"} specifying the variable screening (pre-selection) procedure. Defaults to "ppl". Reset to NA if vs is FALSE.
vsarg	Character vector specifying the parameters of cross-validated variable screening (pre-selection) procedure. Defaults to PPL with its suggested parameters values. Note that $vsarg$ comes as a characters string (between double quotes), with comas separated values without white spaces. All the following parameters

are ignored if `vs` is FALSE

PRSP:

- `alpha` = fraction to peel off at each step. Suggests 0.01.
- `beta` = minimum support size resulting from the peeling sequence. Suggests 0.05.
- `minn` = minimum number of observation that we want to be able to detect in a box. Suggests 5.
- `L` = maximum peeling length in $[1, \text{ceiling}(\log(1/n) / \log(1 - (1/n)))]$. See details below. Suggests $\text{ceiling}(\log(\text{beta}) / \log(1 - \text{alpha})/3)$.
- `S` = maximum variables screening size in $[1, p]$ (i.e. the maximum cardinal subset of top-screened variables), that is used for the cross-validation of the optimal number of screened variables. Setting `S` to NULL will create a grid of $\min(S, \text{floor}(100 * S/p))$ values in $[1, \text{floor}(p/10)]$ that will be used to determine the optimal number of screened variables. Setting `S` to a single upper bound value in $[1, p]$ will fix the number of screened variables up to it. Suggests $S = \text{floor}(p/10)$ to reduce computational complexity.
- `peelcriterion` in `{"lhr", "lrt", "chs"}` standing for Log-Hazard Ratio (LHR), Log-Rank Test (LRT), and Cumulative Hazard Summary (CHS), respectively, specifying the peeling criterion used in the PRSP algorithm. Suggests "lrt".
- `cvcriterion` in `{"lhr", "lrt", "cer"}` standing for Log-Hazard Ratio (LHR), Log-Rank Test (LRT), and Concordance Error Rate (CER), respectively, specifying the cross-validation criterion used for tuning/optimizing the maximum variables screening size in the PRSP variable screening procedure. Suggests "cer".
- `vscons` = numeric scalar in $[1/K, 1]$, specifying the conservativeness of the variable screening (pre-selection) procedure, where $1/K$ is the least conservative and 1 is the most. Suggests 0.5.

PCQR:

- `tau` = quantile in $[0, 0.5]$ used in the censored quantile regression model. It is the tuning parameter of the censored quantile loss. It represents the conditional censored quantile of the survival response to be estimated. It includes the absolute loss when `tau`=0.5. Suggests 0.5.
- `alpha` = elasticnet mixing parameter in $[0, 1]$ that controls the relative contribution from the lasso and the ridge penalty. The penalty is defined as $(1-\text{alpha})/2\|\text{betall}_2\|^2 + \text{alpha}\|\text{betall}_1\|$. `alpha` = 1 is the lasso penalty, and `alpha` = 0 the ridge penalty. If `alpha` is set to NULL, a vector of values of length `nalpha` is used, else `alpha` value is used and `nalpha` is set to 1. Suggests `alpha`=1 (lasso).
- `nalpha` = number of elasticnet penalization `alpha` values to consider in the grid search. Suggests 1 (see above: lasso).
- `nlambd` = number of elasticnet penalization `lambda` values to consider in the grid search. Suggests 100.
- `vscons` = numeric scalar in $[1/K, 1]$, specifying the conservativeness of the variable screening (pre-selection) procedure, where $1/K$ is the least conservative and 1 is the most. Suggests 0.5.

PPL:

- `alpha` = elasticnet mixing parameter in $[0, 1]$ that controls the relative contribution from the lasso and the ridge penalty. See R package **glmnet**. The

penalty is defined as $(1-\alpha)/2\|\beta\|_2^2 + \alpha\|\beta\|_1$. $\alpha = 1$ is the lasso penalty, and $\alpha = 0$ the ridge penalty. If α is set to NULL, a vector of values of length `nalpha` is used, else α value is used and `nalpha` is set to 1. Suggests $\alpha=1$ (lasso).

- `nalpha` = number of elasticnet penalization α values to consider in the grid search. Suggests 1 (see above: lasso).
- `nlambda` = number of elasticnet penalization λ values to consider in the grid search. Suggests 100.
- `vscons` = numeric scalar in $[1/K, 1]$, specifying the conservativeness of the variable screening (pre-selection) procedure, where $1/K$ is the least conservative and 1 is the most. Suggests 0.5.

SPCA:

- `n.thres` = number of thresholds to consider in the grid search. It cannot be less than n (sample size). Suggests 20.
- `n.pcs` = number of cross-validation principal components to use in $\{1, 2, 3\}$. It cannot be less than n (sample size) and more than p (dimensionality), and will be reset to `n.pcs` = $p - 1$ otherwise. Suggests 3.
- `n.var` = minimum number of variables to include in determining range for threshold. It cannot be more than p (dimensionality), and will be reset to `n.var` = $p - 1$ otherwise. Suggests 5.
- `vscons` = numeric scalar in $[1/K, 1]$, specifying the conservativeness of the variable screening (pre-selection) procedure, where $1/K$ is the least conservative and 1 is the most. Suggests 0.5.

<code>cv</code>	logical scalar. Flag for optional cross-validation (CV) of parameters of variable screening (pre-selection) and variable usage (selection) by PRSP algorithm. Defaults to TRUE. If FALSE, no cross-validation at will be performed, the value of K will be overwritten to 1, and traditional log-rank Mantel-Haenszel p -values will be computed, (using the Chi-Squared distribution with 1df for the null distribution) instead of log-rank permutation p -values (using the permutation distribution for the null distribution).
<code>cvtype</code>	character vector in $\{"combined", "averaged"\}$ specifying the cross-validation technique. Defaults to "combined". Reset to NA if <code>cv</code> is FALSE.
<code>cvarg</code>	character vector describing the parameters used in the PRSP algorithm of the Survival Bump Hunting function. Defaults to <code>"alpha=0.01,beta=0.05,minn=5,L=NULL,peelcriterion=\"lrt\",cvcriterion=\"cer\""</code> . Note that <code>cvarg</code> comes as a characters string (between double quotes), with commas separated values without white spaces. <ul style="list-style-type: none"> • <code>alpha</code> = fraction to peel off at each step. Defaults to 0.01. • <code>beta</code> = minimum support size resulting from the peeling sequence. Defaults to 0.05. • <code>minn</code> = minimum number of observation that we want to be able to detect in a box. Defaults to 5. • <code>L</code> = maximum peeling length in $[1, \lceil \log(1/n) / \log(1 - (1/n)) \rceil]$. See details below. Defaults to NULL, that is, with automatic selection. • <code>peelcriterion</code> in $\{"lhr", "lrt", "chs"\}$ standing for Log-Hazard Ratio (LHR), Log-Rank Test (LRT), and Cumulative Hazard Summary (CHS), respectively, specifying the peeling criterion used in the PRSP algorithm. Defaults to "lrt".

	<ul style="list-style-type: none"> • <code>cvcriterion</code> in {"lhr", "lrt", "cer"} standing for Log-Hazard Ratio (LHR), Log-Rank Test (LRT), and Concordance Error Rate (CER), respectively, specifying the cross-validation criterion used for tuning/optimizing the peeling sequence length (i.e. number of peeling steps) in the PRSP algorithm. Defaults to "cer". Ignored if <code>cv</code> is FALSE.
<code>pv</code>	logical scalar. Flag for computation of log-rank p-values. Defaults to FALSE.
<code>decimals</code>	Positive integer of the number of user-specified significant decimals to output results. Defaults to 2.
<code>onese</code>	logical scalar. Flag for using the 1-standard error rule instead of extremum value of the cross-validation criterion when tuning/optimizing model parameters. Defaults to FALSE.
<code>probval</code>	numeric scalar of the survival probability at which we want to get the endpoint box survival time. Defaults to NULL.
<code>timeval</code>	numeric scalar of the survival time at which we want to get the endpoint box survival probability. Defaults to NULL.
<code>parallel.vs</code>	logical. Is parallelization to be performed for variable screening? Defaults to FALSE, because it is not implemented yet.
<code>parallel.rep</code>	logical. Is parallelization to be performed for replications? Defaults to FALSE.
<code>parallel.pv</code>	logical. Is parallelization to be performed for computation of log-rank p-values? Defaults to FALSE.
<code>conf</code>	list of 5 fields containing the parameters values needed for creating the parallel backend (cluster configuration). See details below for usage. Optional, defaults to NULL, but all fields are required if used: <ul style="list-style-type: none"> • <code>type</code> : character vector specifying the cluster type ("SOCKET", "MPI"). • <code>spec</code> : A specification (character vector or integer scalar) appropriate to the type of cluster. • <code>homogeneous</code> : logical scalar to be set to FALSE for inhomogeneous clusters. • <code>verbose</code> : logical scalar to be set to FALSE for quiet mode. • <code>outfile</code> : character vector of an output log file name to direct the stdout and stderr connection output from the workernodes. "" indicates no redirection.
<code>verbose</code>	logical scalar. Is the output to be verbose? Optional, defaults to TRUE.
<code>seed</code>	Positive integer scalar of the user seed to reproduce all the results. Defaults to NULL.

Details

At this point, the main function `sbh` relies on an optional variable screening (pre-selection) procedure that is run before the variable usage (selection) procedure is done by our PRSP algorithm. User can choose between four possible procedures:

- Patient Recursive Survival Peeling (PRSP) (by univariate screening of our algorithm)
- Penalized Censored Quantile Regression (PCQR) (by Semismooth Newton Coordinate Descent fitting algorithm adapted from package **hqreg**)
- Penalized Partial Likelihood (PPL) (by Elasticnet Regularization adapted from package **glmnet**)

- Supervised Principal Component Analysis (SPCA) (by Supervised Principal Component adapted from package **superpc**)

There is no default, but it is recommended to use PPL or SPCA for computational efficiency. Variable screening (pre-selection) is done by computing occurrence frequencies of top-ranking variables over the cross-validation folds and replicates. The conservativeness of the procedure is controlled by the argument `vscons`. Example of calls for pre-selection are as follows:

- '1.0' represents a presence in all the folds (unanimity vote)
- '0.5' represents a presence in at least half of the folds (majority vote)
- '1/K' represents a presence in at least one of the folds (minority vote)

Although any value in the interval $[1/K, 1]$ is accepted, we recommend using the interval $[1/K, 1/2]$ to avoid excessive conservativeness. Final variable usage (selection) is done after running our PRSP algorithm on previously screened variables by collecting those variables that have the maximum occurrence frequency in each peeling step over cross-validation folds and replicates.

In the PRSP algorithm, the maximal number of peeling steps is determined either by `alpha` and `beta` metaparameters or the smallest possible fraction of the training data, i.e. $\frac{1}{n}$:

- `ceiling(log(beta) / log(1 - alpha))`: `alpha` and `beta` are fixed by user
- `ceiling(log(1/n) / log(1 - alpha))`: `alpha` is fixed by user and `beta` is fixed by data
- `ceiling(log(beta) / log(1 - (1/n)))`: `alpha` is fixed by data and `beta` is fixed by user
- `ceiling(log(1/n) / log(1 - (1/n)))`: `alpha` and `beta` are fixed by data

If `L` is not used to specify a fixed number of peeling steps (i.e. `NULL`), then `beta` and `minn` are used in the stopping rule instead.

If a cross-validation is requested, the function performs a supervised (stratified) random splitting of the data based on the outcome, which is in that case the `delta` argument. This is because it is desirable that the data splitting balances the class distributions of the outcome (events) within the cross-validation splits. For each screening method and for building (by PRSP algorithm) the final Survival Bump Hunting (SBH) model, all model tuning parameters are simultaneously estimated by cross-validation. The function offers a number of options for the cross-validation to be performed: the number of replications `B`; the type of technique; the peeling criterion; and the optimization criterion.

The returned S3-class `sbh` object contains cross-validated estimates of all the decision-rules of used (selected) covariates and all other statistical quantities of interest at each iteration of the peeling sequence (inner loop of the PRSP algorithm). This enables the graphical display of results of profiling curves for model tuning, peeling trajectories, covariate traces and survival distributions (see plotting functions for more details).

In case replicated cross-validations are performed, a "summary report" of the outputs is done over the `B` replicates as follows:

- Even though the PRSP algorithm uses only one covariate at a time at each peeling step, the reported matrix of "Replicated CV" box decision rules may show more than one covariate being used in a given step, because these decision rules are averaged over the `B` replicates (see equation #21 in Dazard et al. 2016).
- However, the reported "Replicated CV" trace values are computed (at each peeling step) as a *single* modal trace value of covariate usage over the `B` replicates. This is also reflected in the reported "Replicated CV" importance and usage plots of covariate traces.
- The reported "Replicated CV" box membership indicators are computed (at each peeling step) as the point-wise majority vote over the `B` replicates (right-hand side of equation #22 in Dazard et al. 2016).

- The reported "Replicated CV" box support vector and corresponding box sample size are computed (at each peeling step) based on the above "Replicated CV" box membership indicators (i.e. *not* as equation #23 in Dazard et al. 2016).

If the computation of log-rank p -values is desired, then running with the parallelization option is strongly advised as it may take a while. In case of large ($p > n$) or very large ($p \gg n$) datasets, it is also highly recommended to use the parallelization option.

The function `sbh` relies on the R package **parallel** to create a parallel backend within an R session, enabling access to a cluster of compute cores and/or nodes on a local and/or remote machine(s) and scaling-up with the number of CPU cores available and efficient parallel execution. To run a procedure in parallel (with parallel RNG), argument `parallel` is to be set to `TRUE` and argument `conf` is to be specified (i.e. non `NULL`). Argument `conf` uses the options described in function `makeCluster` of the R packages **parallel** and **snow**. **PRIMsrc** supports two types of communication mechanisms between master and worker processes: 'Socket' or 'Message-Passing Interface' ('MPI'). In **PRIMsrc**, parallel 'Socket' clusters use sockets communication mechanisms only (no forking) and are therefore available on all platforms, including Windows, while parallel 'MPI' clusters use high-speed interconnects mechanism in networks of computers (with distributed memory) and are therefore available only in these architectures. A parallel 'MPI' cluster also requires R package **Rmpi** to be installed. Value type is used to setup a cluster of type 'Socket' ("SOCKET") or 'MPI' ("MPI"), respectively. Depending on this type, values of `spec` are to be used alternatively:

- For 'Socket' clusters (`conf$type="SOCKET"`), `spec` should be a character vector naming the hosts on which to run the job; it can default to a unique local machine, in which case, one may use the unique host name "localhost". Each host name can potentially be repeated to the number of CPU cores available on the local machine. It can also be an integer scalar specifying the number of processes to spawn on the local machine; or a list of machine specifications if you have `ssh` installed (a character value named `host` specifying the name or address of the host to use).
- For 'MPI' clusters (`conf$type="MPI"`), `spec` should be an integer scalar specifying the total number of processes to be spawned across the network of available nodes, counting the `workernodes` and `masternode`.

The actual creation of the cluster, its initialization, and closing are all done internally. For more details, see the reference manual of R package **snow** and examples below.

When random number generation is needed, the creation of separate streams of parallel RNG per node is done internally by distributing the stream states to the nodes. For more details, see the vignette of R package **parallel**. The use of a seed allows to reproduce the results within the same type of session: the same seed will reproduce the same results within a non-parallel session or within a parallel session, but it will not necessarily give the exact same results (up to sampling variability) between a non-parallelized and parallelized session due to the difference of management of the seed between the two (see parallel RNG and value of returned seed below).

Value

Object of class `sbh` (Patient Recursive Survival Peeling) list containing the following 21 fields:

<code>X</code>	numeric matrix of original dataset.
<code>y</code>	numeric vector of observed failure / survival times.
<code>delta</code>	numeric vector of observed event indicator in {1,0}.
<code>B</code>	positive integer of the number of replications used in the cross-validation procedure.
<code>K</code>	positive integer of the number of folds used in the cross-validation procedure.

A	positive integer of the number of permutations used for the computation of log-rank p-values.
vs	logical scalar of returned flag of optional variable pre-selection.
vstype	character vector of the optional variable pre-selection procedure used.
vsarg	character vector of the parameters used in the pre-selection procedure.
cv	logical scalar of returned flag of optional cross-validation.
cvtype	character vector of the cross-validation technique used.
cvarg	character vector of the parameters used in the Survival Bump Hunting procedure.
pv	logical scalar of returned flag of optional computation of log-rank p-values.
onese	logical scalar of returned flag of 1-standard error rule.
decimals	integer of the number of user-specified significant decimals.
probval	Numeric scalar of survival probability used.
timeval	Numeric scalar of survival time used.
cvprofiles	list of 10 fields of cross-validated tuning profiles and estimates, each of length B (one for each replicate): <ul style="list-style-type: none"> • cv.varprofiles: numeric matrix of cross-validation criterion used for tuning/optimizing the variable screening size in the PRSP variable screening (pre-selection) procedure (NULL otherwise). Values are by columns (peeling steps) and replicates (rows). • cv.varprofiles.mean: numeric vector of means (across replicates) of the above cross-validation criterion by peeling steps. • cv.varprofiles.se: numeric vector of standard errors (across replicates) of the above cross-validation criterion by peeling steps. • cv.varset.opt: numeric scalar of optimal variable screening size according to the extremum. • cv.varset.lse: numeric scalar of optimal variable screening size according to 1SE rule. • cv.stepprofiles: numeric matrix of cross-validation criterion used for tuning/optimizing the peeling sequence length (i.e. number of peeling steps) in the PRSP algorithm. Values are by columns (peeling steps) and replicates (rows). • cv.stepprofiles.mean: numeric vector of means (across replicates) of the above cross-validation criterion by peeling steps. • cv.stepprofiles.se: numeric vector of standard errors (across replicates) of the above cross-validation criterion by peeling steps. • cv.nsteps.opt: numeric scalar of optimal number of peeling steps according to the extremum. • cv.nsteps.lse: numeric scalar of optimal number of peeling steps according to 1SE rule.
cvfit	list with 12 fields of cross-validated SBH output estimates, each of length B (one for each replicate): <ul style="list-style-type: none"> • cv.maxsteps: numeric scalar of maximal number of peeling steps over the replicates. • cv.nsteps: numeric scalar of optimal number of peeling steps according to the optimization criterion.

	<ul style="list-style-type: none"> • <code>cv.boxind</code>: logical matrix in TRUE, FALSE of individual observation box membership indicator (columns) for all peeling steps (rows). • <code>cv.boxind.size</code>: numeric vector of box sample size for all peeling steps. • <code>cv.boxind.support</code>: numeric vector of box support for all peeling steps. • <code>cv.rules</code>: data.frame of decision rules on the covariates (columns) for all peeling steps (rows). • <code>cv.screened</code>: numeric vector of screened (pre-selected) covariates, indexed in reference to original index. • <code>cv.trace</code>: numeric vector of the modal trace values of covariate usage for all peeling steps. • <code>cv.sign</code>: numeric vector in $\{-1, +1\}$ of directions of peeling for all used (selected) covariates. • <code>cv.used</code>: numeric vector of covariates used (selected) for peeling, indexed in reference to original index. • <code>cv.stats</code>: numeric matrix of box endpoint quantities of interest (columns) for all peeling steps (rows). • <code>cv.pval</code>: list with 2 fields of two vectors. The first pval is a numeric vector for log-rank p-values of separation of survival distributions, The second seed is is an integer scalar if parallelization is used, or an integer vector of A values, one for each permutation, if parallelization is not used.
success	logical scalar of the returned flag of success at fitting the SBH model.
seed	User seed. An integer scalar if parallelization is used, or an integer vector of B values, one for each replication, if parallelization is not used.

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Note

Unique end-user function for fitting the Survival Bump Hunting model.

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See Also

- makeCluster (R package **parallel**)
- glmnet, cv.glmnet (R package **glmnet**)
- hqreg, cv.hqreg (R package **hqreg**)
- superpc.cv (R package **superpc**)

Examples

```
#####
# Loading the library and its dependencies
#####
library("PRIMsrc")

## Not run:
#####
# PRIMsrc Package news
#####
PRIMsrc.news()

#####
# PRIMsrc Package citation
#####
citation("PRIMsrc")

#####
# Demo with a synthetic dataset
# Use help for descriptions
#####
data("Synthetic.1", package="PRIMsrc")
?Synthetic.1

## End(Not run)

#####
# Simulated dataset #1 (n=250, p=3)
# Peeling criterion = LRT
# Cross-Validation criterion = LRT
# With Combined Cross-Validation (RCCV)
# Without Replications (B = 1)
```

```

# Without variable screening (pre-selection)
# Without computation of log-rank permutation p-values
# Without parallelization
#=====
synt1 <- sbh(X = Synthetic.1[ , -c(1,2), drop=FALSE],
            y = Synthetic.1[ ,1, drop=TRUE],
            delta = Synthetic.1[ ,2, drop=TRUE],
            B = 1,
            K = 3,
            vs = FALSE,
            cv = TRUE,
            cvtype = "combined",
            cvarg = "alpha=0.10,
                    beta=0.05,
                    minn=5,
                    L=NULL,
                    peelcriterion=\"lrt\",
                    cvcriterion=\"lrt\"",
            pv = FALSE,
            decimals = 2,
            onese = FALSE,
            probval = 0.5,
            timeval = NULL,
            parallel.vs = FALSE,
            parallel.rep = FALSE,
            parallel.pv = FALSE,
            conf = NULL,
            verbose = FALSE,
            seed = 123)

summary(object = synt1)
print(x = synt1)

n <- 100
p <- length(synt1$cvfit$cv.used)
x <- matrix(data = runif(n = n*p, min = 0, max = 1),
            nrow = n, ncol = p, byrow = FALSE,
            dimnames=list(1:n, paste("X", 1:p, sep="")))
synt1.pred <- predict(object = synt1,
                     newdata = x,
                     steps = synt1$cvfit$cv.nsteps)

plot(x = synt1,
     main = paste("Scatter plot for model #1", sep=""),
     proj = c(1,2), splom = TRUE, boxes = TRUE,
     steps = synt1$cvfit$cv.nsteps,
     pch = 16, cex = 0.5, col = 2,
     col.box = 2, lty.box = 2, lwd.box = 1,
     add.legend = TRUE, device = NULL)

plot_profile(object = synt1,
             main = "Cross-validated tuning profiles for model #1",
             pch=20, col=1, lty=1, lwd=0.5, cex=0.5,
             add.sd = TRUE, add.legend = TRUE, add.profiles = TRUE,
             device = NULL, file = "Profile Plot", path=getwd(),
             horizontal = FALSE, width = 8.5, height = 5.0)

```

```

plot_boxtraj(object = synt1,
             main = paste("Cross-validated peeling trajectories for model #1", sep=""),
             col=1, lty=1, lwd=0.5, cex=0.5,
             topplot = synt1$cvfit$cv.used,
             device = NULL, file = "Trajectory Plots", path=getwd(),
             horizontal = FALSE, width = 8.5, height = 8.5)

plot_boxtrace(object = synt1,
              main = paste("Cross-validated trace plots for model #1", sep=""),
              xlab = "Box Mass", ylab = "Covariate Range (centered)",
              col=1, lty=1, lwd=0.5, cex=0.5,
              topplot = synt1$cvfit$cv.used,
              center = TRUE, scale = FALSE,
              device = NULL, file = "Covariate Trace Plots", path=getwd(),
              horizontal = FALSE, width = 8.5, height = 8.5)

plot_boxkm(object = synt1,
            main = paste("Cross-validated probability curves for model #1", sep=""),
            xlab = "Time", ylab = "Probability",
            col=2, lty=1, lwd=0.5, cex=0.5,
            device = NULL, file = "Survival Plots", path=getwd(),
            horizontal = TRUE, width = 11.5, height = 8.5)

## Not run:
#####
# Examples of parallel backend parametrization
#####
if (require("parallel")) {
  print("'parallel' is attached correctly \n")
} else {
  stop("'parallel' must be attached first \n")
}
#####
# Example #1 - Quad core PC
# Running WINDOWS with SOCKET communication
#####
cpus <- parallel::detectCores(logical = TRUE)
conf <- list("spec" = rep("localhost", cpus),
            "type" = "SOCKET",
            "homo" = TRUE,
            "verbose" = TRUE,
            "outfile" = "")
#####
# Example #2 - Master node + 3 Worker nodes cluster
# Running LINUX with SOCKET communication
# All nodes equipped with identical setups of
# multicores (8 core CPUs per machine for a total of 32)
#####
masterhost <- Sys.getenv("HOSTNAME")
slavehosts <- c("compute-0-0", "compute-0-1", "compute-0-2")
nodes <- length(slavehosts) + 1
cpus <- 8
conf <- list("spec" = c(rep(masterhost, cpus),
                        rep(slavehosts, cpus)),
            "type" = "SOCKET",
            "homo" = TRUE,
            "verbose" = TRUE,

```

```

"outfile" = "")
#####
# Example #3 - Multinode of multicore per node cluster
# Running LINUX with SLURM scheduler and MPI communication
# Below, variable 'cpus' is the total number
# of requested core CPUs, which is specified from
# within a SLURM script.
#####
if (require("Rmpi")) {
  print("'Rmpi' is attached correctly \n")
} else {
  stop("'Rmpi' must be attached first \n")
}
cpus <- as.numeric(Sys.getenv("SLURM_NTASKS"))
conf <- list("spec" = cpus,
            "type" = "MPI",
            "homo" = TRUE,
            "verbose" = TRUE,
            "outfile" = "")
#####
# Simulated dataset #1 (n=250, p=3)
# Peeling criterion = LRT
# Cross-Validation criterion = LRT
# With Combined Cross-Validation (RCCV)
# With Replications (B = 30)
# With variable screening (pre-selection) (PPL)
# With computation of log-rank permutation p-values
# With parallelization
#####
synt1 <- sbh(X = Synthetic.1[ , -c(1,2), drop=FALSE],
            y = Synthetic.1[ ,1, drop=TRUE],
            delta = Synthetic.1[ ,2, drop=TRUE],
            B = 30,
            K = 5,
            A = 1000,
            vs = TRUE,
            vstype = "ppl",
            vsarg = "alpha=1,
                    nalpha=1,
                    nlambd=100,
                    vscons=0.5",
            cv = TRUE,
            cvtype = "combined",
            cvarg = "alpha=0.01,
                    beta=0.05,
                    minn=5,
                    L=NULL,
                    peelcriterion=\"lrt\",
                    cvcriterion=\"lrt\"",
            pv = TRUE,
            decimals = 2,
            onese = FALSE,
            probval = 0.5,
            timeval = NULL,
            parallel.vs = FALSE,
            parallel.rep = TRUE,
            parallel.pv = TRUE,

```

```

        conf = conf,
        verbose = TRUE,
        seed = 123)

## End(Not run)

```

summary.sbh

*Summary Function***Description**

S3-method summary function to summarize the main parameters used to generate the sbh object.

Usage

```

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

```

Arguments

object	Object of class sbh as generated by the main function sbh .
...	Further generic arguments passed to the summary function.

Value

Summarizes the main parameters used to generate its argument.

Acknowledgments

This work made use of the High Performance Computing Resource in the Core Facility for Advanced Research Computing at Case Western Reserve University. This project was partially funded by the National Institutes of Health NIH - National Cancer Institute (R01-CA160593) to J-E. Dazard and J.S. Rao.

Note

End-user summary function.

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References

- Dazard J-E. and Rao J.S. (2017). "Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods." (in prep).
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- Dazard J-E., Choe M., LeBlanc M. and Rao J.S. (2014). "Cross-Validation of Survival Bump Hunting by Recursive Peeling Methods." In JSM Proceedings, Survival Methods for Risk Estimation/Prediction Section. Boston, MA, USA. American Statistical Association IMS - JSM, p. 3366-3380.
- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

Synthetic.1

Synthetic Dataset #1: $p < n$ case

Description

Dataset from simulated regression survival model #1 as described in Dazard et al. (2015). Here, the regression function uses all of the predictors, which are also part of the design matrix. Survival time was generated from an exponential model with rate parameter λ (and mean $\frac{1}{\lambda}$) according to a Cox-PH model with hazard $\exp(\eta)$, where $\eta(\cdot)$ is the regression function. Censoring indicator were generated from a uniform distribution on $[0, 3]$. In this synthetic example, all covariates are continuous, i.i.d. from a multivariate uniform distribution on $[0, 1]$.

Usage

Synthetic.1

Format

Each dataset consists of a numeric matrix containing $n = 250$ observations (samples) by rows and $p = 3$ variables by columns, not including the censoring indicator and (censored) time-to-event variables. It comes as a compressed Rda data file.

Acknowledgments

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Source

See simulated survival model #1 in Dazard et al., 2015.

References

- Dazard J-E. and Rao J.S. (2017). "Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods." (in prep).
- Yi C. and Huang J. (2016). "Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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Synthetic.1b

Synthetic Dataset #1b: $p < n$ case

Description

Dataset from simulated regression survival model #1b as described in Dazard et al. (2015). Here, the regression function uses all of the predictors, which are also part of the design matrix. In this example, the signal is limited to a box-shaped region R of the predictor space. Survival time was generated from an exponential model with rate parameter λ (and mean $\frac{1}{\lambda}$) according to a Cox-PH model with hazard $\exp(\eta)$, where $\eta(\cdot)$ is the regression function. Censoring indicator were generated from a uniform distribution on $[0, 3]$. In this synthetic example, all covariates are continuous, i.i.d. from a multivariate uniform distribution on $[0, 1]$.

Usage

Synthetic.1b

Format

Each dataset consists of a numeric matrix containing $n = 250$ observations (samples) by rows and $p = 3$ variables by columns, not including the censoring indicator and (censored) time-to-event variables. It comes as a compressed Rda data file.

Acknowledgments

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Source

See simulated survival model #1b in Dazard et al., 2015.

References

- Dazard J-E. and Rao J.S. (2017). "*Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods*." (in prep).
- Yi C. and Huang J. (2016). "*Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression*." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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Synthetic.2

Synthetic Dataset #2: $p < n$ case

Description

Dataset from simulated regression survival model #2 as described in Dazard et al. (2015). Here, the regression function uses some informative predictors. The rest represent un-informative noisy covariates, which are not part of the design matrix. Survival time was generated from an exponential model with rate parameter λ (and mean $\frac{1}{\lambda}$) according to a Cox-PH model with hazard $\exp(\eta)$, where $\eta(\cdot)$ is the regression function. Censoring indicator were generated from a uniform distribution on $[0, 3]$. In this synthetic example, all covariates are continuous, i.i.d. from a multivariate uniform distribution on $[0, 1]$.

Usage

Synthetic.2

Format

Each dataset consists of a numeric matrix containing $n = 250$ observations (samples) by rows and $p = 3$ variables by columns, not including the censoring indicator and (censored) time-to-event variables. It comes as a compressed Rda data file.

Acknowledgments

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Source

See simulated survival model #2 in Dazard et al., 2015.

References

- Dazard J-E. and Rao J.S. (2017). "Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods." (in prep).
- Yi C. and Huang J. (2016). "Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.

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Synthetic.3

Synthetic Dataset #3: $p < n$ case

Description

Dataset from simulated regression survival model #3 as described in Dazard et al. (2015). Here, the regression function does not include any of the predictors. This means that none of the covariates is informative (noisy), and are not part of the design matrix. Survival time was generated from an exponential model with rate parameter λ (and mean $\frac{1}{\lambda}$) according to a Cox-PH model with hazard $\exp(\eta)$, where $\eta(\cdot)$ is the regression function. Censoring indicator were generated from a uniform distribution on $[0, 3]$. In this synthetic example, all covariates are continuous, i.i.d. from a multivariate uniform distribution on $[0, 1]$.

Usage

Synthetic.3

Format

Each dataset consists of a numeric matrix containing $n = 250$ observations (samples) by rows and $p = 3$ variables by columns, not including the censoring indicator and (censored) time-to-event variables. It comes as a compressed Rda data file.

Acknowledgments

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Source

See simulated survival model #3 in Dazard et al., 2015.

References

- Dazard J-E. and Rao J.S. (2017). "Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods." (in prep).
- Yi C. and Huang J. (2016). "Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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- Dazard J-E. and J.S. Rao (2010). "Local Sparse Bump Hunting." J. Comp Graph. Statistics, 19(4):900-92.

Synthetic.4

Synthetic Dataset #4: $p > n$ case

Description

Dataset from simulated regression survival model #4 as described in Dazard et al. (2015). Here, the regression function uses 1/10 of informative predictors in a $p > n$ situation with $p = 1000$ and $n = 100$. The rest represents non-informative noisy covariates, which are not part of the design matrix. Survival time was generated from an exponential model with rate parameter λ (and mean $\frac{1}{\lambda}$) according to a Cox-PH model with hazard $\exp(\eta)$, where $\eta(\cdot)$ is the regression function. Censoring indicator were generated from a uniform distribution on $[0, 2]$. In this synthetic example, all covariates are continuous, i.i.d. from a multivariate standard normal distribution.

Usage

Synthetic.4

Format

Each dataset consists of a numeric matrix containing $n = 100$ observations (samples) by rows and $p = 1000$ variables by columns, not including the censoring indicator and (censored) time-to-event variables. It comes as a compressed Rda data file.

Acknowledgments

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Source

See simulated survival model #4 in Dazard et al., 2015.

References

- Dazard J-E. and Rao J.S. (2017). "*Variable Selection Strategies for High-Dimensional Survival Bump Hunting using Recursive Peeling Methods*." (in prep).
- Yi C. and Huang J. (2016). "*Semismooth Newton Coordinate Descent Algorithm for Elastic-Net Penalized Huber Loss Regression and Quantile Regression*." J. Comp Graph. Statistics, DOI: 10.1080/10618600.2016.1256816.
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