IDENTIFICATION OF KEY PREDICTIVE FACTORS IMPACTING MORTALITY IN HEART FAILURE (HF) PATIENTS

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mimicIII_data1= read.csv("C:\\Users\\CEO\\Documents\\DATA 603\\Project\\mimicIII_data1.csv")
head(mimicIII_data1, 5)

##		age_group	<pre>gender_tx</pre>	t group	ID	outcome	age	gender	BMI	hypertensive
##	1	18-29		F 1	197509	0	25	1	57.92653	0
##	2	18-29		F 1	183860	0	19	1	29.78897	0
##	3	18-29		F 1	165367	1	28	1	80.00000	0
##	4	30-39		F 1	184941	0	39	1	29.35654	1
##	5	30-39		F 1	195748	0	38	1	27.65766	0
##		atrialfib	rillation	CHD_wit	h_no_MI	diabete	s de	ficienc	yanemias (depression
##	1		0		0		0		0	0
##	2		0		0		0		0	0
##	3		0		0		0		0	0
##	4		0		0		0		0	0
##	5		0		0		0		0	0
##		Hyperlipen	nia Renal_	failure	COPD h	eart_rat	e Sy	stolic_	blood_pre	ssure
##	1		0	0	0	90.3600	0		95.0	04545
##	2		0	0	0	90.6521	7		99.	69565
##	3		0	0	0	87.6551	7		111.3	35294
##	4		1	0	0	63.7200	0		133.	66667
##	5		1	0	0	110.8400	0		109.	20833
##		Diastolic_	_blood_pre	ssure Ro	espirat	ory_rate	tem	peratur	e SP_0:	2 Urine_output
##	1		58.	00000		24.72000		36.1888	96.1200	0 4600
##	2		50.	08696		21.69565		38.1666	7 96.82609	9 5050
##	3		58.	76471		23.45946	;	37.4259	3 91.0937	5 4775
##	_		80.	37500		17.27273			2 95.6800	
##	5			29167		26.00000			3 99.4782	
##		hematocrit				MCHC	MC'			cyte Platelets
##										7778 347.4444
##			4.256000							
##			4.758750							
##	4		5.292500							0000 212.0000
##	5		3 4.237778			2222 98.	0000			
##		Neutrophil	_		-	PT			Γ_proBNP (Creatine_kinase
##	1	78.65				17.88571			2982	556.5
	2	79.50				16.65397			1968	556.5
	3	78.70				15.42222			2420	556.5
##	4	92.20	0.	10	4.400	12.83333	1.1	00000	3143	142.0

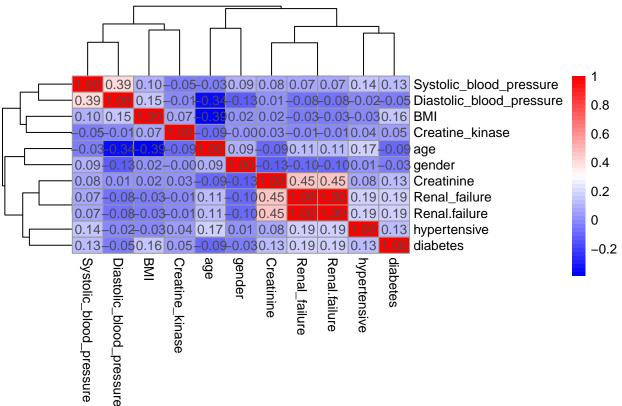
```
## 5
          80.275
                      0.90
                                9.925 19.39444 1.755556
                                                              8474
                                                                             118.0
    Creatinine Urea_nitrogen glucose Blood_potassium Blood_sodium Blood_calcium
                    31.88889 248.4000
## 1 1.3111111
                                              3.666667
                                                            131.3333
                                                                          9.170588
## 2 0.8600000
                     8.40000 84.0000
                                               4.200000
                                                            139.4000
                                                                          8.300000
## 3 0.7545455
                     11.63636 113.8750
                                               3.757143
                                                            142.4545
                                                                          8.687500
## 4 1.1200000
                     26.50000 120.3333
                                               4.470000
                                                            138.5000
                                                                          9.066667
## 5 1.2076923
                     23.38462 97.5000
                                               4.100000
                                                            135.0769
                                                                          8.154545
##
      Chloride Anion_gap Magnesium_ion
                                             PH Bicarbonate Lactic acid
                                                                             PC02
## 1 89.42857 15.33333
                              2.184211 7.311250
                                                    30.05556
                                                                2.800000 65.16667
## 2 103.20000 14.80000
                              1.950000 7.290000
                                                    25.60000
                                                                1.500000 52.00000
## 3 95.45455 11.27273
                              2.377778 7.332500
                                                   39.63636
                                                                1.083333 78.33333
## 4 93.60000 11.60000
                              2.175000 7.441000
                                                    37.80000
                                                                2.350000 60.50000
                              1.950000 7.418333
                                                                3.512500 31.66667
## 5 98.38462 16.61538
                                                    24.23077
##
   EF outcome_txt
## 1 20
              Alive
## 2 55
              Alive
## 3 50
               Dead
## 4 55
              Alive
## 5 15
              Alive
library(dplyr)
##
## Attaching package: 'dplyr'
## The following objects are masked from 'package:stats':
##
##
       filter, lag
## The following objects are masked from 'package:base':
##
##
       intersect, setdiff, setequal, union
# Select relevant columns
selected_data <- mimicIII_data1 %>%
  select(Creatinine, age, gender, BMI, Systolic_blood_pressure, Diastolic_blood_pressure, hypertensive,
# Convert categorical variables (like gender, hypertensive, diabetes, and Renal_failure) to numeric if
selected_data$gender <- as.numeric(factor(selected_data$gender))</pre>
selected_data$hypertensive <- as.numeric(factor(selected_data$hypertensive))</pre>
selected_data$diabetes <- as.numeric(factor(selected_data$diabetes))</pre>
selected_data$Renal_failure <- as.numeric(factor(selected_data$Renal_failure))</pre>
# Calculate correlation matrix
correlation_matrix <- cor(selected_data, use = "complete.obs")</pre>
# Extract correlation of creatinine with other variables
creatinine_correlation <- correlation_matrix["Creatinine", ]</pre>
# Display the result
print(creatinine correlation)
```

```
##
                Creatinine
                                                                     gender
                                                age
##
                1.00000000
                                        -0.08637138
                                                                -0.13092888
##
                       BMI Systolic_blood_pressure Diastolic_blood_pressure
                                       0.07951695
##
                0.01722883
                                                                 0.00566448
##
              hypertensive
                                           diabetes
                                                              Renal_failure
##
                0.07520727
                                         0.12865092
                                                                 0.44895233
summary(mimicIII_data1[c("age", "BMI", "Systolic_blood_pressure", "Diastolic_blood_pressure", "hyperten
##
        age
                        BMI
                                   Systolic_blood_pressure
## Min.
         :19.00
                          :13.35
                                   Min. : 75.0
                   Min.
   1st Qu.:65.00
                  1st Qu.:25.26
                                   1st Qu.:105.5
## Median :77.00
                 Median :28.29
                                   Median :116.0
## Mean
         :74.09
                   Mean
                         :29.82
                                   Mean
                                        :117.9
                   3rd Qu.:32.55
## 3rd Qu.:85.00
                                   3rd Qu.:128.5
## Max.
          :99.00
                   Max.
                          :80.00
                                   Max.
                                        :203.0
## Diastolic_blood_pressure hypertensive
                                                             Renal_failure
                                               diabetes
## Min. : 24.74
                           Min.
                                   :0.0000 Min. :0.0000 Min. :0.0000
## 1st Qu.: 52.28
                                            1st Qu.:0.0000 1st Qu.:0.0000
                            1st Qu.:0.0000
## Median : 58.36
                           Median: 1.0000 Median: 0.0000 Median: 0.0000
## Mean : 59.50
                            Mean :0.7183
                                            Mean :0.4213
                                                             Mean :0.3651
## 3rd Qu.: 65.38
                            3rd Qu.:1.0000
                                            3rd Qu.:1.0000
                                                             3rd Qu.:1.0000
## Max. :107.00
                            Max. :1.0000
                                            Max.
                                                  :1.0000
                                                             Max.
                                                                    :1.0000
# Define the columns you want to compute quantiles for
columns <- c("BMI", "Systolic_blood_pressure", "Diastolic_blood_pressure"</pre>
            )
# Apply the quantile function to each column
quantile_results <- sapply(columns, function(col) {
 quantile(mimicIII_data1[[col]], na.rm = TRUE)
})
quantile_results
##
            BMI Systolic_blood_pressure Diastolic_blood_pressure
## 0%
       13.34680
                                75.0000
                                                       24.73684
## 25%
       25.25823
                               105.4808
                                                       52.28407
## 50%
       28.28862
                               116.0000
                                                       58.36364
## 75% 32.54872
                              128.4929
                                                       65.38455
## 100% 80.00000
                               203.0000
                                                      107.00000
#install.packages("pheatmap")
library(pheatmap)
```

Warning: package 'pheatmap' was built under R version 4.4.2

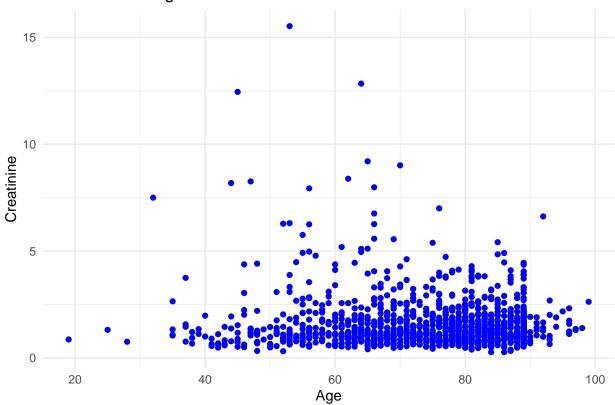
```
# Load necessary library
library(dplyr)
# Select relevant columns (make sure the column names match your data)
selected data <- mimicIII data1 %>%
  select (Creatinine, age, gender, BMI, Systolic_blood_pressure, Diastolic_blood_pressure, hypertensive,
# Convert categorical variables to numeric if necessary
selected_data$gender <- as.numeric(factor(selected_data$gender))</pre>
selected_data$hypertensive <- as.numeric(factor(selected_data$hypertensive))</pre>
selected_data$diabetes <- as.numeric(factor(selected_data$diabetes))</pre>
selected_data$Renal.failure <- as.numeric(factor(selected_data$Renal_failure))</pre>
# Calculate the correlation matrix
correlation_matrix <- cor(selected_data, use = "complete.obs")</pre>
# Generate the heatmap with annotations
pheatmap(correlation_matrix,
         display_numbers = TRUE,
                                       # Annotate with correlation values
         color = colorRampPalette(c("blue", "white", "red"))(50), # Color gradient
         main = "Correlation Matrix of Creatinine with Other Variables",
         fontsize number = 10)
                                       # Adjust the font size of annotations
```

Correlation Matrix of Creatinine with Other Variables

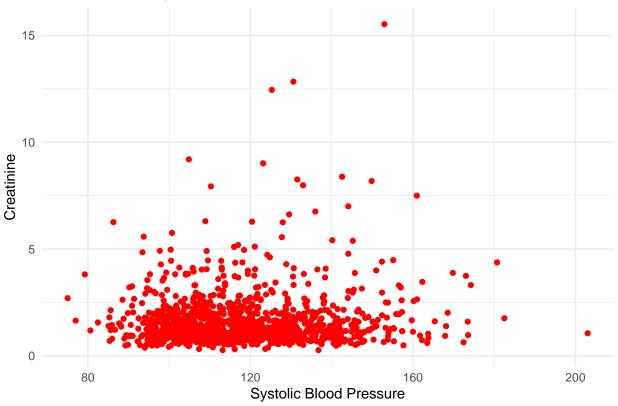


```
##
## Call:
## lm(formula = Creatinine ~ age + gender + BMI + Systolic_blood_pressure +
      Diastolic_blood_pressure + hypertensive + diabetes + Renal_failure,
##
      data = mimicIII_data1)
##
## Residuals:
##
      Min
              1Q Median
                            3Q
## -1.9055 -0.5792 -0.2166 0.2360 13.9176
##
## Coefficients:
                         Estimate Std. Error t value Pr(>|t|)
##
                         ## (Intercept)
                        ## age
## gender
                        ## BMI
                         -0.004286
                                   0.004369 -0.981 0.32686
## Systolic_blood_pressure
                                   0.002168
                                            2.186 0.02899 *
                         0.004739
## Diastolic_blood_pressure -0.004678
                                   0.003712 -1.260 0.20785
## hypertensive
                         0.008698
                                   0.076412 0.114 0.90939
## diabetes
                         0.058274
                                   0.070304
                                            0.829 0.40734
## Renal_failure
                         1.176358
                                   0.071520 16.448 < 2e-16 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## Residual standard error: 1.125 on 1166 degrees of freedom
## Multiple R-squared: 0.2305, Adjusted R-squared: 0.2252
## F-statistic: 43.65 on 8 and 1166 DF, p-value: < 2.2e-16
library(ggplot2)
# Scatter Plot
ggplot(mimicIII_data1, aes(x = age, y = Creatinine)) +
 geom point(color = "blue") +
 labs(title = "Creatinine vs. Age", x = "Age", y = "Creatinine") +
 theme_minimal()
```

Creatinine vs. Age



Creatinine vs. Systolic Blood Pressure



reducedModel= lm(Creatinine~age+gender+Systolic_blood_pressure+Renal_failure, data= mimicIII_data1) summary(reducedModel)

```
##
## Call:
  lm(formula = Creatinine ~ age + gender + Systolic_blood_pressure +
      Renal_failure, data = mimicIII_data1)
##
##
## Residuals:
               1Q Median
##
      Min
                               3Q
                                      Max
## -1.7741 -0.5761 -0.2302 0.2479 14.0357
## Coefficients:
                           Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                           1.993144
                                      0.302217
                                               6.595 6.42e-11 ***
## age
                          -0.012209
                                      0.002489 -4.905 1.07e-06 ***
                                      0.066649 -3.091 0.00204 **
## gender
                          -0.205994
## Systolic_blood_pressure 0.003645
                                      0.001916
                                                 1.902 0.05745 .
## Renal failure
                           1.198152
                                      0.069228 17.307 < 2e-16 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
##
## Residual standard error: 1.125 on 1170 degrees of freedom
## Multiple R-squared: 0.2282, Adjusted R-squared: 0.2256
## F-statistic: 86.48 on 4 and 1170 DF, p-value: < 2.2e-16
```

Null Hypothesis (H_0) : The predictor variables removed has no significant effect on the affects Creatinine.

Alternative Hypothesis (H_a) : The predictor variables removed has a significant effect on the affects Creatinine.

$$H_0: \beta_k = 0$$

$$H_a: \beta_k \neq 0 \quad (k = 1, 2, \dots, p)$$

```
anova(reducedModel,fullModel1)
```

```
## Analysis of Variance Table
##
## Model 1: Creatinine ~ age + gender + Systolic_blood_pressure + Renal_failure
## Model 2: Creatinine ~ age + gender + BMI + Systolic_blood_pressure + Diastolic_blood_pressure +
## hypertensive + diabetes + Renal_failure
## Res.Df RSS Df Sum of Sq F Pr(>F)
## 1 1170 1480.1
## 2 1166 1475.7 4 4.3443 0.8581 0.4885
```

At p-value > 0.05, we fail to reject the null hypothesis;

```
reducedModel1= lm(Creatinine~age+gender+Renal_failure, data= mimicIII_data1)
summary(reducedModel1)
```

```
##
## Call:
## lm(formula = Creatinine ~ age + gender + Renal_failure, data = mimicIII_data1)
##
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -1.7647 -0.5646 -0.2239 0.2250 14.1569
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                 2.416941
                            0.204370 11.826 < 2e-16 ***
                            0.002489 -4.997 6.72e-07 ***
## age
                -0.012438
                -0.193681
                            0.066408 -2.917 0.00361 **
## gender
## Renal_failure 1.209703
                            0.069038 17.522 < 2e-16 ***
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## Residual standard error: 1.126 on 1171 degrees of freedom
## Multiple R-squared: 0.2258, Adjusted R-squared: 0.2238
## F-statistic: 113.8 on 3 and 1171 DF, p-value: < 2.2e-16
```

We choose reduced Model due to reduced Model having higher $R_{adj}^2=0.2256$ and lower RSE 1.125 compared to reduced Model1

InteractReducedModel= lm(Creatinine~(age+gender+Systolic_blood_pressure+Renal_failure)^2, data= mimicII summary(InteractReducedModel)

```
## Call:
## lm(formula = Creatinine ~ (age + gender + Systolic_blood_pressure +
      Renal_failure)^2, data = mimicIII_data1)
##
## Residuals:
               10 Median
      Min
                               30
## -3.1460 -0.5090 -0.2148 0.2194 14.0328
##
## Coefficients:
                                         Estimate Std. Error t value Pr(>|t|)
                                        -0.5202731 1.3788044 -0.377 0.70599
## (Intercept)
## age
                                        0.0430143 0.0169161
                                                               2.543 0.01113
## gender
                                       -0.9130136 0.5874129 -1.554 0.12039
## Systolic_blood_pressure
                                                              1.591 0.11191
                                        0.0175249 0.0110161
## Renal_failure
                                        3.5532837 0.6584807
                                                               5.396 8.24e-08
                                       -0.0003066 0.0048859 -0.063 0.94998
## age:gender
## age:Systolic_blood_pressure
                                       -0.0003662 0.0001336 -2.740 0.00623
## age:Renal_failure
                                       0.0061507 0.0038091
## gender:Systolic_blood_pressure
                                                               1.615 0.10664
## gender:Renal_failure
                                       -0.0407685 0.1364415 -0.299 0.76515
## Systolic_blood_pressure:Renal_failure 0.0049952 0.0038601
                                                              1.294 0.19590
##
## (Intercept)
## age
## gender
## Systolic_blood_pressure
## Renal_failure
## age:gender
## age:Systolic_blood_pressure
## age:Renal_failure
                                        ***
## gender:Systolic_blood_pressure
## gender:Renal_failure
## Systolic_blood_pressure:Renal_failure
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
## Residual standard error: 1.094 on 1164 degrees of freedom
## Multiple R-squared: 0.2737, Adjusted R-squared: 0.2675
## F-statistic: 43.87 on 10 and 1164 DF, p-value: < 2.2e-16
InteractModelBest= lm(Creatinine~age+Systolic_blood_pressure+Renal_failure+age*Systolic_blood_pressure+
summary(InteractModelBest)
##
## Call:
## lm(formula = Creatinine ~ age + Systolic_blood_pressure + Renal_failure +
      age * Systolic_blood_pressure + age * Renal_failure, data = mimicIII_data1)
##
##
## Residuals:
      Min
               1Q Median
                               3Q
## -3.1149 -0.4931 -0.2158 0.1975 13.9799
```

##

Coefficients:

```
##
                                Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                              -1.8227156 1.1518748 -1.582 0.11383
                               0.0386174 0.0155804
                                                     2.479 0.01333 *
## age
## Systolic_blood_pressure
                                                     2.738 0.00628 **
                               0.0267917 0.0097852
## Renal_failure
                               4.0849817 0.4183529
                                                     9.764 < 2e-16 ***
## age:Systolic_blood_pressure -0.0003423 0.0001323 -2.586 0.00982 **
## age:Renal_failure
                              -0.0381749 0.0054792 -6.967 5.39e-12 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 1.099 on 1169 degrees of freedom
## Multiple R-squared: 0.264, Adjusted R-squared: 0.2609
## F-statistic: 83.86 on 5 and 1169 DF, p-value: < 2.2e-16
#summary(reducedModel)
```

Null Hypothesis $((H_0))$: The reduced model is a best model

Alternative Hypothesis $((H_a))$: The reduced model is not a best model

```
reducedModel1= lm(Creatinine~age+gender+Renal_failure, data= mimicIII_data1)
InteractModelBest= lm(Creatinine~age+Systolic_blood_pressure+Renal_failure+age*Systolic_blood_pressure+anova(InteractModelBest, reducedModel)
```

```
## Analysis of Variance Table
##
## Model 1: Creatinine ~ age + Systolic_blood_pressure + Renal_failure +
## age * Systolic_blood_pressure + age * Renal_failure
## Model 2: Creatinine ~ age + gender + Systolic_blood_pressure + Renal_failure
## Res.Df RSS Df Sum of Sq F Pr(>F)
## 1 1169 1411.4
## 2 1170 1480.1 -1 -68.668 56.874 9.296e-14 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
```

The InteractModelBest model has higher $R_{adj}^2 = 0.2609$ and lower RSE of 1.099 meaning that InteractModelBest model is better who compared to the Reduced Model with $R_{adj}^2 = 2256$ and RSE =1125

From the With a p-value of 9.296e-14 < 0.05 significance, we reject the null hypothesis and accept the alternate hypothesis meaning that InteractModelBest model is the best fit model to be used to prediction

The best fit model

```
Creatinine = -1.8227156 + 0.0386174age + 0.0267917Systolic_blood_pressure + 4.0849817Renal_failure \\ -0.0003423age * Systolic_blood_pressure - 0.0381749age * Renal_failure
```

```
newData1 = data.frame(age =72 ,Systolic_blood_pressure=155.8667, Renal_failure= 1 )
predict(InteractModelBest, newData1,interval = "predict")
```

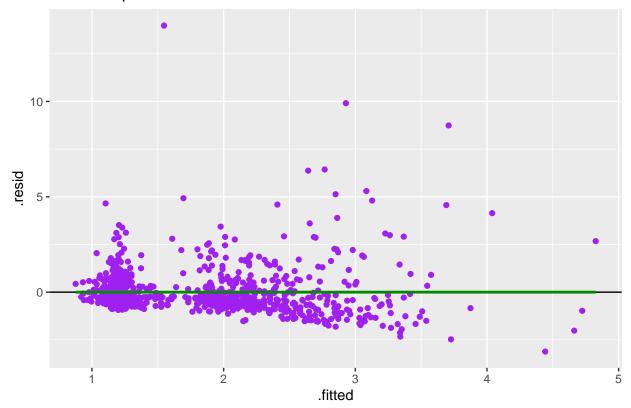
```
## fit lwr upr
## 1 2.629 0.4664461 4.791554
```

The response value is between the lower and upper limit of the predicted value, which signifies that the model predicted correctly and can be used for further prediction Cretinine in indiviuals.

```
library(ggplot2)
ggplot(InteractModelBest, aes(x=.fitted, y=.resid)) +
  geom_point(colour = "purple") +
  geom_hline(yintercept = 0) +
  geom_smooth(colour = "green4")+
  ggtitle("Residual plot: Residual vs Fitted values")
```

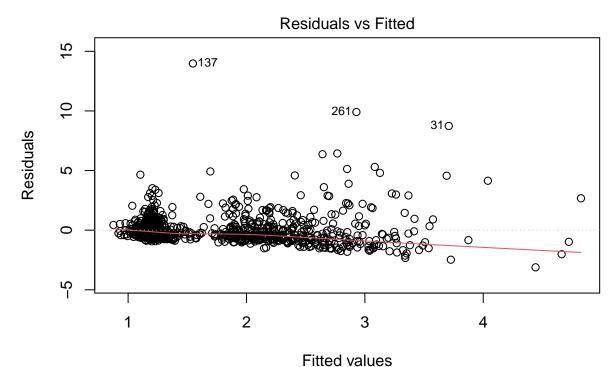
'geom_smooth()' using method = 'gam' and formula = 'y ~ s(x, bs = "cs")'

Residual plot: Residual vs Fitted values



This is a perfect model as there is no patterns, the plot shows no funneling, which means there is no problem with linearity assumption.

```
plot(InteractModelBest, which=1)
```



Im(Creatinine ~ age + Systolic_blood_pressure + Renal_failure + age * Systo ...

This is a perfect model as there is no patterns, the plot shows no funneling, which means there is no problem with linearity assumption.

Test for heteroscedasticity (non constant variance)

H_0: heteroscedasticity is not present (homoscedasticity) H_a: heteroscedasticity is present

library(lmtest)

```
## Warning: package 'lmtest' was built under R version 4.4.2

## Loading required package: zoo

## Warning: package 'zoo' was built under R version 4.4.2

## ## Attaching package: 'zoo'

## The following objects are masked from 'package:base':

## ## as.Date, as.Date.numeric

bptest(InteractModelBest)
```

##

```
## studentized Breusch-Pagan test
##
## data: InteractModelBest
## BP = 49.282, df = 5, p-value = 1.943e-09
```

library(mctest)

We reject the null hypothesis (p-value < 0.05), so we conclude we do have heteroscedasticity

```
nteractModelBest= lm(Creatinine~age+Systolic_blood_pressure+Renal_failure+age*Systolic_blood_pressure+a
imcdiag(InteractModelBest, method="VIF")
##
## Call:
## imcdiag(mod = InteractModelBest, method = "VIF")
##
##
##
   VIF Multicollinearity Diagnostics
##
##
                                    VIF detection
## age
                                42.0464
## Systolic_blood_pressure
                                27.7674
                                                1
## Renal_failure
                                39.4828
```

Multicollinearity may be due to age Systolic_blood_pressure Renal_failure age:Systolic_blood_pressur

1

1

1 --> COLLINEARITY is detected by the test
0 --> COLLINEARITY is not detected by the test

40.6422

age:Systolic_blood_pressure 68.0319

check for normality test

age:Renal_failure

H_0: We have normality H_a: We do not have normality

shapiro.test(residuals(InteractModelBest))

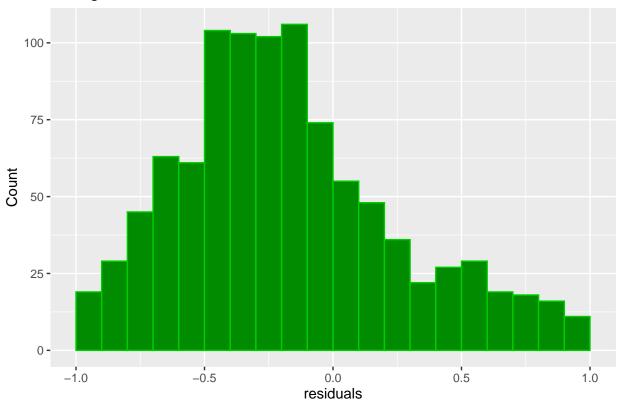
```
##
## Shapiro-Wilk normality test
##
## data: residuals(InteractModelBest)
## W = 0.70846, p-value < 2.2e-16</pre>
```

Shapiro-Wilk normality test confirms that the residuals are not normally distributed as the p- value= 2.2e-16 <0.05 From the Shapiro-Wilk analysis with p-value= 2.2e-16 < 0.05 we reject the null hypothesis

 ${\rm H_0}$: Sample data are significantly normally distributed ${\rm H_a}$: Sample data are not significantly normally distributed

```
ggplot(data=mimicIII_data1, aes(residuals(InteractModelBest))) +
geom_histogram(breaks = seq(-1,1,by=0.1), col="green3", fill="green4") +
labs(title="Histogram for residuals") +
labs(x="residuals", y="Count")
```

Histogram for residuals



p-value =2.2e-16 < 0.05, We reject the null hypothesis, therefore the Sample data are not significantly normally distributed