

Initial Practice: Parkinson's Disease

Statistical Learning with Deep Artificial Neural Networks

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1 Introduction

We are working on a dataset describing Parkinson's disease. Click [here](#) for more information regarding the dataset. In short, the dataset is composed of a range of biomedical voice measurements from 42 people with early-stage Parkinson's disease recruited to a six-month trial of a telemonitoring device for remote symptom progression monitoring.

The main objective is to predict the severity of Parkinson's disease based on the data. More details are given in the following.

2 Load the Parkinsons Data

```
data <- read.csv("parkinsons_updrs.data")
str(data)
```

```
#> 'data.frame':    5875 obs. of  22 variables:
#> $ subject.      : int  1 1 1 1 1 1 1 1 1 1 ...
#> $ age           : int  72 72 72 72 72 72 72 72 72 72 ...
#> $ sex           : int  0 0 0 0 0 0 0 0 0 0 ...
#> $ test_time     : num  5.64 12.67 19.68 25.65 33.64 ...
#> $ motor_UPDRS   : num  28.2 28.4 28.7 28.9 29.2 ...
#> $ total_UPDRS   : num  34.4 34.9 35.4 35.8 36.4 ...
#> $ Jitter...     : num  0.00662 0.003 0.00481 0.00528 0.00335 0.00353 0.00422 0.00476 0.00432 0.00496 ...
#> $ Jitter.Abs.   : num  3.38e-05 1.68e-05 2.46e-05 2.66e-05 2.01e-05 ...
#> $ Jitter.RAP    : num  0.00401 0.00132 0.00205 0.00191 0.00093 0.00119 0.00212 0.00226 0.00156 0.002 ...
#> $ Jitter.PPQ5   : num  0.00317 0.0015 0.00208 0.00264 0.0013 0.00159 0.00221 0.00259 0.00207 0.00253 ...
#> $ Jitter.DDP    : num  0.01204 0.00395 0.00616 0.00573 0.00278 ...
#> $ Shimmer       : num  0.0256 0.0202 0.0168 0.0231 0.017 ...
#> $ Shimmer.dB.   : num  0.23 0.179 0.181 0.327 0.176 0.214 0.445 0.212 0.371 0.31 ...
#> $ Shimmer.APQ3  : num  0.01438 0.00994 0.00734 0.01106 0.00679 ...
#> $ Shimmer.APQ5  : num  0.01309 0.01072 0.00844 0.01265 0.00929 ...
#> $ Shimmer.APQ11: num  0.0166 0.0169 0.0146 0.0196 0.0182 ...
#> $ Shimmer.DDA   : num  0.0431 0.0298 0.022 0.0332 0.0204 ...
#> $ NHR           : num  0.0143 0.0111 0.0202 0.0278 0.0116 ...
#> $ HNR           : num  21.6 27.2 23 24.4 26.1 ...
#> $ RPDE          : num  0.419 0.435 0.462 0.487 0.472 ...
#> $ DFA           : num  0.548 0.565 0.544 0.578 0.561 ...
#> $ PPE           : num  0.16 0.108 0.21 0.333 0.194 ...
```

```
summary(data)
```

```
#>      subject.      age      sex      test_time
#> Min.   : 1.00   Min.   :36.0   Min.   :0.0000   Min.   : -4.263
#> 1st Qu.:10.00   1st Qu.:58.0   1st Qu.:0.0000   1st Qu.: 46.847
#> Median :22.00   Median :65.0   Median :0.0000   Median : 91.523
#> Mean   :21.49   Mean   :64.8   Mean   :0.3178   Mean   : 92.864
#> 3rd Qu.:33.00   3rd Qu.:72.0   3rd Qu.:1.0000   3rd Qu.:138.445
#> Max.   :42.00   Max.   :85.0   Max.   :1.0000   Max.   :215.490
#>      motor_UPDRS      total_UPDRS      Jitter...      Jitter.Abs.
#> Min.   : 5.038   Min.   : 7.00   Min.   :0.000830   Min.   :2.250e-06
#> 1st Qu.:15.000   1st Qu.:21.37   1st Qu.:0.003580   1st Qu.:2.244e-05
#> Median :20.871   Median :27.58   Median :0.004900   Median :3.453e-05
#> Mean   :21.296   Mean   :29.02   Mean   :0.006154   Mean   :4.403e-05
#> 3rd Qu.:27.596   3rd Qu.:36.40   3rd Qu.:0.006800   3rd Qu.:5.333e-05
#> Max.   :39.511   Max.   :54.99   Max.   :0.099990   Max.   :4.456e-04
```

```

#>      Jitter.RAP      Jitter.PPQ5      Jitter.DDP      Shimmer
#> Min.   :0.000330   Min.   :0.000430   Min.   :0.000980   Min.   :0.00306
#> 1st Qu.:0.001580   1st Qu.:0.001820   1st Qu.:0.004730   1st Qu.:0.01912
#> Median :0.002250   Median :0.002490   Median :0.006750   Median :0.02751
#> Mean   :0.002987   Mean   :0.003277   Mean   :0.008962   Mean   :0.03404
#> 3rd Qu.:0.003290   3rd Qu.:0.003460   3rd Qu.:0.009870   3rd Qu.:0.03975
#> Max.   :0.057540   Max.   :0.069560   Max.   :0.172630   Max.   :0.26863
#>      Shimmer.dB.      Shimmer.APQ3      Shimmer.APQ5      Shimmer.APQ11
#> Min.   :0.026       Min.   :0.00161       Min.   :0.00194       Min.   :0.00249
#> 1st Qu.:0.175       1st Qu.:0.00928       1st Qu.:0.01079       1st Qu.:0.01566
#> Median :0.253       Median :0.01370       Median :0.01594       Median :0.02271
#> Mean   :0.311       Mean   :0.01716       Mean   :0.02014       Mean   :0.02748
#> 3rd Qu.:0.365       3rd Qu.:0.02057       3rd Qu.:0.02375       3rd Qu.:0.03272
#> Max.   :2.107       Max.   :0.16267       Max.   :0.16702       Max.   :0.27546
#>      Shimmer.DDA      NHR      HNR      RPDE
#> Min.   :0.00484       Min.   :0.000286       Min.   : 1.659       Min.   :0.1510
#> 1st Qu.:0.02783       1st Qu.:0.010955       1st Qu.:19.406       1st Qu.:0.4698
#> Median :0.04111       Median :0.018448       Median :21.920       Median :0.5423
#> Mean   :0.05147       Mean   :0.032120       Mean   :21.680       Mean   :0.5415
#> 3rd Qu.:0.06173       3rd Qu.:0.031463       3rd Qu.:24.444       3rd Qu.:0.6140
#> Max.   :0.48802       Max.   :0.748260       Max.   :37.875       Max.   :0.9661
#>      DFA      PPE
#> Min.   :0.5140       Min.   :0.02198
#> 1st Qu.:0.5962       1st Qu.:0.15634
#> Median :0.6436       Median :0.20550
#> Mean   :0.6532       Mean   :0.21959
#> 3rd Qu.:0.7113       3rd Qu.:0.26449
#> Max.   :0.8656       Max.   :0.73173

```

3 Description of the Variables

As we have seen above, the dataset contains 5875 rows, i.e. 5875 measurements. The columns consist of **patient ID**, **age**, **sex**, **time interval since enrollment date**, **motor_UPDRS**, **total_UPDRS** and 16 voice biomedical measurements. The variables are the following

- subject. - The patient ID. Integer that uniquely identifies each subject.
- age - Age of each subject.
- sex - Gender of the subject; '0' = male and '1' = female.
- test_time - Time since recruitment into the trial. The integer part is the number of days since recruitment.
- motor_UPDRS - Clinician's motor UPDRS score, linearly interpolated.
- ctotat_UPDRS - Clinician's total UPDRS score, linearly interpolated.
- Jitter(%), Jitter(Abs), Jitter:RAP, Jitter:PPQ5, Jitter:DDP - Several measures of variation in fundamental frequency.
- Shimmer, Shimmer(dB), Shimmer:APQ3, Shimmer:APQ5, Shimmer:APQ11, Shimmer:DDA - Several measures of variation in amplitude.
- NHR, HNR - Two measures of ratio of noise to tonal components in the voice.
- RPDE - A nonlinear dynamical complexity measure.
- DFA - Signal fractal scaling exponent.
- PPE - A nonlinear measure of fundamental frequency variation.

As noted, the objective is to predict the severity of the disease, where severity is defined based on the variable **total_UPDRS**: The disease is severe if **total_UPDRS** > **25**. This variable is created below.

BESKRIV LITT MER KRING HVA DE 16 BIOMEDISINSKE VARIABLENE ER (IKKE BARE KOPIERE

FRA NETTSIDEN SLIK JEG HAR GJORT OVER!)

4 Create the Binary Variable of Parkinson's Severity

```
data$severity <- data$total_UPDRS > 25
dim(data)
```

```
#> [1] 5875 23
```

```
summary(data$severity)
```

```
#>   Mode  FALSE   TRUE
#> logical 2188  3687
```

5 Normalization

The variables of the 16 voice measurements are normalized by means of the min-max transformation.

```
normalize <- function(x) {
  return((x- min(x))/(max(x)-min(x)))
}
```

```
for (i in 1:16){
  data[, 6+i] <- normalize(data[,6+i])
}
```

```
summary(data)
```

```
#>   subject.      age      sex  test_time
#> Min.   : 1.00  Min.   :36.0  Min.   :0.0000  Min.   : -4.263
#> 1st Qu.:10.00  1st Qu.:58.0  1st Qu.:0.0000  1st Qu.: 46.847
#> Median :22.00  Median :65.0  Median :0.0000  Median : 91.523
#> Mean   :21.49  Mean   :64.8  Mean   :0.3178  Mean   : 92.864
#> 3rd Qu.:33.00  3rd Qu.:72.0  3rd Qu.:1.0000  3rd Qu.:138.445
#> Max.   :42.00  Max.   :85.0  Max.   :1.0000  Max.   :215.490
#> motor_UPDRS  total_UPDRS  Jitter...  Jitter.Abs.
#> Min.   : 5.038  Min.   : 7.00  Min.   :0.00000  Min.   :0.00000
#> 1st Qu.:15.000  1st Qu.:21.37  1st Qu.:0.02773  1st Qu.:0.04553
#> Median :20.871  Median :27.58  Median :0.04104  Median :0.07281
#> Mean   :21.296  Mean   :29.02  Mean   :0.05369  Mean   :0.09423
#> 3rd Qu.:27.596  3rd Qu.:36.40  3rd Qu.:0.06021  3rd Qu.:0.11523
#> Max.   :39.511  Max.   :54.99  Max.   :1.00000  Max.   :1.00000
#> Jitter.RAP  Jitter.PPQ5  Jitter.DDP  Shimmer
#> Min.   :0.00000  Min.   :0.00000  Min.   :0.00000  Min.   :0.00000
#> 1st Qu.:0.02185  1st Qu.:0.02011  1st Qu.:0.02185  1st Qu.:0.06047
#> Median :0.03356  Median :0.02980  Median :0.03361  Median :0.09207
#> Mean   :0.04645  Mean   :0.04118  Mean   :0.04650  Mean   :0.11664
#> 3rd Qu.:0.05174  3rd Qu.:0.04383  3rd Qu.:0.05179  3rd Qu.:0.13816
#> Max.   :1.00000  Max.   :1.00000  Max.   :1.00000  Max.   :1.00000
#> Shimmer.dB.  Shimmer.APQ3  Shimmer.APQ5  Shimmer.APQ11
#> Min.   :0.0000  Min.   :0.00000  Min.   :0.00000  Min.   :0.00000
#> 1st Qu.:0.0716  1st Qu.:0.04762  1st Qu.:0.05361  1st Qu.:0.04827
#> Median :0.1091  Median :0.07507  Median :0.08481  Median :0.07407
#> Mean   :0.1369  Mean   :0.09652  Mean   :0.11027  Mean   :0.09155
```

```

#> 3rd Qu.:0.1629 3rd Qu.:0.11775 3rd Qu.:0.13215 3rd Qu.:0.11073
#> Max. :1.0000 Max. :1.00000 Max. :1.00000 Max. :1.00000
#> Shimmer.DDA NHR HNR RPDE
#> Min. :0.00000 Min. :0.00000 Min. :0.0000 Min. :0.0000
#> 1st Qu.:0.04758 1st Qu.:0.01426 1st Qu.:0.4900 1st Qu.:0.3911
#> Median :0.07507 Median :0.02428 Median :0.5594 Median :0.4800
#> Mean :0.09650 Mean :0.04256 Mean :0.5528 Mean :0.4790
#> 3rd Qu.:0.11775 3rd Qu.:0.04168 3rd Qu.:0.6291 3rd Qu.:0.5681
#> Max. :1.00000 Max. :1.00000 Max. :1.0000 Max. :1.0000
#> DFA PPE severity
#> Min. :0.0000 Min. :0.0000 Mode :logical
#> 1st Qu.:0.2336 1st Qu.:0.1893 FALSE:2188
#> Median :0.3685 Median :0.2586 TRUE :3687
#> Mean :0.3959 Mean :0.2784
#> 3rd Qu.:0.5612 3rd Qu.:0.3417
#> Max. :1.0000 Max. :1.0000

```

6 Separation into Train and Test Data

I will use (pseudo-) random sampling to separate the data into a training and test set.

```

set.seed(1)
ratio <- 0.9
sample.size <- floor(nrow(data) * ratio)
train.indices <- sample(1:nrow(data), size = sample.size)
train <- data[train.indices, ]
test <- data[-train.indices, ]

x_train <- data.matrix(train[,-23])
y_train <- to_categorical(train[, 23], num_classes = 2)

#> Loaded Tensorflow version 2.7.1

x_test <- data.matrix(test[,-23])
y_test <- to_categorical(test[, 23], num_classes = 2)

```

7 Implementation of a Dense DNN

A dense deep neural network (DNN) for severity prediction is made. It has two hidden layers, with 10 nodes in each hidden layer. I have implemented two variants of this DNN; variant A has two output nodes, while variant B has only one output node.

7.1 Variant A: Two Output Nodes

Since we have two output nodes in this variant, we should use the *softmax* activation function in the output and the *categorical_crossentropy* loss function.

```

set.seed(1)
# defining the model and layers
model <- keras_model_sequential() %>%
  layer_dense(units = 10, activation = 'relu', input_shape = c(ncol(x_train))) %>%
  layer_dense(units = 10, activation = 'relu') %>%
  layer_dense(units = ncol(y_train), activation = 'softmax')

```

```
summary(model)
```

```
#> Model: "sequential"
```

```
#>
```

```
#> -----  
#> Layer (type)                Output Shape          Param #  
#> =====  
#> dense_2 (Dense)              (None, 10)            230  
#>  
#> dense_1 (Dense)              (None, 10)            110  
#>  
#> dense (Dense)                (None, 2)             22  
#>  
#> =====  
#> Total params: 362  
#> Trainable params: 362  
#> Non-trainable params: 0  
#> -----
```

```
# compile (define loss and optimizer)
```

```
model %>% compile(loss = 'categorical_crossentropy',  
                  optimizer = optimizer_rmsprop(),  
                  metrics = c('accuracy'))
```

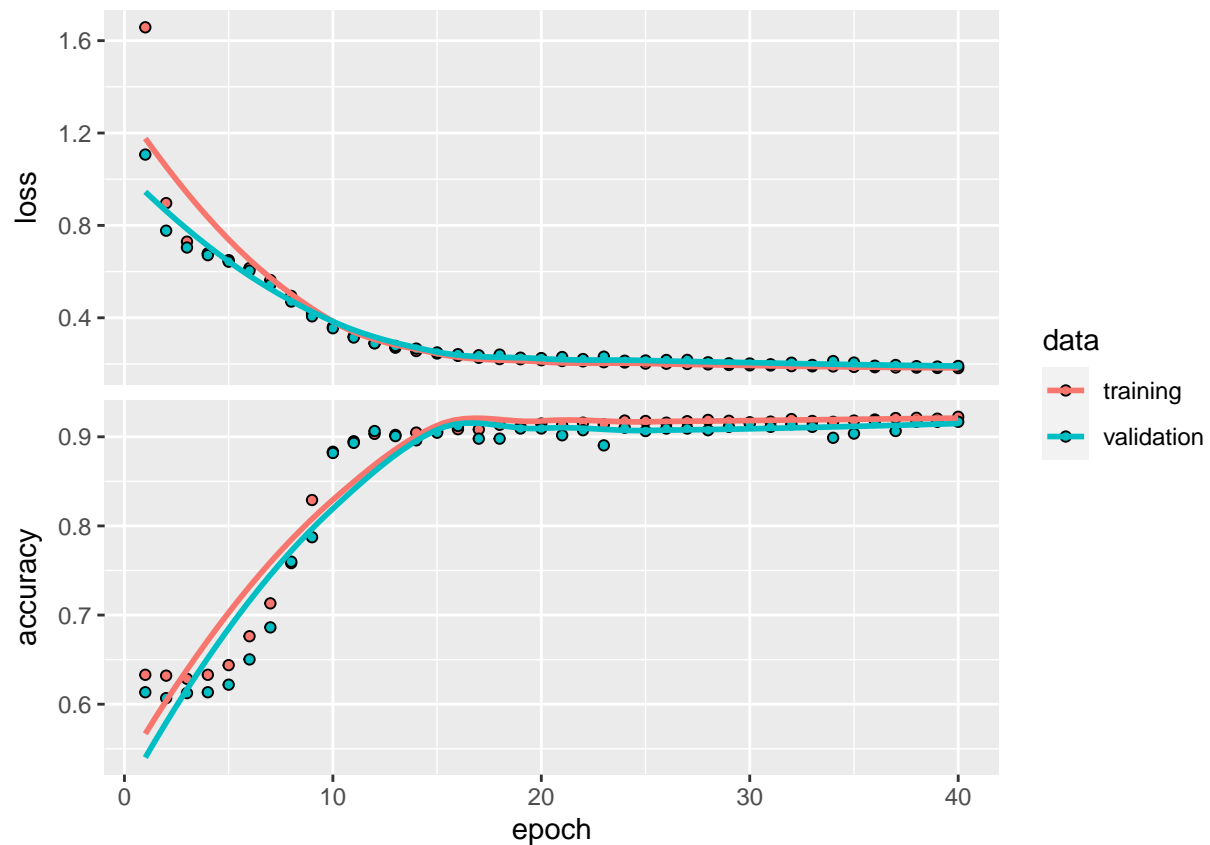
```
# train (fit)
```

```
history <- model %>% fit(data.matrix(x_train), y_train, epochs = 40,  
                        batch_size = 256, validation_split = 0.2)
```

```
# plot
```

```
plot(history)
```

```
#> `geom_smooth()` using formula 'y ~ x'
```



```
# evaluate on training data.
model %>% evaluate(x_train, y_train)
```

```
#>      loss  accuracy
#> 0.1847276 0.9175336
```

```
# evaluate on test data.
model %>% evaluate(x_test, y_test)
```

```
#>      loss  accuracy
#> 0.1473642 0.9438776
```

7.2 Variant B

Since we have one output nodes in this variant, we should use the *sigmoid* activation function in the output and the *binary_crossentropy* loss function.

```
set.seed(1)
# defining the model and layers
model2 <- keras_model_sequential() %>%
  layer_dense(units = 10, activation = 'relu', input_shape = c(ncol(x_train))) %>%
  layer_dense(units = 10, activation = 'relu') %>%
  layer_dense(units = 1, activation = 'sigmoid')

summary(model2)
```

```
#> Model: "sequential_1"
```

```
#> -----
#> Layer (type)                                     Output Shape                                     Param #
```

```
#> =====
#> dense_5 (Dense)                (None, 10)                230
#>
#> dense_4 (Dense)                (None, 10)                110
#>
#> dense_3 (Dense)                (None, 1)                 11
#>
#> =====
#> Total params: 351
#> Trainable params: 351
#> Non-trainable params: 0
#> -----
```

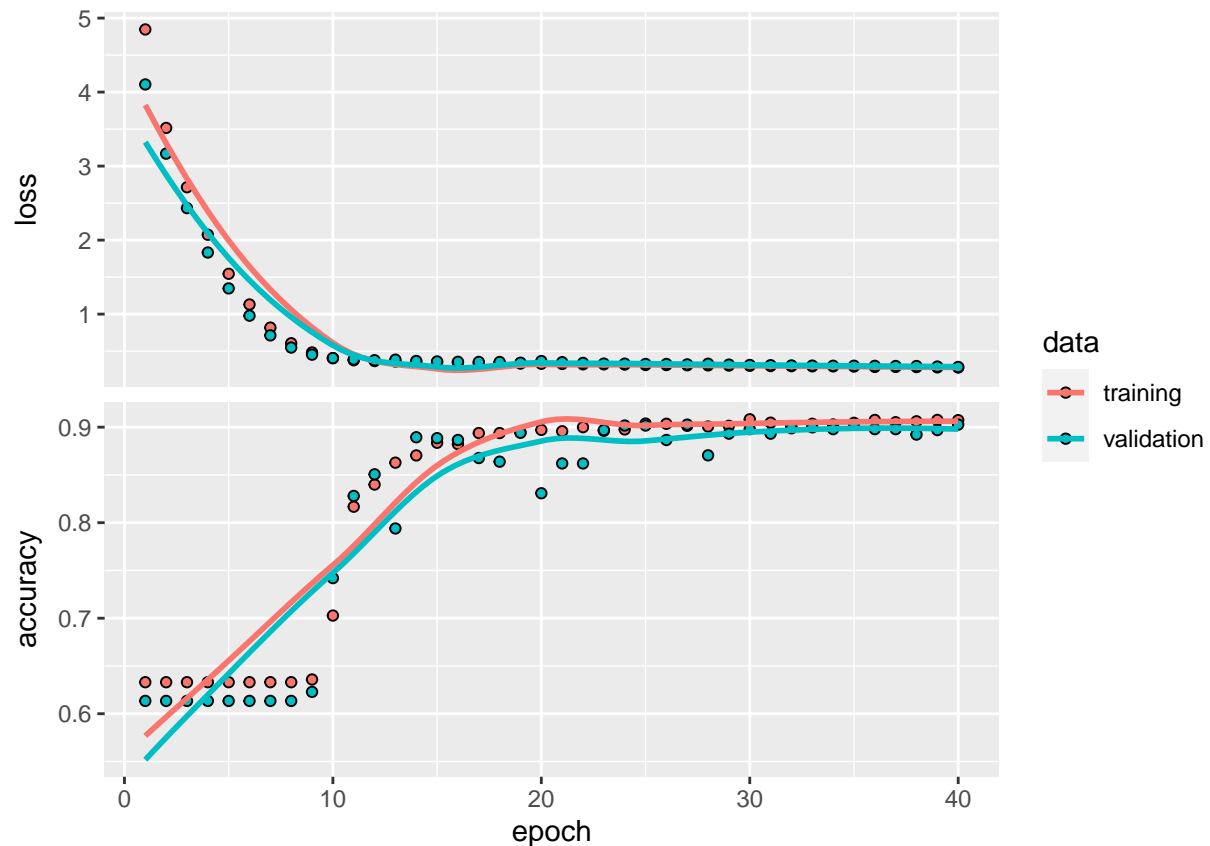
```
y_train2 <- as.numeric(data.matrix(train[,23]))
y_test2 <- as.numeric(data.matrix(test[,23]))

# compile (define loss and optimizer)
model2 %>% compile(loss = 'binary_crossentropy',
                  optimizer = optimizer_rmsprop(),
                  metrics = c('accuracy'))

# train (fit)
history2 <- model2 %>% fit(data.matrix(x_train), y_train2, epochs = 40,
                          batch_size = 256, validation_split = 0.2)

# plot
plot(history2)
```

```
#> `geom_smooth()` using formula 'y ~ x'
```




```
# evaluate on training data.
model2 %>% evaluate(x_train, y_train2)
```

```
#>      loss  accuracy
#> 0.2774411 0.9103462
```

```
# evaluate on testing data.
model2 %>% evaluate(x_test, y_test2)
```

```
#>      loss  accuracy
#> 0.2632757 0.9251701
```

8 Predictions

8.1 Predictions for Variant A

```
y_pred <- model %>% predict(x_test) %>% k_argmax()
y_pred <- as.array(y_pred)
(tab <- table("Predictions" = y_pred, "Labels" = test[, 23]))
```

```
#>      Labels
#> Predictions FALSE TRUE
#>      0    215    21
#>      1     12   340
```

```
# accuracy in predictions (as shown with the "evaluate" above).
(tab[1]+tab[4])/sum(tab)
```

```
#> [1] 0.9438776
```

8.2 Predictions for Variant B

```
# Predictions for one output node
y_pred2 <- model2 %>% predict(x_test) %>% `>`(0.5) %>% k_cast("int32")
y_pred2 <- as.array(y_pred2)
(tab2 <- table("Predictions" = y_pred2, "Labels" = test[, 23]))
```

```
#>      Labels
#> Predictions FALSE TRUE
#>      0    206    23
#>      1     21   338
```

```
# accuracy in predictions (as shown with the "evaluate" above).
(tab2[1]+tab2[4])/sum(tab2)
```

```
#> [1] 0.9251701
```