

HarvardX: PH125.9x Data Science

IDV Learners Capstone Project

Cancer treatment survival analysis and prediction project

Denis Korolskii

November 23, 2019

Overview

IDV Learners Capstone Project of the HarvardX: PH125.9x Data Science: Capstone course. Current task is to create recommendation system using a chosen dataset. Also, current task is to train a machine learning algorithm using the inputs in one subset to predict survival time in the test set.

Introduction

For this project we will focus on create a hypothesis testing and training prediction model system using Cameron and Pauling investigation dataset “Intravenous vitamin C in the supportive care of cancer patients: a review and rational approach”, <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5927785/>. Literature demonstrates that cancer patients experience vitamin C deficiency correlated with reduced oral intake, inflammation, infection, disease processes, and treatments such as radiation, chemotherapy, and surgery. Reaserch of the statistical significance of a possible difference in the effect of the Ascorbate treatment, as well as training the machine learning algorithm on the basis of the data obtained are the goals of this project.

##Executive summary

The evaluation of algorithm performance is the Root Mean Square Error. RMSE is a frequently used measure of the differences between values (sample or population values) predicted by a model or an estimator and the values observed. The RMSE represents the square root of the second sample moment of the differences between predicted values and observed values or the quadratic mean of these differences. These deviations are called residuals when the calculations are performed over the data sample that was used for estimation and are called errors (or prediction errors) when computed out-of-sample. The RMSE serves to aggregate the magnitudes of the errors in predictions for various times into a single measure of predictive power. RMSE is a measure of accuracy, to compare forecasting errors of different models for a particular dataset and not between datasets, as it is scale-dependent.

The function that computes the RMSE for vectors of ratings and their corresponding predictors will be the following:

$$RMSE = \sqrt{\frac{1}{N} \sum_{u,i} (\hat{y}_{u,i} - y_{u,i})^2}$$

Dataset

The cancer_1 dataset is uploaded at GitHub repository

https://github.com/DKorolski/homework-0/raw/8d656ba0abb62a8a48e611f8d5a2cebe4250bc97/cancer_1.xlsx

```
#Loading libraries
if(!require(tidyverse)) install.packages ("tidyverse", repos = "http://cran.us.r-project.org")
if(!require(caret)) install.packages ("caret", repos = "http://cran.us.r-project.org")
if(!require(data.table)) install.packages ("data.table", repos = "http://cran.us.r-project.org")
if(!require(car)) install.packages("car", repos = "http://cran.us.r-project.org")
if(!require(ggplot2)) install.packages("ggplot2", repos = "http://cran.us.r-project.org")
if(!require(readxl)) install.packages("readxl", repos = "http://cran.us.r-project.org")
#Loading dataset
z <- read_excel (path = "cancer_1.xlsx", sheet = "cancer")
```

Metadata Reference: Cameron, E. and Pauling, L. (1978) Supplemental ascorbate in the supportive treatment of cancer: re-evaluation of prolongation of survival times in terminal human cancer. Proceedings of the National Academy of Science USA. Also found in: Manly, B.F.J. (1986) Multivariate Statistical Methods: A Primer, New York: Chapman & Hall, 11. Also found in: Hand, D.J., et al. (1994) A Handbook of Small Data Sets, London: Chapman & Hall, 255. Description: Patients with advanced cancers of the stomach, bronchus, colon, ovary or breast were treated with ascorbate. The purpose of the study was to determine if the survival times differ with respect to the organ affected by the cancer. Number of cases: 64 Variable Names: Survival: Survival time (in days) Organ: Organ affected by the cancer relation 'Survival' numeric 'Organ' {"Breast", "Bronchus", "Colon", "Ovary", "Stomach"}

Methods and Analysis

Data Analysis

To get familiar with the dataset, we find the first rows of “cancer_1” subset as below. The subset contain the two variables “Survival”, “Organ”. Each row represent a single case.

Preprocessing

Testing for any N/A

```
colSums(is.na(z))
```

```
## Survival    Organ
##           0      0
```

Exploring dataset

```
str(z)
```

```
## Classes 'tbl_df', 'tbl' and 'data.frame':   64 obs. of  2 variables:
## $ Survival: num  124 42 25 45 412 ...
## $ Organ   : chr  "Stomach" "Stomach" "Stomach" "Stomach" ...
```

```
summary(z)
```

```
##      Survival      Organ
## Min.   : 20.0   Length:64
## 1st Qu.: 102.5   Class :character
## Median : 265.5   Mode  :character
## Mean    : 558.6
## 3rd Qu.: 721.0
## Max.    :3808.0
```

Converting characters to factor

```
z$Organ <- factor(z$Organ)
```

```
table(z$Organ)
```

```
##  
##   Breast Bronchus   Colon   Ovary Stomach  
##      11       17      17      6     13
```

```
table(z)
```

```
##           Organ  
## Survival Breast Bronchus Colon Ovary Stomach  
##    20         0         1     1     0     0  
##    24         1         0     0     0     0  
##    25         0         0     0     0     1  
##    37         0         1     0     0     0  
##    40         1         0     0     0     0  
##    42         0         0     0     0     1  
##    45         0         0     0     0     1  
##    46         0         0     0     0     1  
##    51         0         0     0     0     1  
##    63         0         1     0     0     0  
##    64         0         1     0     0     0  
##    72         0         1     0     0     0  
##    81         0         1     0     0     0  
##    89         0         0     0     1     0  
##   101         0         0     1     0     0  
##   103         0         0     0     0     1  
##   124         0         0     0     0     1  
##   138         0         1     0     0     0  
##   146         0         0     0     0     1  
##   151         0         1     0     0     0  
##   155         0         1     0     0     0  
##   163         0         0     1     0     0  
##   166         0         2     0     0     0  
##   180         0         0     1     0     0  
##   189         0         0     1     0     0  
##   201         0         0     0     1     0  
##   223         0         1     0     0     0  
##   245         0         1     0     0     0  
##   246         0         1     0     0     0  
##   248         0         0     1     0     0  
##   283         0         0     1     0     0  
##   340         0         0     0     0     1  
##   356         0         0     0     1     0  
##   365         0         0     1     0     0  
##   372         0         0     1     0     0  
##   377         0         0     1     0     0  
##   396         0         0     0     0     1  
##   406         0         0     1     0     0
```

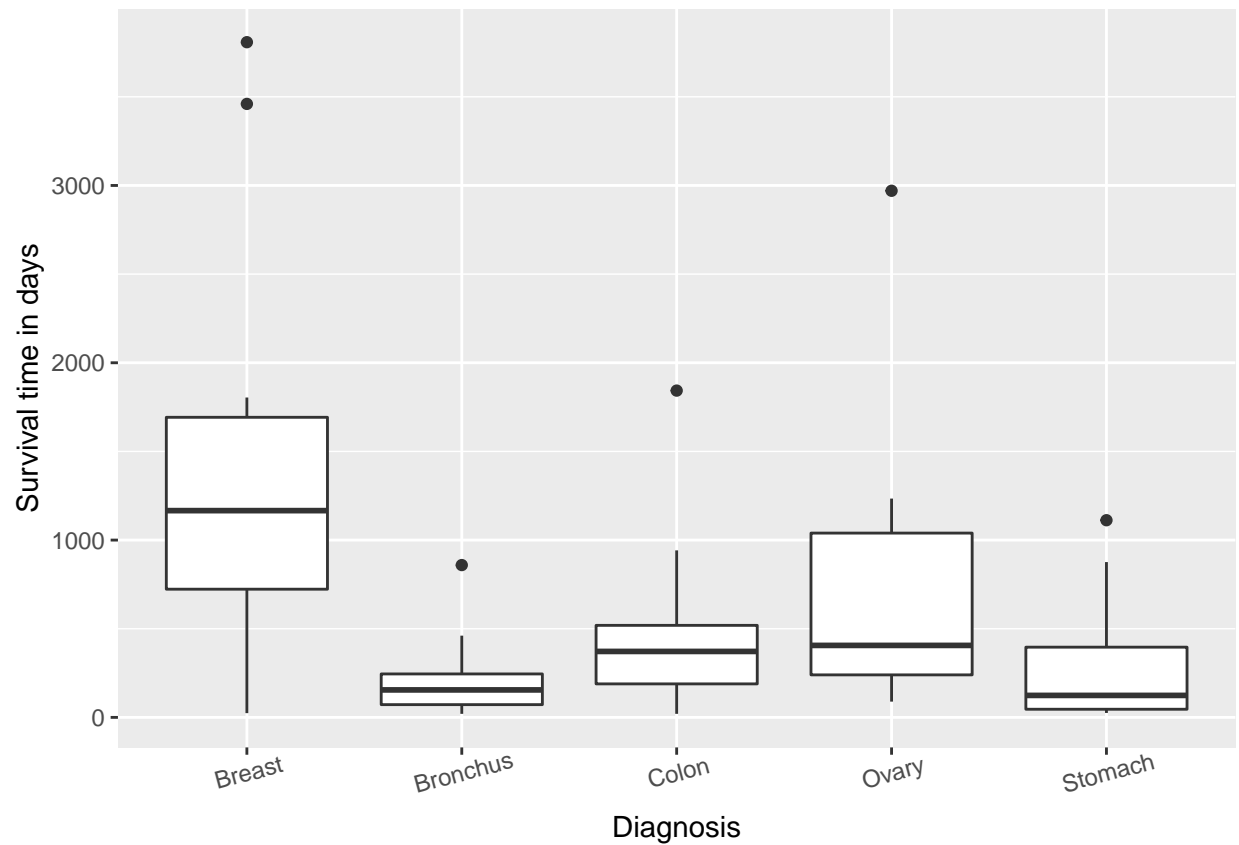
##	412	0	0	0	0	1
##	450	0	1	0	0	0
##	455	0	0	1	0	0
##	456	0	0	0	1	0
##	461	0	1	0	0	0
##	519	0	0	1	0	0
##	537	0	0	1	0	0
##	719	1	0	0	0	0
##	727	1	0	0	0	0
##	776	0	0	1	0	0
##	791	1	0	0	0	0
##	859	0	1	0	0	0
##	876	0	0	0	0	1
##	942	0	0	1	0	0
##	1112	0	0	0	0	1
##	1166	1	0	0	0	0
##	1234	0	0	0	1	0
##	1235	1	0	0	0	0
##	1581	1	0	0	0	0
##	1804	1	0	0	0	0
##	1843	0	0	1	0	0
##	2970	0	0	0	1	0
##	3460	1	0	0	0	0
##	3808	1	0	0	0	0

Dataset is small. Data is set of independent medical cases H0_hypothesis - survival time is equally dependent on organ (traditional) H1_hypothesis - survival time is not equally dependent on organ

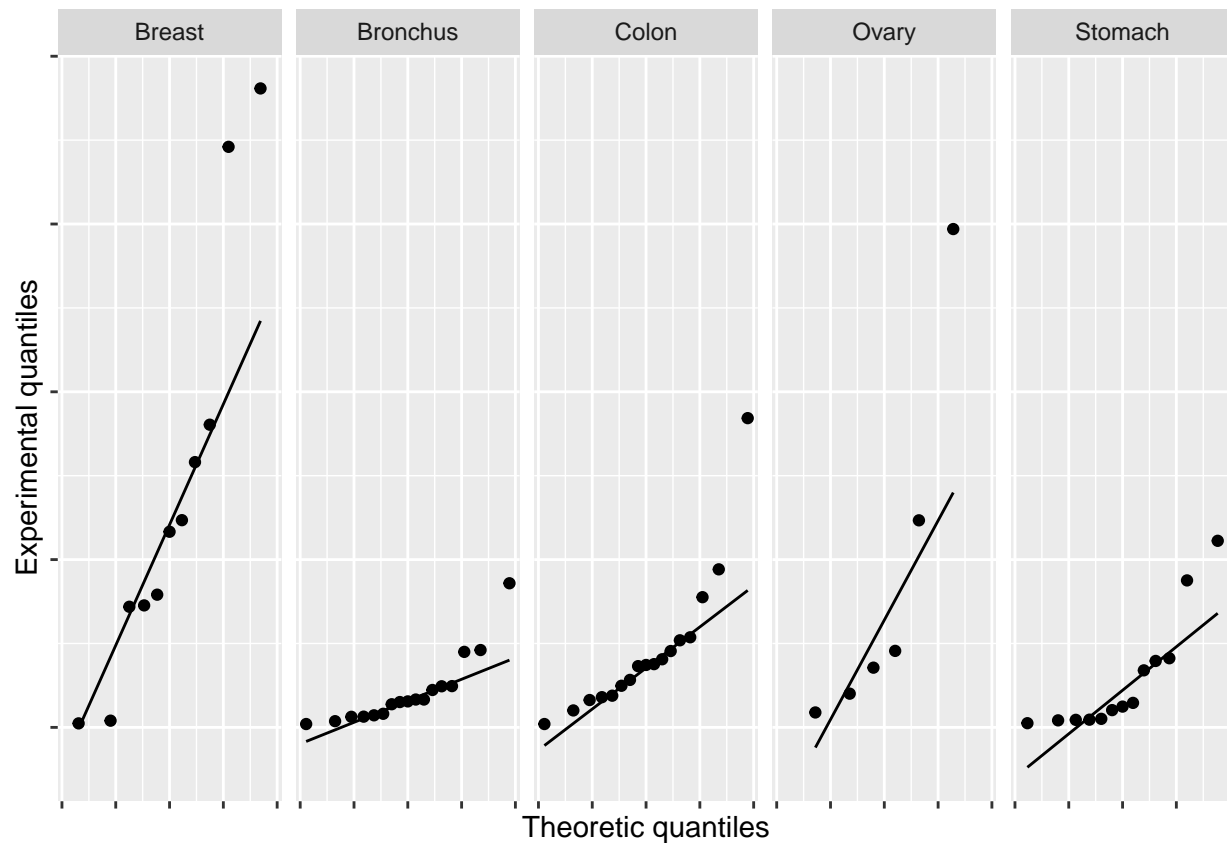
Plotting dataset distribution

The plotted boxplot shows the presence of extremely high survival rates in each group Since these outliers are uniformly distributed across all groups, it is more likely to conclude that these are not sample artifacts, but a strong sign of an asymmetric distribution. Their exclusion will lead to a distortion of the initial nature of the distribution, so it was decided to leave all the data in the array for further analysis.

```
#check for normal distribution
ggplot(data = z, aes(x = Organ, y = Survival)) +
  geom_boxplot() +
  labs(x = "Diagnosis", y = "Survival time in days") +
  theme(axis.text.x = element_text(angle = 15, vjust = 0.9, hjust = 0.5))
```



```
ggplot(aes(sample = Survival), data = z) +  
  geom_qq() + geom_qq_line() +  
  scale_x_continuous(labels = NULL, name = "Theoretic quantiles") +  
  scale_y_continuous(labels = NULL, name = "Experimental quantiles") +  
  facet_wrap(~Organ, ncol = 5)
```



Performing t-test

```
#Performing t-test comparing selected organ (breast) with others
Stomach <- (z$Survival[z$Organ == "Stomach"])
Breast <- (z$Survival[z$Organ == "Breast"])
Bronchus <- (z$Survival[z$Organ == "Bronchus"])
Colon <- (z$Survival[z$Organ == "Colon"])
Ovary <- (z$Survival[z$Organ == "Ovary"])
t_St <- t.test(Breast, Stomach)
t_Br <- t.test(Breast, Bronchus)
t_C <- t.test(Breast, Colon)
t_O <- t.test(Breast, Ovary)
p_vals <- c(t_St$p.value, t_Br$p.value, t_C$p.value, t_O$p.value)
#Performing Holm adjustment
p_holm <- p.adjust(p_vals, method = 'holm')
sum(p_holm <= 0.05)
```

```
## [1] 2
```

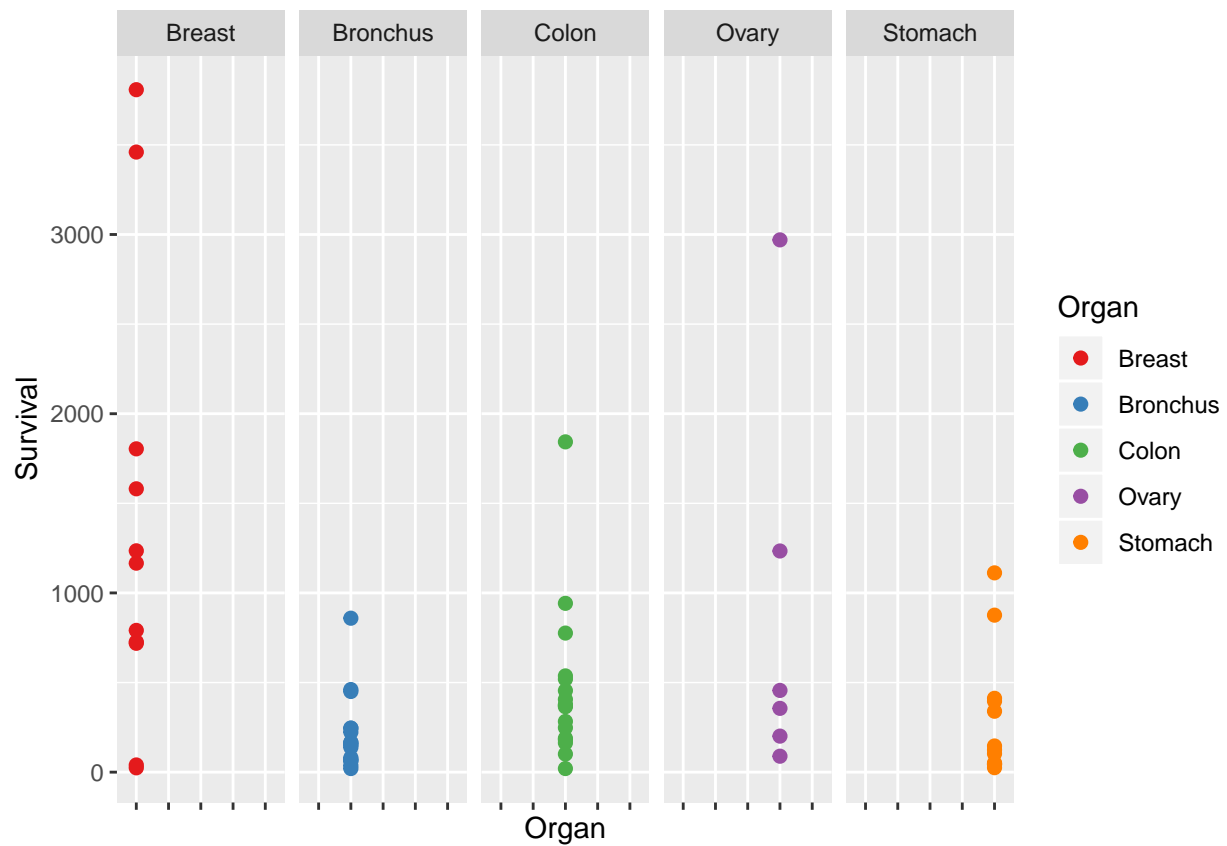
```
p_holm
```

```
## [1] 0.04391906 0.04018233 0.06595352 0.39862981
```

H1_hypothesis is correct for Breast-Stomach, Breast-Bronchus pairs

```
#plotting results
```

```
ggplot(z, aes(x = Organ, y = Survival, colour = Organ)) + geom_point(size = 2) + scale_colour_brewer(p
```



Regression model

Preparing train and test datasets for small dataset with normal distribution

```
# Set seed
set.seed (42)
#making permutations
n_obs <- nrow(z)
permuted_rows<-sample(n_obs)

z_shuffled <- z[permuted_rows, ]

# Identify row to split on: split
split <- round(n_obs * 0.6)

# Create train
train <- z_shuffled[1:split, ]
str(train)
```

```
## Classes 'tbl_df', 'tbl' and 'data.frame':   38 obs. of  2 variables:
## $ Survival: num  89 519 124 166 876 537 246 719 283 151 ...
## $ Organ   : Factor w/ 5 levels "Breast","Bronchus",...: 4 3 5 2 5 3 2 1 3 2 ...
```

```
# Create test
test <- z_shuffled[(split+1):n_obs, ]
str(test)
```

```
## Classes 'tbl_df', 'tbl' and 'data.frame': 26 obs. of 2 variables:
## $ Survival: num 45 776 72 40 396 406 163 42 859 180 ...
## $ Organ : Factor w/ 5 levels "Breast","Bronchus",...: 5 3 2 1 5 3 3 5 2 3 ...
```

1. Fit lm model on train: model

```
model <- lm (Survival ~ Organ , train)
head (model)
```

```
## $coefficients
## (Intercept) OrganBronchus OrganColon OrganOvary OrganStomach
## 1209.7143 -1021.2143 -637.0893 -325.3810 -818.1429
##
## $residuals
## 1 2 3 4 5 6
## -795.33333 -53.62500 -267.57143 -22.50000 484.42857 -35.62500
## 7 8 9 10 11 12
## 57.50000 -490.71429 -289.62500 -37.50000 720.42857 -482.71429
## 13 14 15 16 17 18
## -418.71429 2250.28571 -552.62500 -125.50000 -151.50000 371.28571
## 19 20 21 22 23 24
## -366.57143 369.37500 2085.66667 34.50000 -428.33333 -528.33333
## 25 26 27 28 29 30
## -324.62500 20.42857 -43.71429 1270.37500 -50.50000 -683.33333
## 31 32 33 34 35 36
## -383.62500 -1185.71429 56.50000 -245.57143 272.50000 -33.50000
## 37 38
## 349.66667 -345.57143
##
## $effects
## (Intercept) OrganBronchus OrganColon OrganOvary OrganStomach
## -3728.01048 -1533.48933 -605.08448 171.51444 1530.60513
##
## -10.68558 152.39015 -234.68337 -264.68558 57.39015
##
## 718.53414 -226.68337 -162.68337 2506.31663 -527.68558
##
## -30.60985 -56.60985 627.31663 -368.46586 394.31442
##
## 2248.05769 129.39015 -265.94231 -365.94231 -299.68558
##
## 18.53414 212.31663 1295.31442 44.39015 -520.94231
##
## -358.68558 -929.68337 151.39015 -247.46586 367.39015
##
## 61.39015 512.05769 -347.46586
##
## $rank
```



```
## [1] 5
##
## $fitted.values
##      1      2      3      4      5      6      7
## 884.3333 572.6250 391.5714 188.5000 391.5714 572.6250 188.5000
##      8      9     10     11     12     13     14
## 1209.7143 572.6250 188.5000 391.5714 1209.7143 1209.7143 1209.7143
##     15     16     17     18     19     20     21
## 572.6250 188.5000 188.5000 1209.7143 391.5714 572.6250 884.3333
##     22     23     24     25     26     27     28
## 188.5000 884.3333 884.3333 572.6250 391.5714 1209.7143 572.6250
##     29     30     31     32     33     34     35
## 188.5000 884.3333 572.6250 1209.7143 188.5000 391.5714 188.5000
##     36     37     38
## 188.5000 884.3333 391.5714
##
## $assign
## [1] 0 1 1 1 1
```

```
# Predict on test: p
p <- predict(model,test)
length(p)
```

```
## [1] 26
```

```
# Compute errors: error
error <- p - test[["Survival"]]
length(p)
```

```
## [1] 26
```

```
nrow(test)
```

```
## [1] 26
```

```
length(error)
```

```
## [1] 26
```

```
# Calculate RMSE
sqrt(mean(error^2))
```

```
## [1] 627.9954
```

2.Cross-Validation. This method is a good choice when we have a minimum amount of data and we get sufficiently big difference in quality or different optimal parameters between folds. As a general rule, we choose $k=5$ or $k=10$, as these values have been shown empirically to yield test error estimates that suffer neither from excessively high bias nor high variance.

```

#CROSS-VALIDATION
# Fit lm model using 10-fold CV: model
model <- train(
  Survival~Organ ,
  Z,
  method = "lm",
  trControl = trainControl(          #train-control func
    method = "cv",
    number = 10, #10-fold cross validation
    verboseIter = TRUE
  )
)

```

```

## + Fold01: intercept=TRUE
## - Fold01: intercept=TRUE
## + Fold02: intercept=TRUE
## - Fold02: intercept=TRUE
## + Fold03: intercept=TRUE
## - Fold03: intercept=TRUE
## + Fold04: intercept=TRUE
## - Fold04: intercept=TRUE
## + Fold05: intercept=TRUE
## - Fold05: intercept=TRUE
## + Fold06: intercept=TRUE
## - Fold06: intercept=TRUE
## + Fold07: intercept=TRUE
## - Fold07: intercept=TRUE
## + Fold08: intercept=TRUE
## - Fold08: intercept=TRUE
## + Fold09: intercept=TRUE
## - Fold09: intercept=TRUE
## + Fold10: intercept=TRUE
## - Fold10: intercept=TRUE
## Aggregating results
## Fitting final model on full training set

```

```

model

```

```

## Linear Regression
##
## 64 samples
## 1 predictor
##
## No pre-processing
## Resampling: Cross-Validated (10 fold)
## Summary of sample sizes: 58, 58, 59, 58, 58, 56, ...
## Resampling results:
##
##   RMSE      Rsquared   MAE
## 638.0567  0.3917217  456.2529
##
## Tuning parameter 'intercept' was held constant at a value of TRUE

```

```

# Fit lm model using 5 x 5-fold CV: model
model <- train(
  Survival~Organ,
  Z,
  method = "lm",
  trControl = trainControl(
    method = "repeatedcv",
    number = 5,
    repeats = 5,
    verboseIter = TRUE
  )
)

```

```

## + Fold1.Rep1: intercept=TRUE
## - Fold1.Rep1: intercept=TRUE
## + Fold2.Rep1: intercept=TRUE
## - Fold2.Rep1: intercept=TRUE
## + Fold3.Rep1: intercept=TRUE
## - Fold3.Rep1: intercept=TRUE
## + Fold4.Rep1: intercept=TRUE
## - Fold4.Rep1: intercept=TRUE
## + Fold5.Rep1: intercept=TRUE
## - Fold5.Rep1: intercept=TRUE
## + Fold1.Rep2: intercept=TRUE
## - Fold1.Rep2: intercept=TRUE
## + Fold2.Rep2: intercept=TRUE
## - Fold2.Rep2: intercept=TRUE
## + Fold3.Rep2: intercept=TRUE
## - Fold3.Rep2: intercept=TRUE
## + Fold4.Rep2: intercept=TRUE
## - Fold4.Rep2: intercept=TRUE
## + Fold5.Rep2: intercept=TRUE
## - Fold5.Rep2: intercept=TRUE
## + Fold1.Rep3: intercept=TRUE
## - Fold1.Rep3: intercept=TRUE
## + Fold2.Rep3: intercept=TRUE
## - Fold2.Rep3: intercept=TRUE
## + Fold3.Rep3: intercept=TRUE
## - Fold3.Rep3: intercept=TRUE
## + Fold4.Rep3: intercept=TRUE
## - Fold4.Rep3: intercept=TRUE
## + Fold5.Rep3: intercept=TRUE
## - Fold5.Rep3: intercept=TRUE
## + Fold1.Rep4: intercept=TRUE
## - Fold1.Rep4: intercept=TRUE
## + Fold2.Rep4: intercept=TRUE
## - Fold2.Rep4: intercept=TRUE
## + Fold3.Rep4: intercept=TRUE
## - Fold3.Rep4: intercept=TRUE
## + Fold4.Rep4: intercept=TRUE
## - Fold4.Rep4: intercept=TRUE
## + Fold5.Rep4: intercept=TRUE
## - Fold5.Rep4: intercept=TRUE

```

```
## + Fold1.Rep5: intercept=TRUE
## - Fold1.Rep5: intercept=TRUE
## + Fold2.Rep5: intercept=TRUE
## - Fold2.Rep5: intercept=TRUE
## + Fold3.Rep5: intercept=TRUE
## - Fold3.Rep5: intercept=TRUE
## + Fold4.Rep5: intercept=TRUE
## - Fold4.Rep5: intercept=TRUE
## + Fold5.Rep5: intercept=TRUE
## - Fold5.Rep5: intercept=TRUE
## Aggregating results
## Fitting final model on full training set
```

Calculating RMSE

```
model
```

```
## Linear Regression
##
## 64 samples
## 1 predictor
##
## No pre-processing
## Resampling: Cross-Validated (5 fold, repeated 5 times)
## Summary of sample sizes: 52, 50, 51, 51, 52, 51, ...
## Resampling results:
##
##      RMSE      Rsquared    MAE
##  668.4124  0.2849913  430.9667
##
## Tuning parameter 'intercept' was held constant at a value of TRUE
```

```
# Show the coefficients
mod <- lm(Survival~Organ, z)
coef(mod)
```

```
##      (Intercept) OrganBronchus   OrganColon   OrganOvary   OrganStomach
##      1395.9091   -1184.3209   -938.4973   -511.5758   -1109.9091
```

```
# Show the full output
summary(mod)
```

```
##
## Call:
## lm(formula = Survival ~ Organ, data = z)
##
## Residuals:
##      Min       1Q   Median       3Q      Max
## -1371.91  -241.75  -111.50    87.19   2412.09
##
## Coefficients:
##              Estimate Std. Error t value Pr(>|t|)
```

```
## (Intercept)      1395.9      201.9   6.915 3.77e-09 ***
## OrganBronchus   -1184.3      259.1  -4.571 2.53e-05 ***
## OrganColon      -938.5      259.1  -3.622 0.000608 ***
## OrganOvary      -511.6      339.8  -1.506 0.137526
## OrganStomach    -1109.9      274.3  -4.046 0.000153 ***
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 669.5 on 59 degrees of freedom
## Multiple R-squared:  0.3037, Adjusted R-squared:  0.2565
## F-statistic: 6.433 on 4 and 59 DF,  p-value: 0.0002295
```

```
# View summary of model
summary(mod)
```

```
##
## Call:
## lm(formula = Survival ~ Organ, data = z)
##
## Residuals:
##      Min       1Q   Median       3Q      Max
## -1371.91  -241.75  -111.50    87.19   2412.09
##
## Coefficients:
##              Estimate Std. Error t value Pr(>|t|)
## (Intercept)      1395.9      201.9   6.915 3.77e-09 ***
## OrganBronchus   -1184.3      259.1  -4.571 2.53e-05 ***
## OrganColon      -938.5      259.1  -3.622 0.000608 ***
## OrganOvary      -511.6      339.8  -1.506 0.137526
## OrganStomach    -1109.9      274.3  -4.046 0.000153 ***
## ---
## Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 669.5 on 59 degrees of freedom
## Multiple R-squared:  0.3037, Adjusted R-squared:  0.2565
## F-statistic: 6.433 on 4 and 59 DF,  p-value: 0.0002295
```

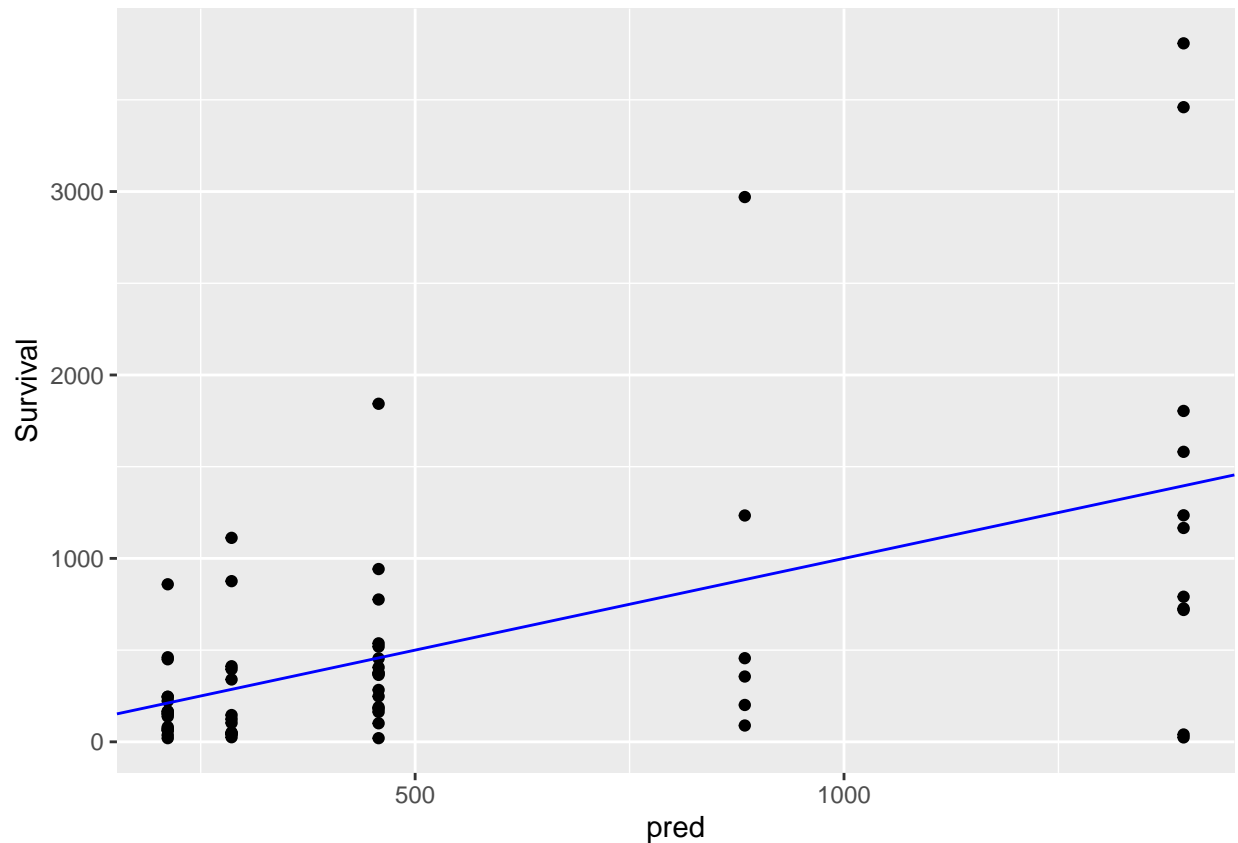
```
# Compute the mean of the residuals
mean(residuals(mod))
```

```
## [1] -2.4712e-14
```

```
# Compute RMSE
sqrt(sum(residuals(mod)^2) / df.residual(mod))
```

```
## [1] 669.5324
```

```
z$pred <- predict(model)
# Make a plot to compare predictions to actual (prediction on x axis).
ggplot(z, aes(x = pred, y = Survival)) +
  geom_point() +
  geom_abline(color = "blue")
```



```
3.Average+Organ effect system mu <- mean(train$Survival)
survival_avg <- train$Survival - mu
rmse_1 <- RMSE(test$Survival, predicted_ratings_bi)
rmse_1
```

#Conclusion.This IDV project was examined to observe data, check hypothesis and to predict survival time. The model evaluation performance through the RMSE (root mean squared error) showed that the Linear regression models are useful to predict survival time on the test set.

Appendix - Enviroment

```
print("Operating System:")
```

```
## [1] "Operating System:"
```

```
version
```

```
##
## platform      _
## arch          x86_64-w64-mingw32
## os            x86_64
## system        mingw32
## status        x86_64, mingw32
## major         3
## minor         6.1
```

```
## year          2019
## month         07
## day           05
## svn rev       76782
## language      R
## version.string R version 3.6.1 (2019-07-05)
## nickname      Action of the Toes
```