PH 716 Applied Survival Analysis

Part VIII: Model/variable/subset selection

Zhiyang Zhou (zhou67@uwm.edu, zhiyanggeezhou.github.io)

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Comparing nested models

- The value of (partial) log-likelihood always increasing with added covariates
 - But a parsimonious model preferred if the incremental (partial) log-likelihood is insignificant, because
 - * Reduced overfitting
 - * Increased computational efficiency
 - * Enhanced stability (lower risk of multicollinearity)
 - * Ease of application
 - * Improved interpretability
- ullet Comparing nested models \Leftrightarrow testing the significance of the increment attributable to added covariates

Ex. 8.1: A toy example

- Dataset asaur::pharmacoSmoking
 - ageGroup4: Age group with levels 21-34, 35-49, 50-64, or 65+
 - employment: ft (full-time), pt (part-time), or other
- Candidate models
 - Model A: ageGroup4
 - Model B: employment
 - Model C: ageGroup4+employment
- Models A and B both nested to Model C

```
options(digits=4)
library(survival)
data.ex81 = asaur::pharmacoSmoking
fit.ex81.a = coxph(Surv(ttr, relapse)~ageGroup4, data=data.ex81)
fit.ex81.b = coxph(Surv(ttr, relapse)~employment, data=data.ex81)
fit.ex81.c = coxph(Surv(ttr, relapse)~ageGroup4+employment, data=data.ex81)
logLik(fit.ex81.a)
logLik(fit.ex81.b)
logLik(fit.ex81.c)
anova(fit.ex81.a, fit.ex81.c)
anova(fit.ex81.b, fit.ex81.c)
```

Comparing non-nested models

• Akaike Information Criterion (AIC)

$$AIC = -2p\ell(\hat{\beta}_1, \dots, \hat{\beta}_p) + 2p$$

- $-p\ell(\hat{\beta}_1,\ldots,\hat{\beta}_k)$: (partial) log-likelihood evaluated at estimates $\hat{\beta}_1,\ldots,\hat{\beta}_k$
- The smaller the better
- Bayesian Information Criterion (BIC)

$$BIC = -2p\ell(\hat{\beta}_1, \dots, \hat{\beta}_p) + (\ln n)p$$

- $-p\ell(\hat{\beta}_1,\ldots,\hat{\beta}_k)$: (partial) log-likelihood evaluated at estimates $\hat{\beta}_1,\ldots,\hat{\beta}_k$
- -n: sample size
- The smaller the better

Revisit Ex. 8.1

- Dataset asaur::pharmacoSmoking
 - ageGroup4: Age group with levels 21-34, 35-49, 50-64, or 65+
 - employment: ft (full-time), pt (part-time), or other
- Candidate models
 - Model A: ageGroup4
 - Model B: employment
 - Model C: ageGroup4+employment
- Model A not nested to B

```
AIC(fit.ex81.a)
AIC(fit.ex81.b)
AIC(fit.ex81.c)

BIC(fit.ex81.a)
BIC(fit.ex81.b)
BIC(fit.ex81.c)
```

Generic procedure of model selection

- Forward selection
 - 1. Start with the minimal model.
 - Add predictors one by one: consider adding each of the remaining predictors and evaluate how
 much it improves the model. The evaluation metric can be the improvement in likelihood, AIC,
 BIC, or any other appropriate metric.
 - 3. Select the best predictor: Add the predictor that improves the model the most (e.g., results in the largest decrease in AIC).
 - 4. Repeat steps 2–3 until no significant improvement.
 - 5. Final model: The process stops when no adding improves the model.
- Backward selection (NOT always doable)
 - 1. Start with the maximal model.
 - 2. Remove predictors one by one: consider removing each predictor in the current model and evaluate how much it improves the model. The evaluation metric can be the improvement in likelihood, AIC, BIC, or any other appropriate metric.
 - 3. Select the best predictor: Delete the predictor that improves the model the most (e.g., results in the largest decrease in AIC).
 - 4. Repeat steps 2–3 until no significant improvement.
 - 5. Final model: The process stops when no deletion improves the model.
- (Bidirectional) stepwise selection
 - 1. Initiate by adding the best variable.
 - 2. After adding a new variable, it checks whether any of the previously included variables have become unnecessary and should be removed.
 - 3. Each forward selection step can be followed by one or more backward elimination steps.
 - 4. This process continues until no variable can be added/removed to improve the model.

Revisit Ex. 8.1

```
options(digits=4)
library(survival)
data.ex81 = asaur::pharmacoSmoking
minimal = coxph(Surv(ttr, relapse)~grp, data=data.ex81)
maximal = coxph(
  Surv(ttr, relapse)~grp+gender+race+employment+yearsSmoking+levelSmoking+ageGroup4,
  data=data.ex81
scope = list(lower = minimal, upper = maximal)
# Selection with AIC
forward.AIC = step(minimal, scope = scope, direction = "forward", k = 2)
backward.AIC = step(maximal, scope = scope, direction = "backward", k = 2)
stepwise.AIC = step(minimal, scope = scope, direction = "both", k = 2)
# Selection with BIC
forward.BIC = step(minimal, scope = scope, direction = "forward", k = log(nrow(data.ex81)))
backward.BIC = step(maximal, scope = scope, direction = "backward", k = log(nrow(data.ex81)))
stepwise.BIC = step(minimal, scope = scope, direction = "both", k = log(nrow(data.ex81)))
```

Penalization/regularization/shirinkage

- model selection \Leftrightarrow variable selection \Leftrightarrow compressing certain coefficients to zeros
- Introducing bias but may significantly decrease variances of the estimates
- Ridge: minimizing a penalized (partial) log-likelihood

$$-p\ell(\beta_1,\ldots,\beta_p) + \lambda \sum_{j=1}^p \beta_j^2$$

with respect to β_1, \ldots, β_p

- $-\lambda \geq 0$: a tuning parameter
- Penalizing large estimates of coefficients (BUT setting NO coefficient to exactly zero)
- Equv. minimizing $-p\ell(\beta_1,\ldots,\beta_p)$ subject to $\sum_{j=1}^p \beta_j^{\frac{j}{2}} \leq \eta(\lambda)$ * $\eta > 0$: a scalar corresponding to λ
- Least absolute shrinkage and selection operator (lasso): minimizing a penalized (partial) log-likelihood

$$-p\ell(\beta_1,\ldots,\beta_p) + \lambda \sum_{j=1}^p |\beta_j|$$

with respect to β_1, \ldots, β_p

- $-\lambda \geq 0$: a tuning parameter controlling the overall penalty strength
- Shrinking some estimates of coefficients to zero
- Equv. minimizing $-p\ell(\beta_1,\ldots,\beta_p)$ subject to $\sum_{j=1}^p |\beta_j| \le \eta(\lambda)$ * $\eta > 0$: a scalar corresponding to λ
- Elastic net: minimizing a penalized (partial) log-likelihood

$$-p\ell(\beta_1,\ldots,\beta_p) + \lambda \left(\frac{1-\alpha}{2}\sum_{j=1}^p \beta_j^2 + \alpha\sum_{j=1}^p |\beta_j|\right)$$

with respect to β_1, \ldots, β_p

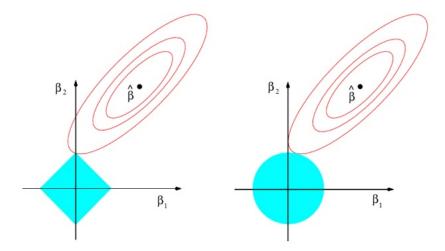


Figure 1: Contours of the error and constraint functions for the lasso (left) and ridge regression (right). The solid blue areas are the constraint regions, $|\beta_1| + |\beta_2| \le \eta$ and $\beta_1^2 + \beta_2^2 \le \eta$, while the red ellipses are the contours of (partial) log-likelihood. (G. James, D. Witten, T. Hastie, & R. Tibshirani, 2021, An Introduction to Statistical Learning, 2nd Ed., pp. 245)

- $-\lambda \geq 0$: a tuning parameter controlling the overall penalty strength
- $-\alpha \in [0,1]$: a tuning parameter controlling the regression type
- A mixture of ridge ($\alpha = 0$) and lasso ($\alpha = 1$)
- R function glmnet::glmnet

Revisit Ex. 8.1

```
options(digits=4)
library(survival)
library(glmnet)
data.ex81 = asaur::pharmacoSmoking
sapply(data.ex81, class)
maximal = coxph(
  Surv(ttr, relapse)~
    grp+gender+race+employment+yearsSmoking+levelSmoking+ageGroup4,
  data=data.ex81,
  x = T
)
zero.ttr.idx = (data.ex81$ttr == 0 & data.ex81$relapse == 1) # subjects with zero event time
# ridge-penalized Cox PH (with the lambda sequence generated automatically)
cox.ridge = glmnet(
  x=maximal$x[!zero.ttr.idx,],
 y=maximal$y[!zero.ttr.idx],
 family = 'cox',
  alpha = 0, # ridge
 nlambda = 100 # number of different lambda values
## paths of estimates against log(lambda)
plot(cox.ridge, xvar="lambda", label=TRUE)
```

```
# lasso-penalized Cox PH (with the lambda sequence generated automatically)
cox.lasso = glmnet(
    x=maximal$x[!zero.ttr.idx,],
    y=maximal$y[!zero.ttr.idx],
    family = 'cox',
    alpha = 1, # lasso
    nlambda = 100 # number of different lambda values
)
## paths of estimates against log(lambda)
plot(cox.lasso, xvar="lambda", label=TRUE)
```

Cross-validation (CV)

- To determine the value of tuning parameter(s)
- K-fold CV
 - 1. Partition the dataset into K subsets (called "folds")
 - 2. Set up a grid of values of λ
 - 3. For each potential value of λ
 - a. Take one fold as a test set and the remaining K-1 folds as a training set.
 - b. Fit the model on the training set.
 - c. Evaluate the model fitting in terms of certain measure on the test set.
 - d. Repeat a–c for all K folds and compute the average measure value, say $\mathrm{CV}(\lambda) = n^{-1} \sum_{k=1}^K \mathrm{measure}_k$.
 - 4. Select the value of λ that minimizes $CV(\lambda)$.
- Leave-one-out $CV \Leftrightarrow n$ -fold CV
- Measure used for CV for Cox PH models
 - (Partial) likelihood deviance: $2\{p\ell_{\text{saturated}} p\ell(\beta_1, \dots, \beta_p)\}$
 - * $p\ell_{\text{saturated}} = -\sum_{m} d_m \ln d_m$ with d_m as the number of events at the mth ordered failure time
 - Concordance index (C-index)

$$Cidx = \frac{\sum_{i_1, i_2} \delta_{i_2} \cdot 1_{\tilde{t}_{i_1} > \tilde{t}_{t_2}} \cdot 1_{\lambda_{i_1} < \lambda_{t_2}}}{\sum_{i_1, i_2} \delta_{i_2} \cdot 1_{\tilde{t}_{i_1} > \tilde{t}_{t_2}}}$$

- * Where
 - · $1_{\tilde{t}_{i_1} > \tilde{t}_{t_2}} = 1$ if $\tilde{t}_{i_1} > \tilde{t}_{t_2}$ and 0 otherwise
 - · $1_{\lambda_{i_1} < \lambda_{t_2}}$ if $\lambda_{i_1} < \lambda_{t_2}$ and 0 otherwise
- * To evaluate the discrimination/concordance of a survival model
 - · Discrimination: the ability to correctly provide a reasonable ranking of survival times based on the predicted individual risks
 - · Concordance: the subject with a higher predicted risk corresponds to a longer lifetime
- * Ranging from .5 to 1, the higher the better
 - Cidx = .5: a model with no predictive ability, equivalent to random guessing
 - · Cidx = 1: a perfect model where the predictions and outcomes are in complete concordance
- R function glmnet::cv.glmnet

Revisit Ex. 8.1

```
options(digits=4)
library(survival)
library(glmnet)
data.ex81 = asaur::pharmacoSmoking
```

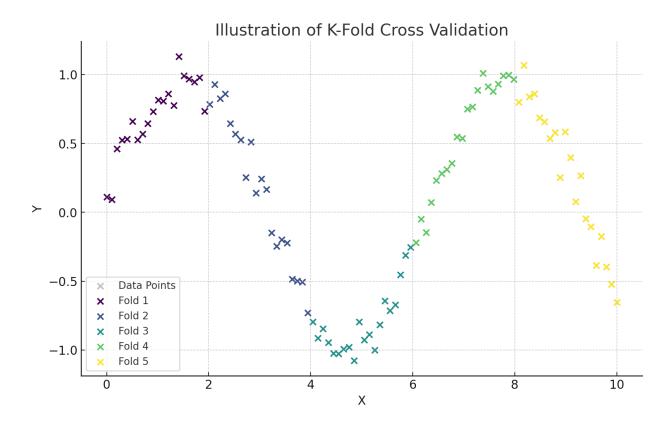


Figure 2: 5-fold CV

```
sapply(data.ex81, class)
maximal = coxph(
  Surv(ttr, relapse)~
    grp+gender+race+employment+yearsSmoking+levelSmoking+ageGroup4,
  data=data.ex81,
  x = T
)
zero.ttr.idx = (data.ex81$ttr == 0 & data.ex81$relapse == 1) # subjects with zero event time
# 5-fold CV
cv.fit = cv.glmnet(
  x=maximal$x[!zero.ttr.idx,],
  y=maximal$y[!zero.ttr.idx],
  family = 'cox',
  alpha = 1, # by default alpha = 1
  type.measure = c("deviance"), # "deviance" or "C"
  nfolds = 5
cv.fit$lambda.min
cv.fit$lambda # values of lambda used in the fits
cv.fit$cvm # mean cross-validated errors
plot(cv.fit)
# fit with the lambda given by CV
cox.lasso = glmnet(
```

```
x=maximal$x[!zero.ttr.idx],
y=maximal$y[!zero.ttr.idx],
family = 'cox',
alpha = 1, # by default alpha = 1
lambda = cv.fit$lambda.min
)
cox.lasso$beta

# fit with the lambda given by CV
cox.lasso = glmnet(
x=maximal$x[!zero.ttr.idx,],
y=maximal$y[!zero.ttr.idx],
family = 'cox',
alpha = 1, # lasso
)
plot(cox.lasso, xvar="lambda", label=TRUE)
```