Supplement-zc2556

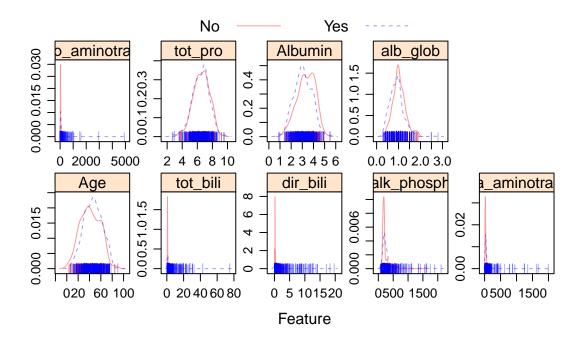
Zhe Chen

2021/03/30

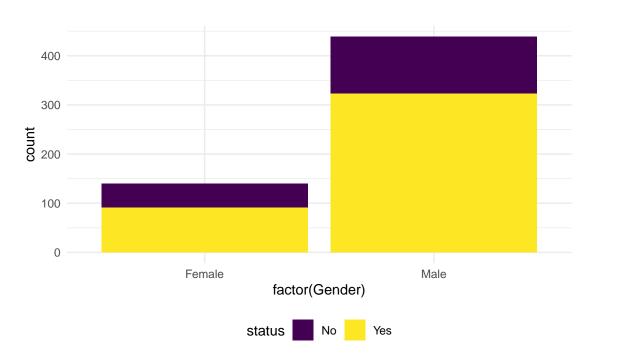
Data Preparation

```
liver.df = read.csv("./indian_liver_patient.csv") %>%
  mutate(
    status = ifelse(Dataset == 1, "Yes", "No"),
    status = as.factor(status),
  ) %>%
  rename(
    tot_bili = Total_Bilirubin,
    dir_bili = Direct_Bilirubin,
    alk_phosph = Alkaline_Phosphotase,
    ala_aminotrans = Alamine_Aminotransferase,
    asp_aminotrans = Aspartate_Aminotransferase,
    tot_pro = Total_Protiens,
    alb_glob = Albumin_and_Globulin_Ratio
  )%>%
  na.omit() %>%
  dplyr::select(-Dataset)
set.seed(621)
liver.t = createDataPartition(y = liver.df$status,
                                  p = 0.8,
                                  list = FALSE)
liver.train = liver.df[liver.t, ]
liver.test = liver.df[-liver.t, ]
```

Exploratory Plot



```
#plot for gender
ggplot(liver.df, aes(factor(Gender), fill = status))+
  geom_bar()
```



```
#test for gender differences
fm.trail = liver.df %>% filter(Gender == "Female") %>%
    nrow()
fm.case = liver.df %>% filter(Gender == "Female" & status == "Yes") %>%
    nrow()
```

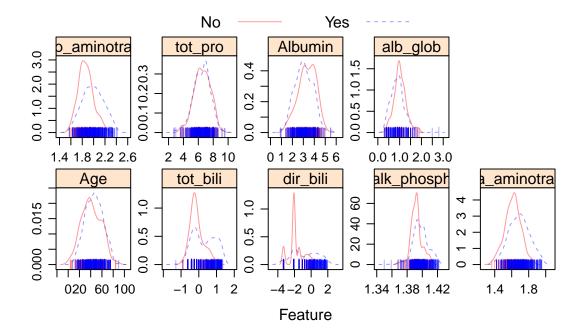
```
m.trail = liver.df %>% filter(Gender == "Male") %>%
  nrow()
m.case = liver.df %>% filter(Gender == "Male" & status == "Yes") %>%
prop.test(x=c(fm.case,m.case), n=c(fm.trail, m.trail),
          conf.level=0.95)
##
##
  2-sample test for equality of proportions with continuity correction
##
## data: c(fm.case, m.case) out of c(fm.trail, m.trail)
## X-squared = 3.4223, df = 1, p-value = 0.06432
## alternative hypothesis: two.sided
## 95 percent confidence interval:
## -0.179600360 0.008074164
## sample estimates:
##
     prop 1
               prop 2
## 0.6500000 0.7357631
Transformation
preprocesspred <- preProcess(liver.df[,3:7], method=c("BoxCox"))</pre>
print(preprocesspred)
## Created from 579 samples and 5 variables
##
## Pre-processing:
    - Box-Cox transformation (5)
     - ignored (0)
##
## Lambda estimates for Box-Cox transformation:
## -0.7, -0.3, -0.7, -0.5, -0.4
```

```
dir_bili
##
        Age
                    Gender
                                      tot_bili
## Min. : 4.0
                 Length:464
                                  Min. :-1.2845
                                                        :-3.3175
                                                  Min.
  1st Qu.:33.0
                 Class :character
                                  1st Qu.:-0.2415
                                                  1st Qu.:-2.0689
## Median :45.0
                 Mode :character
                                  Median : 0.0000
                                                  Median :-1.4501
## Mean
        :44.6
                                   Mean : 0.1868
                                                   Mean :-1.0003
  3rd Qu.:57.0
                                   3rd Qu.: 0.7158
##
                                                   3rd Qu.: 0.2523
          :90.0
                                   Max. : 1.3590
                                                   Max.
                                                          : 1.9702
##
  {\tt Max.}
##
     alk_phosph
                ala_aminotrans asp_aminotrans
                                                  tot_pro
                Min. :1.368
## Min.
         :1.350
                                 Min. :1.575 Min.
                                                      :2.700
## 1st Qu.:1.390
                  1st Qu.:1.583
                                 1st Qu.:1.821
                                                1st Qu.:5.800
## Median :1.395
                 Median :1.662
                                 Median :1.937
                                               Median :6.600
## Mean :1.396
                Mean :1.669 Mean :1.955 Mean :6.479
```

liver.df.boxcox <- predict(preprocesspred, liver.df)
liver.train.boxcox <- predict(preprocesspred, liver.train)
liver.test.boxcox <- predict(preprocesspred, liver.test)</pre>

summary(liver.train.boxcox)

```
3rd Qu.:1.402
                   3rd Qu.:1.744
                                    3rd Qu.:2.083
                                                    3rd Qu.:7.200
##
   Max.
          :1.422
                   Max. :1.955
                                    Max.
                                          :2.417
                                                    Max.
                                                           :9.600
      Albumin
                       alb glob
##
                                     status
           :0.900
                          :0.3000
                                     No :132
##
  Min.
                   Min.
                   1st Qu.:0.7000
##
   1st Qu.:2.500
                                     Yes:332
##
  Median :3.100
                   Median :0.9000
   Mean
         :3.127
                   Mean :0.9417
   3rd Qu.:3.800
                    3rd Qu.:1.1000
##
          :5.500
   Max.
                   Max.
                           :2.8000
featurePlot(x = liver.train.boxcox[, c(1,3:10)],
  y = liver.train.boxcox$status,
  scales = list(x = list(relation = "free"),
                y = list(relation = "free")),
  plot = "density", pch = "|",
  auto.key = list(columns = 2))
```



Model Building

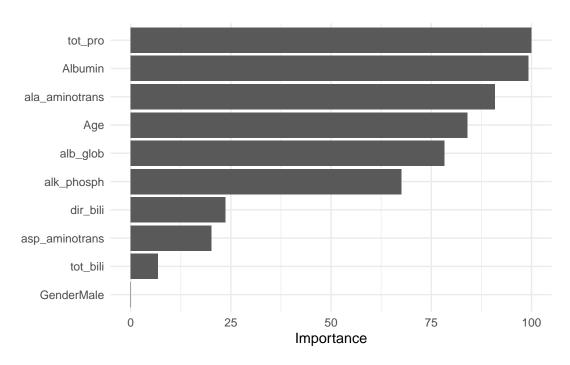
Feature Selection

```
log.fit = glm(status ~ ., data = liver.train.boxcox,family = binomial(link = "logit"))
summary(log.fit)

##
## Call:
## glm(formula = status ~ ., family = binomial(link = "logit"),
## data = liver.train.boxcox)
##
```

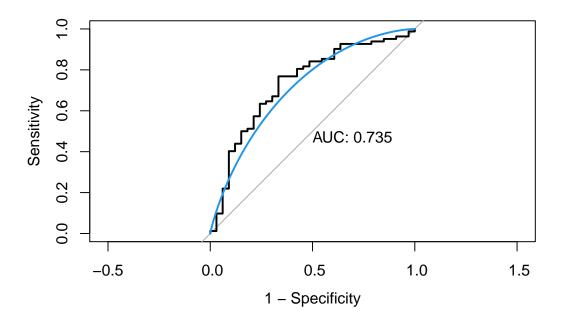
```
## Deviance Residuals:
##
      Min
                10
                    Median
                                  30
                                          Max
## -2.3096 -0.9950
                    0.4138
                              0.8246
                                        1.6669
##
## Coefficients:
                   Estimate Std. Error z value Pr(>|z|)
##
                 -52.178881 19.599619 -2.662 0.00776 **
## (Intercept)
                                         2.514 0.01194 *
## Age
                   0.018305
                              0.007281
## GenderMale
                   -0.023442
                              0.265936 -0.088 0.92976
## tot_bili
                   0.208988
                             0.733432
                                         0.285 0.77569
## dir_bili
                   0.222032
                              0.288827
                                         0.769 0.44205
## alk_phosph
                   28.842930 14.154091
                                         2.038 0.04157 *
## ala_aminotrans
                  4.574380
                             1.686871
                                         2.712 0.00669 **
                  0.752375
                                         0.670 0.50273
## asp_aminotrans
                             1.122608
## tot_pro
                              0.446144
                                         2.974 0.00294 **
                   1.326969
## Albumin
                   -2.596596
                              0.879445 -2.953 0.00315 **
                                         2.348 0.01889 *
## alb_glob
                   3.198564
                              1.362463
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
       Null deviance: 554.14 on 463 degrees of freedom
## Residual deviance: 455.17 on 453 degrees of freedom
## AIC: 477.17
## Number of Fisher Scoring iterations: 5
#logistic regression
set.seed(621)
ctrl <-trainControl(method = "repeatedcv", repeats = 10, summaryFunction = twoClassSummary,classProbs =</pre>
glm.fit <- train(x = liver.train.boxcox[,1:10],</pre>
                y = liver.train.boxcox$status,
                method = "glm",
                metric = "ROC",
                 trControl = ctrl)
glm.fit
## Generalized Linear Model
##
## 464 samples
## 10 predictor
##
    2 classes: 'No', 'Yes'
##
## No pre-processing
## Resampling: Cross-Validated (10 fold, repeated 10 times)
## Summary of sample sizes: 418, 417, 418, 418, 417, 417, ...
## Resampling results:
##
##
    ROC
                Sens
                          Spec
    0.7495863 0.2913736 0.8876827
```

vip(glm.fit)



```
## Generalized Linear Model
##
## 464 samples
     6 predictor
##
     2 classes: 'No', 'Yes'
##
##
## No pre-processing
## Resampling: Cross-Validated (10 fold, repeated 10 times)
## Summary of sample sizes: 417, 417, 418, 418, 418, 417, ...
## Resampling results:
##
##
    ROC
                Sens
                           Spec
    0.7452946 0.2698901 0.8951693
##
```

```
anova(glm.fit2$finalModel, glm.fit$finalModel, test = "LRT")
## Analysis of Deviance Table
##
## Model 1: .outcome ~ tot_pro + Albumin + ala_aminotrans + Age + alb_glob +
##
       alk_phosph
## Model 2: .outcome ~ Age + Gender + tot_bili + dir_bili + alk_phosph +
       ala_aminotrans + asp_aminotrans + tot_pro + Albumin + alb_glob
##
     Resid. Df Resid. Dev Df Deviance Pr(>Chi)
## 1
           457
                   465.95
                   455.17 4
## 2
           453
                               10.774 0.02922 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
#ROC for logistics
test.pred.prob.glm2 <- predict(glm.fit2, newdata = liver.test.boxcox ,type = "prob")[,2]</pre>
roc.glm2 <- roc(liver.test.boxcox$status, test.pred.prob.glm2)</pre>
## Setting levels: control = No, case = Yes
## Setting direction: controls < cases
plot(roc.glm2, legacy.axes = TRUE, print.auc = TRUE)
plot(smooth(roc.glm2), col = 4, add = TRUE)
```



LDA and QDA

```
#LDA
set.seed(621)
model.lda <-train(</pre>
  x = liver.train.boxcox[,c(8,9,6,1,10,5)],
  y = liver.train.boxcox$status,
  method = "lda",
 metric = "ROC",
  trControl = ctrl)
model.lda
## Linear Discriminant Analysis
##
## 464 samples
##
    6 predictor
##
     2 classes: 'No', 'Yes'
##
## No pre-processing
## Resampling: Cross-Validated (10 fold, repeated 10 times)
## Summary of sample sizes: 418, 417, 418, 418, 417, 417, ...
## Resampling results:
##
##
     ROC
                Sens
                            Spec
     0.7449128 0.2435165 0.9116667
##
#QDA
set.seed(621)
model.qda \leftarrow train(x = liver.train.boxcox[,c(8,9,6,1,10,5)],
                  y = liver.train.boxcox$status,
                  method = "qda",
                  metric = "ROC",
                  trControl = ctrl)
model.qda
## Quadratic Discriminant Analysis
##
## 464 samples
    6 predictor
##
     2 classes: 'No', 'Yes'
## No pre-processing
## Resampling: Cross-Validated (10 fold, repeated 10 times)
## Summary of sample sizes: 418, 417, 418, 418, 417, 417, ...
## Resampling results:
##
##
                Sens
                           Spec
     0.7201886 0.724011 0.6490463
##
```

KNN

```
set.seed(621)
model.knn \leftarrow-train(x = liver.train.boxcox[,c(8,9,6,1,10,5)],
                 y = liver.train.boxcox$status,
                 method = "knn",
                 metric = "ROC",
                 trControl = ctrl,
                 tuneGrid = data.frame(k=1:12))
model.knn
## k-Nearest Neighbors
##
## 464 samples
    6 predictor
##
##
    2 classes: 'No', 'Yes'
## No pre-processing
## Resampling: Cross-Validated (10 fold, repeated 10 times)
## Summary of sample sizes: 418, 417, 418, 418, 417, 417, ...
## Resampling results across tuning parameters:
##
##
        ROC
    k
                   Sens
                              Spec
##
     1 0.6135313 0.4474725 0.7795900
##
     2 0.6329099 0.4032967 0.7583779
##
     3 0.6527855 0.4035714 0.8205080
##
     4 0.6525063 0.3795604 0.8046346
##
     5 0.6523965 0.3400000 0.8374332
##
     6 0.6568939 0.3089011 0.8334759
##
     7 0.6656735 0.2319231 0.8578610
##
     8 0.6683748 0.2326923 0.8641800
     9 0.6678607 0.2060440 0.8677986
##
##
    10 0.6629104 0.2097802 0.8687879
    11 0.6558820 0.1644505 0.8732353
##
     12 0.6499859 0.1566484 0.8855793
##
## ROC was used to select the optimal model using the largest value.
## The final value used for the model was k = 8.
```

Naive Bey

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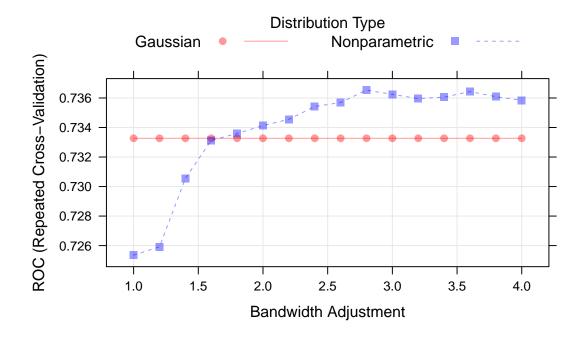
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## observation 43
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## observation 43
```

plot(model.nb)



Box Plot of Resamples

Spec

```
summary(res)
##
## Call:
## summary.resamples(object = res)
## Models: LDA, QDA, KNN, glm, NB
## Number of resamples: 100
##
## ROC
##
            Min.
                   1st Qu.
                               Median
                                           Mean
                                                  3rd Qu.
                                                                Max. NA's
## LDA 0.5735931 0.7038378 0.7452547 0.7449128 0.7948718 0.9208145
  QDA 0.5216450 0.6771611 0.7202797 0.7201886 0.7808858 0.9072398
                                                                        0
## KNN 0.4545455 0.6250000 0.6699967 0.6683748 0.7122583 0.8647186
                                                                        0
## glm 0.5757576 0.7031229 0.7447552 0.7452946 0.7906211 0.8881119
                                                                        0
      0.5571096 0.6965718 0.7331002 0.7365308 0.7814685 0.8927739
                                                                        0
##
## Sens
##
             Min.
                    1st Qu.
                                Median
                                            Mean
                                                   3rd Qu.
## LDA 0.00000000 0.1538462 0.2307692 0.2435165 0.3076923 0.6428571
                                                                         0
```

QDA 0.42857143 0.6153846 0.7142857 0.7240110 0.8461538 1.0000000

KNN 0.00000000 0.1538462 0.2307692 0.2326923 0.3076923 0.6153846

glm 0.00000000 0.2142857 0.2857143 0.2698901 0.3076923 0.5384615

NB 0.07142857 0.2307692 0.3076923 0.3364835 0.4285714 0.6153846

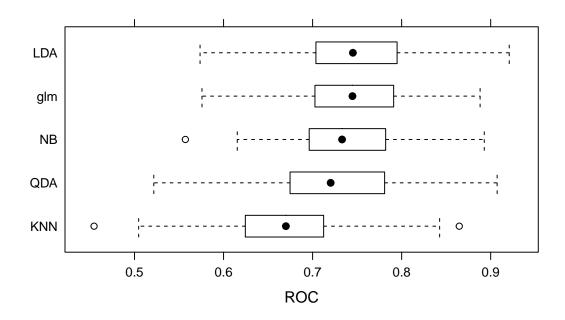
res = resamples(list(LDA = model.lda, QDA = model.qda, KNN = model.knn, glm = glm.fit2, NB = model.nb))

0

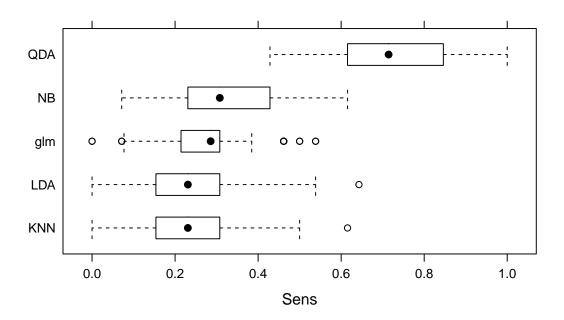
0

0

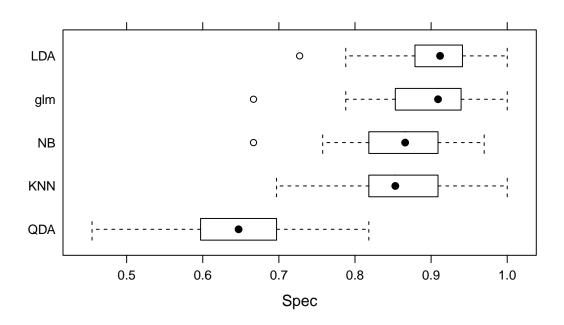
bwplot(res, metric = "ROC")



bwplot(res, metric = "Sens")



```
bwplot(res, metric = "Spec")
```



ROC Plot

```
lda.pred <- predict(model.lda, newdata = liver.test.boxcox, type = "prob")[,2]

## Warning in predict.lda(modelFit, newdata): variable names in 'newdata' do not

## match those in 'object'

knn.pred <- predict(model.knn, newdata = liver.test.boxcox, type = "prob")[,2]

qda.pred <- predict(model.qda, newdata = liver.test.boxcox, type = "prob")[,2]

## Warning in predict.qda(modelFit, newdata): variable names in 'newdata' do not

## match those in 'object'

nb.pred <- predict(model.nb, newdata = liver.test.boxcox, type = "prob")[,2]

roc.lda <- roc(liver.test.boxcox$status, lda.pred)

## Setting levels: control = No, case = Yes

## Setting levels: control = No, case = Yes

## Setting levels: control = No, case = Yes

## Setting direction: controls < cases</pre>
```

```
roc.qda <- roc(liver.test.boxcox$status, qda.pred)

## Setting levels: control = No, case = Yes
## Setting direction: controls < cases

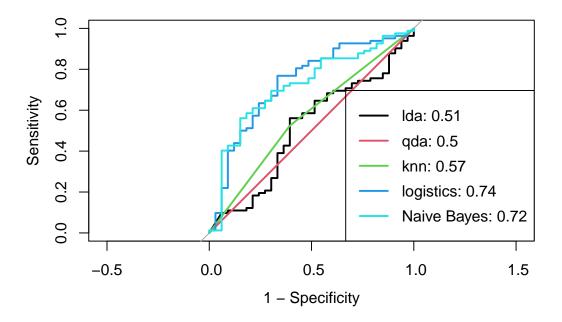
roc.nb <- roc(liver.test.boxcox$status, nb.pred)

## Setting levels: control = No, case = Yes
## Setting direction: controls < cases

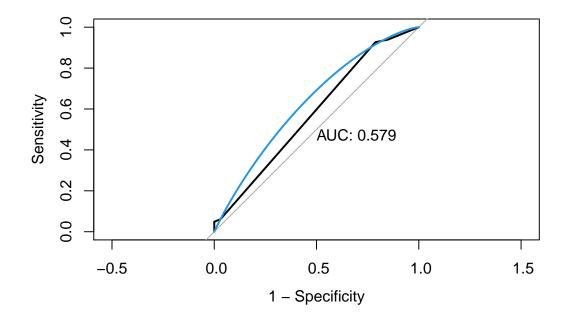
auc <- c(roc.lda$auc[1], roc.qda$auc[1], roc.knn$auc[1], roc.glm2$auc[1], roc.nb$auc[1])

plot(roc.lda, legacy.axes = TRUE)
plot(roc.qda, col = 2, add = TRUE)
plot(roc.knn, col = 3, add = TRUE)
plot(roc.glm2, col = 4, add = TRUE)
plot(roc.nb, col = 5, add = TRUE)

modelNames <- c("lda","qda","knn","logistics", "Naive Bayes")
legend("bottomright", legend = paste0(modelNames, ": ", round(auc,2)),
col = 1:5, lwd = 2)</pre>
```

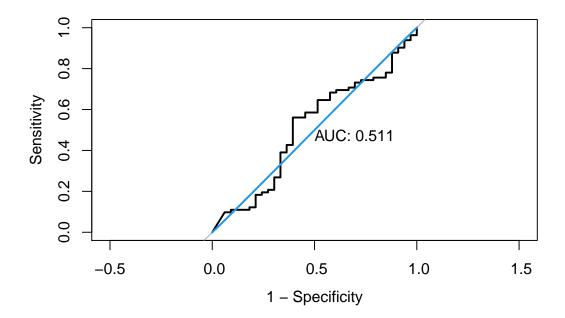


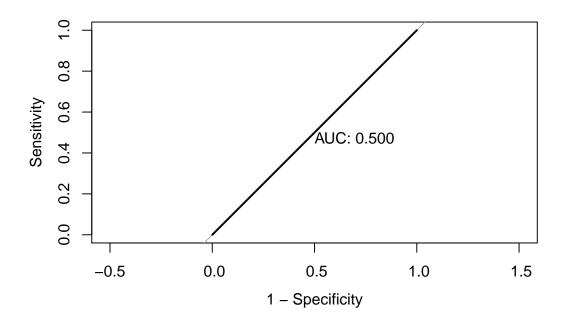
This shows overfitting for KNN and QDA. Thus, new models needed to be build for KNN and QDA. Now, AUC tested for KNN model and QDA model with different cross validation parameters.

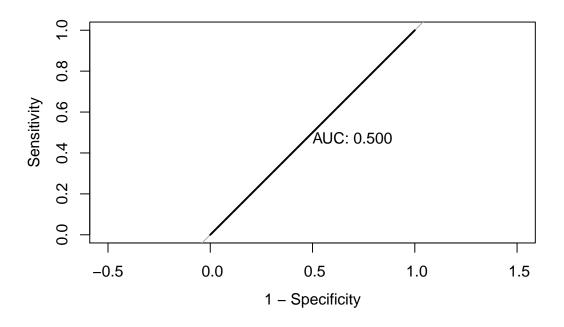


```
## Setting direction: controls < cases
```

```
plot(roc.lda.new, legacy.axes = TRUE, print.auc = TRUE)
plot(smooth(roc.lda.new), col = 4, add = TRUE)
```







Confusion Matrix

```
#confusion matrix for logistics regression with all features
test.pred.prob <- predict(log.fit, newdata = liver.test.boxcox ,type = "response")
test.pred <- rep("No", length(test.pred.prob))
test.pred[test.pred.prob>0.5] <- "Yes"
CM.log = confusionMatrix(data = as.factor(test.pred),
    reference = liver.test.boxcox$status,
    positive = "Yes",
    prevalence = preval)</pre>
CM.log
```

```
## Confusion Matrix and Statistics
##
##
             Reference
##
  Prediction No Yes
##
          No
             13
                   8
          Yes 20
##
                  74
##
##
                  Accuracy : 0.7565
##
                    95% CI : (0.6677, 0.8317)
##
       No Information Rate: 0.713
       P-Value [Acc > NIR] : 0.17730
##
##
##
                     Kappa: 0.3325
##
##
    Mcnemar's Test P-Value : 0.03764
##
##
               Sensitivity: 0.9024
```

```
##
               Specificity: 0.3939
##
            Pos Pred Value: 1.6776
##
            Neg Pred Value: 1.7000
##
                Prevalence: 2.5091
##
            Detection Rate: 0.6435
##
      Detection Prevalence: 0.8174
##
         Balanced Accuracy: 0.6482
##
##
          'Positive' Class : Yes
##
#confusion matrix for logistics regression selected features
test.pred.glm2 <- rep("No", length(test.pred.prob.glm2))</pre>
test.pred.glm2[test.pred.prob.glm2>0.5] <- "Yes"</pre>
CM.log.fea = confusionMatrix(data = as.factor(test.pred.glm2),
  reference = liver.test.boxcox$status,
  positive = "Yes",
 prevalence = preval)
CM.log.fea
## Confusion Matrix and Statistics
##
##
             Reference
## Prediction No Yes
##
          No 11
##
          Yes 22 76
##
##
                  Accuracy : 0.7565
##
                    95% CI : (0.6677, 0.8317)
##
       No Information Rate: 0.713
       P-Value [Acc > NIR] : 0.177296
##
##
##
                     Kappa: 0.3042
##
   Mcnemar's Test P-Value: 0.004586
##
##
               Sensitivity: 0.9268
##
##
               Specificity: 0.3333
##
            Pos Pred Value: 1.7625
##
            Neg Pred Value: 1.5747
##
                Prevalence: 2.5091
##
            Detection Rate: 0.6609
##
      Detection Prevalence: 0.8522
##
         Balanced Accuracy: 0.6301
##
##
          'Positive' Class : Yes
##
#confusion matrix for LDA
test.pred.lda <- rep("No", length(lda.pred))</pre>
test.pred.lda[lda.pred>0.5] <- "Yes"</pre>
CM.lda = confusionMatrix(data = as.factor(test.pred.lda),
 reference = liver.test.boxcox$status,
```

```
positive = "Yes",
  prevalence = preval)
CM.lda
## Confusion Matrix and Statistics
##
            Reference
## Prediction No Yes
         No 2
##
         Yes 31 76
##
##
##
                  Accuracy : 0.6783
##
                    95% CI: (0.5847, 0.7623)
##
      No Information Rate: 0.713
      P-Value [Acc > NIR] : 0.8238
##
##
##
                     Kappa: -0.0162
##
##
  Mcnemar's Test P-Value: 7.961e-05
##
##
              Sensitivity: 0.92683
##
              Specificity: 0.06061
##
            Pos Pred Value: 2.56149
            Neg Pred Value : -0.99271
##
##
                Prevalence: 2.50909
##
            Detection Rate: 0.66087
##
     Detection Prevalence: 0.93043
##
        Balanced Accuracy: 0.49372
##
##
          'Positive' Class : Yes
##
#confusion matrix for QDA
test.pred.qda <- rep("No", length(qda.pred))</pre>
test.pred.qda[qda.pred>0.5] <- "Yes"
CM.qda = confusionMatrix(data = as.factor(test.pred.qda),
 reference = liver.test.boxcox$status,
 positive = "Yes",
 prevalence = preval)
## Warning in confusionMatrix.default(data = as.factor(test.pred.qda), reference =
## liver.test.boxcox$status, : Levels are not in the same order for reference and
## data. Refactoring data to match.
CM.qda
## Confusion Matrix and Statistics
##
##
             Reference
## Prediction No Yes
         No O
         Yes 33 82
##
```

```
##
##
                  Accuracy: 0.713
                    95% CI: (0.6212, 0.7935)
##
##
       No Information Rate: 0.713
##
       P-Value [Acc > NIR] : 0.5468
##
##
                     Kappa: 0
##
##
   Mcnemar's Test P-Value: 2.54e-08
##
##
               Sensitivity: 1.000
##
               Specificity: 0.000
            Pos Pred Value: 2.509
##
##
            Neg Pred Value :
##
                Prevalence: 2.509
##
            Detection Rate: 0.713
##
      Detection Prevalence : 1.000
##
         Balanced Accuracy: 0.500
##
##
          'Positive' Class : Yes
##
#confusion matrix for knn
CM.pred.knn <- rep("No", length(knn.pred))</pre>
CM.pred.knn[knn.pred>0.5] <- "Yes"</pre>
CM.knn = confusionMatrix(data = as.factor(CM.pred.knn),
  reference = liver.test.boxcox$status,
  positive = "Yes",
 prevalence = preval)
## Warning in confusionMatrix.default(data = as.factor(CM.pred.knn), reference =
## liver.test.boxcox$status, : Levels are not in the same order for reference and
## data. Refactoring data to match.
CM.knn
## Confusion Matrix and Statistics
##
             Reference
##
## Prediction No Yes
         No 33 82
         Yes 0
##
##
##
                  Accuracy: 0.287
##
                    95% CI: (0.2065, 0.3788)
##
       No Information Rate: 0.713
##
       P-Value [Acc > NIR] : 1
##
##
                     Kappa: 0
##
## Mcnemar's Test P-Value : <2e-16
##
##
               Sensitivity: 0.000
```

```
##
               Specificity: 1.000
##
            Pos Pred Value :
            Neg Pred Value : -1.509
##
##
                Prevalence: 2.509
            Detection Rate: 0.000
##
##
      Detection Prevalence: 0.000
##
         Balanced Accuracy: 0.500
##
##
          'Positive' Class : Yes
##
#confusion matrix for Naive Bayes
test.pred.nb <- rep("No", length(nb.pred))</pre>
test.pred.nb[nb.pred>0.5] <- "Yes"</pre>
CM.nb = confusionMatrix(data = as.factor(test.pred.nb),
  reference = liver.test.boxcox$status,
  positive = "Yes",
  prevalence = preval)
CM.nb
## Confusion Matrix and Statistics
##
##
             Reference
## Prediction No Yes
          No 11 12
##
##
          Yes 22 70
##
##
                  Accuracy : 0.7043
##
                    95% CI: (0.6121, 0.7858)
##
       No Information Rate: 0.713
       P-Value [Acc > NIR] : 0.6264
##
##
##
                     Kappa: 0.2056
##
##
   Mcnemar's Test P-Value: 0.1227
##
##
               Sensitivity: 0.8537
##
               Specificity: 0.3333
##
            Pos Pred Value: 1.8857
##
            Neg Pred Value: 3.7029
                Prevalence: 2.5091
##
##
            Detection Rate: 0.6087
##
      Detection Prevalence: 0.8000
##
         Balanced Accuracy: 0.5935
##
##
          'Positive' Class : Yes
##
```