Package 'penAFT'

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tive to the Cox proportional hazards model. This package provides a suite of functions for fit-

Description The semiparametric accelerated failure time (AFT) model is an attractive alterna-

Title Fit the Regularized Gehan Estimator with Elastic Net and Sparse

Type Package

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Group Lasso Penalties

ting one popular estimator of the semiparametric AFT model, the regularized Gehan estimator. Specifically, we provide functions for cross-validation, prediction, coefficient extraction, and visualizing both trace plots and cross-validation curves. For further details, please see Suder, P. M. and Molstad, A. J., (2022+) Scalable algorithms for semiparametric accelerated failure time models in high dimensions, to appear in Statistics in Medicine <doi:10.1002 sim.9264="">.</doi:10.1002>
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penAFT-package

Fit and tune the a semiparameteric accelerated failure time model with weight elastic net or weighted sparse group-lasso penalties.

Description

This package contains numerous functions related to the penalized Gehan estimator. In particular, the main functions are for solution path computation, cross-validation, prediction, and coefficient extraction.

Details

The primary functions are penAFT and penAFT.cv, the latter of which performs cross-validation. In general, both functions fit the penalized Gehan estimator. Given $(\log(y_1), x_1, \delta_1), \ldots, (\log(y_n), x_n, \delta_n)$ where y_i is the minimum of the survival time and censoring time, x_i is a p-dimensional predictor, and δ_i is the indicator of censoring, penAFT fits the solution path for the argument minimizing

$$\frac{1}{n^2} \sum_{i=1}^n \sum_{j=1}^n \delta_i \{ \log(y_i) - \log(y_j) - (x_i - x_j)' \beta \}^- + \lambda g(\beta)$$

where $\{a\}^- := \max(-a, 0)$, $\lambda > 0$, and g is either the weighted elastic net penalty or weighted sparse group lasso penalty. The weighted elastic net penalty is defined as

$$\alpha \|w \circ \beta\|_1 + \frac{(1-\alpha)}{2} \|\beta\|_2^2$$

where w is a set of non-negative weights (which can be specified in the weight.set argument). The weighted sparse group-lasso penalty we consider is

$$\alpha \| w \circ \beta \|_1 + (1 - \alpha) \sum_{l=1}^{G} v_l \| \beta_{\mathcal{G}_l} \|_2$$

where again, w is a set of non-negative weights and v_l are weights applied to each of the G (user-specified) groups.

For a comprehensive description of the algorithm, and more details about rank-based estimation in general, please refer to the referenced manuscript.

Author(s)

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genSurvData	Generate a survival dataset from the log-logistic accelerated failure time model.

Description

This is a function for generating synthetic datasets from the log-logistic accelerated failure time model. The purpose of this function is to provide structured data for the examples of the other functions' usage.

Usage

```
genSurvData(n, p, s, mag, cens.quant = 0.6)
```

Arguments

n	The numer of subjects to be included in the dataset.
p	Dimension of the predictor. Note that the function computes the square-root of a $p \times p$ covariance matrix, so setting p large may be time-consuming.
S	The number of nonzero regression coefficients in β .
mag	The magnitude of the s nonzero regression coefficients. Signs of coefficients are assigned at random.
cens.quant	The quantile of true survival times used to set the mean of the exponential distribution from which censoring times are drawn. Default is 0.6.

Details

This function generates predictors to follow a p-dimensional multivariate normal distribution whose covariance has an AR(1) structure with lag 0.7. Then, log survival times are generated as

$$\log(T) = X\beta + \epsilon$$

where ϵ has independent components drawn from a logistic distribution with location parmeter zero and scale parameter two. Then censoring times are drawn from an exponential distribution with mean equal to the quantile cens. quant of T.

Value

beta	The true data generating regression coefficient vector.
logY	The observed failure times or censoring times on the log scale.
status	Indicator of censoring; a value of 1 indicates the corresponding component of logY is an observed log failure time and a value of 0 indicates a log censoring time.
Xn	The $n \times p$ matrix of predictors.

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Examples

penAFT

Fit the solution path for the regularized semiparametric accelerated failure time model with weighted elastic net or weighted sparse group lasso penalties.

Description

A function to fit the solution path for the regularized semiparametric accelerated failure time model estimator.

Usage

```
penAFT(X, logY, delta, nlambda = 50,
  lambda.ratio.min = 0.1, lambda = NULL,
  penalty = NULL, alpha = 1, weight.set = NULL,
  groups = NULL, tol.abs = 1e-8, tol.rel = 2.5e-4,
  gamma = 0, standardize = TRUE,
  admm.max.iter = 1e4, quiet=TRUE)
```

Arguments

X An $n \times p$ matrix of predictors. Observations should be organized by row.

logY An *n*-dimensional vector of log-survival or log-censoring times.

delta An n-dimensional binary vector indicating whether the jth component of logY

is an observed log-survival time ($\delta_j=1$) or a log-censoring time ($\delta_j=0$) for

 $j=1,\ldots,n$.

nlambda The number of candidate tuning parameters to consider.

lambda.ratio.min

The ratio of maximum to minimum candidate tuning parameter value. As a default, we suggest 0.1, but standard model selection procedures should be applied

to select λ .

lambda An optional (not recommended) prespecified vector of candidate tuning param-

eters. Should be in descending order.

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Either "EN" or "SG" for elastic net or sparse group lasso penalties. penalty alpha The tuning parameter α . See documentation. weight.set A list of weights. For both penalties, w is an n-dimensional vector of nonnegative weights. For "SG" penalty, can also include v – a non-negative vector the length of the number of groups. See documentation for usage example. When using penalty "SG", a p-dimensional vector of integers corresponding the groups to group assignment of each predictor (i.e., column of X). Absolute convergence tolerance. tol.abs tol.rel Relative convergence tolerance. gamma A non-negative optimization parameter which can improve convergence speed in certain settings. It is highly recommended to set equal to zero. standardize Should predictors be standardized (i.e., column-wise average zero and scaled to

have unit variance) for model fitting?

r Maximum number of ADMM iterations.

admm.max.iter Maximum number of ADMM iterations.

quiet TRUE or FALSE variable indicating whether progress should be printed.

Details

Given $(\log y_1, x_1, \delta_1), \dots, (\log y_n, x_n, \delta_n)$ where y_i is the minimum of the survival time and censoring time, x_i is a p-dimensional predictor, and δ_i is the indicator of censoring, penAFT fits the solution path for the argument minimizing

$$\frac{1}{n^2} \sum_{i=1}^n \sum_{j=1}^n \delta_i \{ \log y_i - \log y_j - (x_i - x_j)'\beta \}^- + \lambda g(\beta)$$

where $\{a\}^- := \max(-a,0)$, $\lambda > 0$, and g is either the weighted elastic net penalty (penalty = "EN") or weighted sparse group lasso penalty (penalty = "SG"). The weighted elastic net penalty is defined as

 $\alpha \| w \circ \beta \|_1 + \frac{(1-\alpha)}{2} \| \beta \|_2^2$

where w is a set of non-negative weights (which can be specified in the weight.set argument). The weighted sparse group-lasso penalty we consider is

$$\alpha \|w \circ \beta\|_1 + (1 - \alpha) \sum_{l=1}^G v_l \|\beta_{\mathcal{G}_l}\|_2$$

where again, w is a set of non-negative weights and v_l are weights applied to each of the G groups.

Value

beta A $p \times$ nlambda sparse matrix consisting of the estimates of β for the candidate values of λ . It is recommended to use penAFT. coef to extract coefficients.

lambda The candidate tuning parameter values.

standardize Were predictors standardized to have unit variance for model fitting?

X.mean The mean of the predictors.

X.sd The standard deviation of the predictors.

The tuning parameter α . See documentation.

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```
# -----
# Generate data
# -----
set.seed(1)
genData \leftarrow genSurvData(n = 50, p = 50, s = 10, mag = 2, cens.quant = 0.6)
X <- genData$X
logY <- genData$logY</pre>
delta <- genData$status</pre>
# -----
# Fit elastic net penalized estimator
# -----
fit.en \leftarrow penAFT(X = X, logY = logY, delta = delta,
               nlambda = 50, lambda.ratio.min = 0.01,
               penalty = "EN",
               alpha = 1)
coef.en.10 <- penAFT.coef(fit.en, lambda = fit.en$lambda[10])</pre>
# -----
# Fit weighted elastic net penalized estimator
# ------
weight.set <- list("w" = c(0, 0, rep(1, 48)))
fit.weighted.en <- penAFT(X = X, logY = logY, delta = delta,
               nlambda = 50, weight.set = weight.set,
               penalty = "EN",
               alpha = 1)
coef.wighted.en.10 <- penAFT.coef(fit.weighted.en, lambda = fit.weighted.en$lambda[10])</pre>
# Fit ridge penalized estimator with user-specified lambda
# -----
fit.ridge <- penAFT(X = X, logY = logY, delta = delta,</pre>
               lambda = 10^seq(-4, 4, length=50),
               penalty = "EN",
               alpha = 0)
# -----
# Fit sparse group penalized estimator
# -----
groups \leftarrow rep(1:5, each = 10)
fit.sg <- penAFT(X = X, logY = logY, delta = delta,
               nlambda = 50, lambda.ratio.min = 0.01,
               penalty = "SG", groups = groups,
               alpha = 0.5)
```

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penAFT.coef

Extract regression coefficients from fitted model object

Description

A function to extract coefficients along the solution path for the regularized semiparametric acceleratred failure time model estimator.

Usage

```
penAFT.coef(fit, lambda = NULL)
```

Arguments

fit A fitted model from penAFT or penAFT.cv.

lambda The tuning parameter value at which to extract coefficients. If NULL and fit is

a penAFT.cv object, will use the tuning parameter value with minimum cross-

validation linear predictor score.

Details

The regression coefficients stored in the fitted model objects coming from penAFT or penAFT.cv will (i) be on the scale of standardized predictors if standardization was used (which is the default) and (ii) are stored as a specific sparse matrix so that coefficient extraction is cumbersome. This function returns the regression coefficient estimates on the original scale of the predictors for a particular tuning parmaeter value. It is important to note that this method does not return an estimate of the intercept: the intercept is absored into the error term as the Gehan loss function is invariant to location change.

Value

beta The coefficient estimates

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```
# -----
# Generate data
# -----
set.seed(1)
genData \leftarrow genSurvData(n = 100, p = 50, s = 10, mag = 1, cens.quant = 0.6)
X <- genData$X
logY <- genData$logY</pre>
delta <- genData$status</pre>
# -----
# Fit elastic net penalized estimator without CV
# -----
fit <- penAFT(X = X, logY = logY, delta = delta,</pre>
                nlambda = 50.
                penalty = "EN",
                alpha = 1)
coef.10 <- penAFT.coef(fit, lambda = fit$lambda[10])</pre>
coef.20 <- penAFT.coef(fit, lambda = fit$lambda[20])</pre>
# Cannot obtain fit at lambda not in fit$lambda
## Not run: coef.error <- penAFT.coef(fit, lambda = 10) # throws error
 # ------
 # Fit elastic net penalized estimator with CV
 # -----
 fit.cv <- penAFT.cv(X = X, logY = logY, delta = delta,</pre>
                nlambda = 50,
                penalty = "EN",
                alpha = 1, nfolds = 5)
 ## --- coefficients at lambda minimizing cross-validation error
 coef.cv <- penAFT.coef(fit.cv)</pre>
 ## ---- coefficients at 10th considered lambda
 coef.cv10 <- penAFT.coef(fit.cv, lambda = fit.cv$full.fit$lambda[10])</pre>
 # -----
 # Repeat with sparse group lasso without CV
 # -----
 groups \leftarrow rep(1:10, each = 5)
 fit.sg <- penAFT(X = X, logY = logY, delta = delta,
                nlambda = 50, groups = groups,
                penalty = "SG",
                alpha = 0.5)
 coef.sg.10 <- penAFT.coef(fit.sg, lambda = fit.sg$lambda[10])</pre>
 coef.sg.20 <- penAFT.coef(fit.sg, lambda = fit.sg$lambda[20])</pre>
```

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penAFT.cv

Cross-validation function for fitting a regularized semiparametric accelerated failure time model

Description

A function to perform cross-validation and compute the solution path for the regularized semiparametric accelerated failure time model estimator.

Usage

```
penAFT.cv(X, logY, delta, nlambda = 50,
  lambda.ratio.min = 0.1, lambda = NULL,
  penalty = NULL, alpha = 1,weight.set = NULL,
  groups = NULL, tol.abs = 1e-8, tol.rel = 2.5e-4,
  standardize = TRUE, nfolds = 5, cv.index = NULL,
  admm.max.iter = 1e4,quiet = TRUE)
```

Arguments

X An $n \times p$ matrix of predictors. Observations should be organized by row.

logY An *n*-dimensional vector of log-survival or log-censoring times.

delta An n-dimensional binary vector indicating whether the jth component of logY

is an observed log-survival time ($\delta_j=1$) or a log-censoring time ($\delta_j=0$) for

 $j=1,\ldots,n.$

nlambda The number of candidate tuning parameters to consider.

lambda.ratio.min

The ratio of maximum to minimum candidate tuning parameter value. As a default, we suggest 0.1, but standard model selection procedures should be applied

to select λ .

lambda An optional (not recommended) prespecified vector of candidate tuning param-

eters. Should be in descending order.

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penalty Either "EN" or "SG" for elastic net or sparse group lasso penalties. alpha The tuning parameter α . See documentation. weight.set A list of weights. For both penalties, w is an n-dimensional vector of nonnegative weights. For "SG" penalty, can also include v-a non-negative vector the length of the number of groups. See documentation for usage example. When using penalty "SG", a p-dimensional vector of integers corresponding the groups to group assignment of each predictor (i.e., column of X). tol.abs Absolute convergence tolerance. tol.rel Relative convergence tolerance. standardize Should predictors be standardized (i.e., scaled to have unit variance) for model fitting? nfolds The number of folds to be used for cross-validation. Default is five. Ten is recommended when sample size is especially small. A list of length nfolds of indices to be used for cross-validation. This is to be cv.index used if trying to perform cross-validation for both α and λ . Use with extreme caution: this overwrites nfolds. admm.max.iter Maximum number of ADMM iterations. quiet TRUE or FALSE variable indicating whether progress should be printed.

Details

Given $(\log y_1, x_1, \delta_1), \ldots, (\log y_n, x_n, \delta_n)$ where for subject i $(i = 1, \ldots, n)$, y_i is the minimum of the survival time and censoring time, x_i is a p-dimensional predictor, and δ_i is the indicator of censoring, penAFT.cv performs nfolds cross-validation for selecting the tuning parameter to be used in the argument minimizing

$$\frac{1}{n^2} \sum_{i=1}^n \sum_{j=1}^n \delta_i \{ \log y_i - \log y_j - (x_i - x_j)'\beta \}^- + \lambda g(\beta)$$

where $\{a\}^- := \max(-a,0)$, $\lambda > 0$, and g is either the weighted elastic net penalty (penalty = "EN") or weighted sparse group lasso penalty (penalty = "SG"). The weighted elastic net penalty is defined as

$$\alpha\|w\circ\beta\|_1+\frac{(1-\alpha)}{2}\|\beta\|_2^2$$

where w is a set of non-negative weights (which can be specified in the weight.set argument). The weighted sparse group-lasso penalty we consider is

$$\alpha \|w \circ \beta\|_1 + (1 - \alpha) \sum_{l=1}^G v_l \|\beta_{\mathcal{G}_l}\|_2$$

where again, w is a set of non-negative weights and v_l are weights applied to each of the G groups.

Next, we define the cross-validation errors. Let V_1, \ldots, V_K be a random nfolds = K element partition of [n] (the subjects) with the cardinality of each V_k (the "kth fold"") approximately equal for $k = 1, \ldots, K$. Let $\hat{\beta}_{\lambda(-V_k)}$ be the solution with tuning parameter λ using only data indexed by

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 $[n] \setminus \{\mathcal{V}_k\}$ (i.e., outside the kth fold). Then, definining $e_i(\beta) := \log y_i - \beta' x_i$ for $i = 1, \dots, n$, we call

$$\sum_{k=1}^{K} \left[\frac{1}{|\mathcal{V}_k|^2} \sum_{i \in \mathcal{V}_k} \sum_{j \in \mathcal{V}_k} \delta_i \{ e_i(\hat{\beta}_{\lambda(-\mathcal{V}_k)}) - e_j(\hat{\beta}_{\lambda(-\mathcal{V}_k)}) \}^- \right],$$

the cross-validated Gehan loss at λ in the kth fold, and refer to the sum over all nfolds = K folds as the cross-validated Gehan loss. Similarly, letting letting

$$\tilde{e}_i(\hat{\beta}_{\lambda}) = \sum_{k=1}^{K} (\log y_i - x_i' \hat{\beta}_{\lambda(-\mathcal{V}_k)}) \mathbf{1}(i \in \mathcal{V}_k)$$

for each $i \in [n]$, we call

$$\left[\sum_{i=1}^{n}\sum_{j=1}^{n}\delta_{i}\{\tilde{e}_{i}(\hat{\beta}_{\lambda})-\tilde{e}_{j}(\hat{\beta}_{\lambda})\}^{-}\right]$$

the cross-validated linear predictor score at λ .

Value

full. fit A model fit with the same output as a model fit using penAFT. See documentation for penAFT for more.

cv.err.linPred Anlambda-dimensional vector of cross-validated linear predictor scores.

cv.err.obj A nfolds ×nlambda matrix of cross-valdiation Gehan losses.

cv.index A list of length nfolds. Each element contains the indices for subjects belonging to that particular fold.

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```
# ---- predict at 8th tuning parameter from full fit
Xnew <- matrix(rnorm(10*p), nrow=10)</pre>
predict.en <- penAFT.predict(fit.en, Xnew = Xnew, lambda = fit.en$full.fit$lambda[8])</pre>
 # -----
 # Fit sparse group penalized estimator
 groups \leftarrow rep(1:5, each = 10)
 fit.sg <- penAFT.cv(X = X, logY = logY, delta = delta,
                   nlambda = 50, lambda.ratio.min = 0.01,
                   penalty = "SG", groups = groups, nfolds = 5,
                   alpha = 0.5)
 # ------
 # Pass fold indices
 groups \leftarrow rep(1:5, each = 10)
 cv.index <- list()</pre>
 for(k in 1:5){
   cv.index[[k]] <- which(rep(1:5, length=50) == k)</pre>
 fit.sg.cvIndex <- penAFT.cv(X = X, logY = logY, delta = delta,
                   nlambda = 50, lambda.ratio.min = 0.01,
                   penalty = "SG", groups = groups,
                   cv.index = cv.index,
                   alpha = 0.5)
 # --- compare cv indices
 ## Not run: fit.sg.cvIndex$cv.index == cv.index
```

penAFT.plot

Plot cross-validation curves

Description

A function for plotting the cross-validation curves for the regularized semiparametric accelerated failure time model estimator.

Usage

```
penAFT.plot(fit)
```

Arguments

fit

A fitted model from penAFT.cv.

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Details

This function returns a plot with the cross-validation curves for the regularized Gehan estimator. The vertical blue line indicates the tuning parameter which minimized cross-validated linear predictor scores and the vertical black line indicates the tuning parameter which minimized the cross-validated Gehan loss according to the one-standard-error rule. The vertical axis (and blue line) denotes the cross-validated linear predictor scores whereas the right vertical axis (and black line) denotes cross-validated Gehan loss and standard errors). To make matters simple, we do not allow for customization of the plot: please refer to the source code if extensive customization is desired.

Value

No return value; prints a plot of cross-validation curves as described in Details.

Examples

penAFT.predict

Obtain linear predictor for new subjects using fitted model from penAFT or penAFT.cv

Description

A function for prediction along the solution path of the regularized semiparametric accelerated failure time model estimator.

Usage

```
penAFT.predict(fit, Xnew, lambda = NULL)
```

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Arguments

fit A fitted model from penAFT or penAFT.cv.

Xnew A matrix of dimension $n_{\text{new}} \times p$. Must be a matrix, even if $n_{\text{new}} = 1$.

lambda The value of λ used to estimate β . If NULL and fit was obtained using nfolds

non-NULL, the function will use the tuning parameter which minimized cross-

validation linear predictor scores.

Details

It is important to note that the output of this function should not be treated as an estimate of the log-survival time. Because the Gehan loss function is location invariant, the intercept is absored into the error. If predictors were standardized for model fitting, this function returns $\tilde{X}_{\text{new}}\hat{\beta}$ where \tilde{X}_{new} is the version of input Xnew which has been centered and standardized according to the design matrix used to fit the penAFT or penAFT. cv object. If predictors were not standardized, this function returns $X_{\text{new}}\hat{\beta}$.

We recommend input Xnew as a matrix, although if a *p*-dimensional vector is input, the function will detect this.

Value

preds The matrix of linear predictors: rows correspond to rows of Xnew.

```
# Generate data
# -----
set.seed(1)
genData \leftarrow genSurvData(n = 50, p = 50, s = 10, mag = 2, cens.quant = 0.6)
X <- genData$X
logY <- genData$logY</pre>
delta <- genData$status
# --- generate data for two new subjects
p \leftarrow dim(X)[2]
Xnew <- rbind(rnorm(p), rnorm(p))</pre>
# Fit elastic net penalized estimator without CV
# -----
fit <- penAFT(X = X, logY = logY, delta = delta,
                nlambda = 10, lambda.ratio.min = 0.1,
                 penalty = "EN",
                 alpha = 1)
# predict at 10th candidate tuning parameter
linPred.10 <- penAFT.predict(fit, Xnew = Xnew, lambda = fit$lambda[10])</pre>
 # -----
```

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```
# Fit elastic net penalized estimator with CV
# -----
fit.cv <- penAFT.cv(X = X, logY = logY, delta = delta,
                  nlambda = 50,
                  penalty = "EN",
                   alpha = 1, nfolds = 5)
# --- return linear predictor at lambda minimizing cross-validation error
linPred.cv <- penAFT.predict(fit.cv, Xnew = Xnew)</pre>
# --- predict at 10th candidate tuning parameter
linPred.cv10 <- penAFT.predict(fit.cv, Xnew = Xnew, lambda = fit.cv$full.fit$lambda[10])</pre>
# Fit penAFT with cross-validation
groups \leftarrow rep(1:5, each = 10)
fit.sg.cv <- penAFT.cv(X = X, logY = logY, delta = delta,
                  nlambda = 50, groups = groups,
                  penalty = "SG",
                  alpha = 0.5, nfolds = 5)
# ---- return linear predictor at lambda minimizing cross-validation error
linPred.sg.cv <- penAFT.predict(fit.sg.cv, Xnew = Xnew)</pre>
# --- predict at 10th candidate tuning parameter
linPred.sg.cv10 <- penAFT.predict(fit.sg.cv, Xnew = Xnew, lambda = fit.sg.cv$full.fit$lambda[10])</pre>
```

penAFT.trace

Print trace plot for the regularized Gehan estimator fit using penAFT or penAFT.cv

Description

Print the trace plot for the regularized Gehan estimator.

Usage

```
penAFT.trace(fit, groupNames=NULL)
```

Arguments

fit A fitted model from penAFT or penAFT.cv.

groupNames A list of groupnames to be printed when fit used penalty "SG".

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Details

The function penAFT.trace returns a trace plot for a fitted model obtained from either penAFT or penAFT.cv. If the model is fit using the sparse group-lasso penalty, you may provide names for the groups (in order of the integer values specifying the groups). This feature may not be desired if there are a large number of groups, however. The vertical blue line indicates the tuning parameter which minimized cross-validated linear predictor scores and the vertical black line indicates the tuning parameter minimizing the cross-validated Gehan loss according to the one-standard error rule.

Value

No return value; prints a trace plot as described in Details.

```
# -----
# Generate data
set.seed(1)
genData \leftarrow genSurvData(n = 50, p = 50, s = 10, mag = 2, cens.quant = 0.6)
X <- genData$X
logY <- genData$logY</pre>
delta <- genData$status
# -----
# Fit elastic net penalized estimator with CV
# ------
fit.cv <- penAFT.cv(X = X, logY = logY, delta = delta,</pre>
               nlambda = 10.
               penalty = "EN"
               alpha = 1, nfolds = 5)
# -- print plot
penAFT.trace(fit.cv)
 # ------
 # Fit sparse group-lasso estimator with CV
 # -----
 groups <- rep(1:5, length=10)</pre>
 fit.sg.cv <- penAFT.cv(X = X, logY = logY, delta = delta,
                nlambda = 50, groups = groups,
                penalty = "SG", tol.rel= 1e-5,
                alpha = 0, nfolds = 5)
 penAFT.trace(fit.sg.cv, groupNames = paste("Group", 1:5, sep="-"))
```

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