Problme1

a)

The Least Squares line equation:

$$\hat{Y} = \hat{\beta_0} + \hat{\beta_1} X_i$$

Proof: According to method of Least Squares, $\hat{\beta}_0 = \bar{Y} - \hat{\beta}_1 \bar{X}$, therefore, when $X = \bar{X}$

$$\hat{Y} = \hat{\beta}_0 + \hat{\beta}_1 \bar{X}$$

$$= \bar{Y} - \hat{\beta}_1 \bar{X} + \hat{\beta}_1 \bar{X}$$

$$= \bar{Y}$$

Therefore, it always goes through the point (\bar{X}, \bar{Y}) .

b)

First, there are some useful properties of LS:

$$\sum e_i = 0$$
$$\sum X_i e_i = 0$$
$$\sum \hat{Y}_i e_i = 0$$

Proof: To minimize Q, we have:

$$\frac{\partial Q}{\partial \beta_0} = 2\sum_i (Y_i - \beta_0 - \beta_1 X_i) \cdot (-\beta_0)'$$

$$= -2\sum_i (Y_i - \beta_0 - \beta_1 X_i) = 0$$

$$\frac{\partial Q}{\partial \beta_1} = -2\sum_i (Y_i - \beta_0 - \beta_1 X_i) \cdot X_i = 0$$

Then these two equation imply that $\sum e_i = 0$ and $\sum X_i e_i = 0$

$$\sum_{i} \hat{Y}_{i} e_{i} = \sum_{i} \left(\hat{\beta}_{0} + \hat{\beta}_{1} X_{i} \right) e_{i}$$
$$= \hat{\beta}_{0} \sum_{i} e_{i} + \hat{\beta}_{1} \sum_{i} e_{i} X_{i}$$
$$= 0$$

Therefore,

$$\begin{split} corr(e_i, \hat{Y}_i) &= corr(Y_i - \hat{Y}_i, \hat{Y}_i) \\ &= corr(Y_i, \hat{Y}_i) - corr(\hat{Y}_i, \hat{Y}_i) \\ &= \frac{1}{n} \sum (Y_i - \bar{Y})(\hat{Y}_i - \hat{Y}) - \frac{1}{n} \sum (\hat{Y}_i - \bar{Y})^2 \\ &= \frac{1}{n} \sum (Y_i - \bar{Y})(\hat{Y}_i - \bar{Y}) - \frac{1}{n} \sum (\hat{Y}_i - \bar{Y})^2 \\ &= \frac{1}{n} \sum (Y_i - \bar{Y})(\hat{Y}_i - \bar{Y}) - (\hat{Y}_i - \bar{Y})^2 \\ &= \frac{1}{n} \sum \hat{Y}_i(Y_i - \hat{Y}_i) \\ &= \frac{1}{n} \sum \hat{Y}_i e_i \\ &= 0 \end{split}$$

One of the possible explanations is that the variance of ϵ is not constant.

Problem2

a)

$$Q = (\mathbf{Y} - \mathbf{X}\boldsymbol{\beta})'(\mathbf{Y} - \mathbf{X}\boldsymbol{\beta})$$

= $\mathbf{Y}'\mathbf{Y} - \boldsymbol{\beta}'\mathbf{X}'\mathbf{Y} - \mathbf{Y}'\mathbf{X}\boldsymbol{\beta} + \boldsymbol{\beta}'\mathbf{X}'\mathbf{X}\boldsymbol{\beta}$
= $\mathbf{Y}'\mathbf{Y} - 2\boldsymbol{\beta}'\mathbf{X}'\mathbf{Y} + \boldsymbol{\beta}'\mathbf{X}'\mathbf{X}\boldsymbol{\beta}$

Therefore, let

$$\frac{\partial}{\partial \beta}(Q) = -2\mathbf{X}'\mathbf{Y} + 2\mathbf{X}'\mathbf{X}\boldsymbol{\beta} = 0$$

 $\hat{\boldsymbol{\beta}}_{2\times 1} = (\mathbf{X}'\mathbf{X})^{-1}\mathbf{X}'\mathbf{Y}_{2\times 1}$

then

$$E(\hat{\boldsymbol{\beta}}) = E((\mathbf{X}'\mathbf{X})^{-1}\mathbf{X}'\mathbf{Y})$$
$$= (\mathbf{X}'\mathbf{X})^{-1}\mathbf{X}'E(\mathbf{Y})$$
$$= (\mathbf{X}'\mathbf{X})^{-1}\mathbf{X}'X\beta$$
$$= I\beta$$
$$= \beta$$

b)

$$Var(\tilde{\beta}) = Var\left((X'X)^{-1} X'Y\right)$$

$$= (X'X)^{-1} X' \sigma^{2} IX (X'X)^{-1}$$

$$= \sigma^{2} I (X'X)^{-1} X'X (X'X)^{-1}$$

$$= \sigma^{2} (X'X)^{-1}$$

$$= \sigma^{2} \left[\sum_{i=1}^{n} X_{i} \sum_{i=1}^{n} X_{i} \right]^{-1}$$

$$= \sigma^{2} \left[\frac{1}{n} + \frac{\bar{X}^{2}}{\sum_{i=1}^{n} (X_{i} - \bar{X})^{2}} \frac{-\bar{X}}{\sum_{i=1}^{n} (X_{i} - \bar{X})^{2}} \right]$$

$$= \sigma^{2} \left[\frac{1}{n} + \frac{\bar{X}^{2}}{\sum_{i=1}^{n} (X_{i} - \bar{X})^{2}} \frac{1}{\sum_{i=1}^{n} (X_{i} - \bar{X})^{2}} \right]$$

Problem3

a) Regression model for the nonhuman data

```
lm_non_human <-</pre>
       brain_df %>%
       filter(species != "Homo sapiens") %>%
       lm(glia_neuron_ratio ~ ln_brain_mass, data = .)
summary(lm_non_human)
##
## Call:
## lm(formula = glia_neuron_ratio ~ ln_brain_mass, data = .)
## Residuals:
##
       Min
                 1Q Median
## -0.24150 -0.12030 -0.01787 0.15940 0.25563
##
## Coefficients:
                Estimate Std. Error t value Pr(>|t|)
                 0.16370 0.15987
                                     1.024 0.322093
## (Intercept)
## ln_brain_mass 0.18113
                            0.03604 5.026 0.000151 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.1699 on 15 degrees of freedom
## Multiple R-squared: 0.6274, Adjusted R-squared: 0.6025
## F-statistic: 25.26 on 1 and 15 DF, p-value: 0.0001507
Therefore, the regression equation is:
```

b)

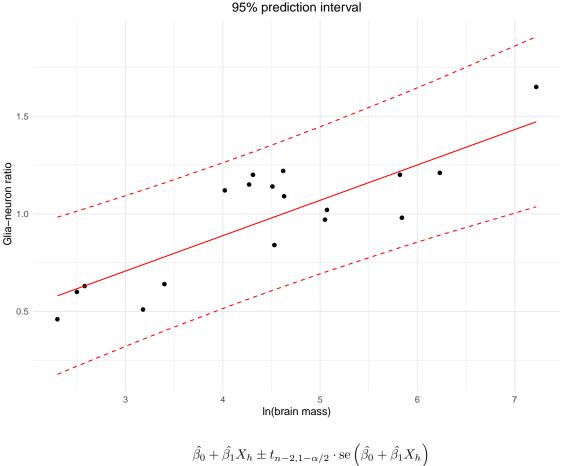
```
\hat{Y}_{human} = 0.164 + 0.181 \ln(\text{brain mass})
= 0.164 + 0.181 \times 7.22
= 1.471
```

Therefore, the predicted glia-neuron ratio for humans is 1.471

c)

The most-plausible range of values for the prediction is (2.30,6.23), which is the range of ln(brain mass) for non-human. An interval for the prediction of a single new observation is more relevant for our prediction of human glia-neuron ratio, because in the case, we want to predict the value of a new observation(human data).

d)



$$\beta_0 + \beta_1 X_h \pm t_{n-2,1-\alpha/2} \cdot \operatorname{se}\left(\beta_0 + \beta_1 X_h\right)$$

$$\operatorname{se}\left(\widehat{\beta_0} + \widehat{\beta_1}X_h\right) = \sqrt{MSE\left\{\frac{1}{n} + \left[\left(X_h - \bar{X}\right)^2 / \sum_{i=1}^n \left(X_i - \bar{X}\right)^2\right] + 1\right\}}$$

Therefore, 95% prediction interval for huamn is (1.04,1.91). As we can see, the huamn data point falls within the prediction interval for its given ln(brain mass), therefore, human brain doesn't have an excessive glia-neuron ratio for its mass compared with other primates

e)

As we can see in the graph, the human data point falls outside the scope of non-human data points, from which we generate the regression line. Therefore, though it's not far away from the scope, we may still need to be cautious in using the regression line to predict.

Problem4

a)

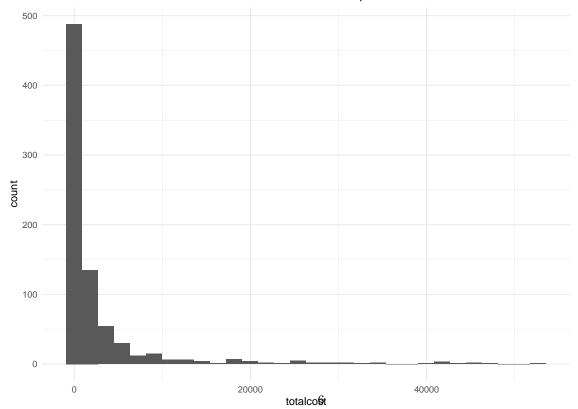
The main outcome is "total cost" (in dollars) of patients diagnosed with heart disease, and the main predictor is the "number of emergency room (ER) visits". Other important covariates include age, gender, number or intervetion carried out, drugs prescribed, complications, comorbidities, and duraion of treatment condition.

Table 1: Characteristics of patients

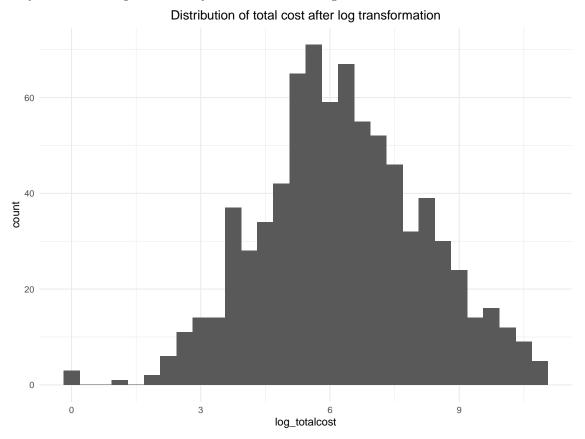
	Overall (N=788)
totalcost	
- Mean (SD)	2799.96 (6690.26)
- Median (Q1, Q3)	507.20 (161.12, 1905.45)
- Min - Max	0.00 - 52664.90
e_rvisits	
- Mean (SD)	3.43 (2.64)
- Median (Q1, Q3)	3.00 (2.00, 5.00)
- Min - Max	0.00 - 20.00
age	
- Mean (SD)	58.72 (6.75)
- Median (Q1, Q3)	60.00 (55.00, 64.00)
- Min - Max	24.00 - 70.00
gender	
- female	608 (77.2%)
- male	180 (22.8%)
complications	
- Mean (SD)	0.06 (0.25)
- Median (Q1, Q3)	0.00 (0.00, 0.00)
- Min - Max	0.00 - 3.00
duration	
- Mean (SD)	164.03 (120.92)
- Median (Q1, Q3)	165.50 (41.75, 281.00)
- Min - Max	0.00 - 372.00
	•

b)

Distribution of total cost of patients



As we can see in the first histgram, the distribution for "total cost" is highly right-skewed. Therefore, we may need to use *Logarithm transformation* to reduce right skewness.



After log tranformation, the distribution for "total cost" is a bell-shaped curve, and the transformed total cost follows a normal distribution.

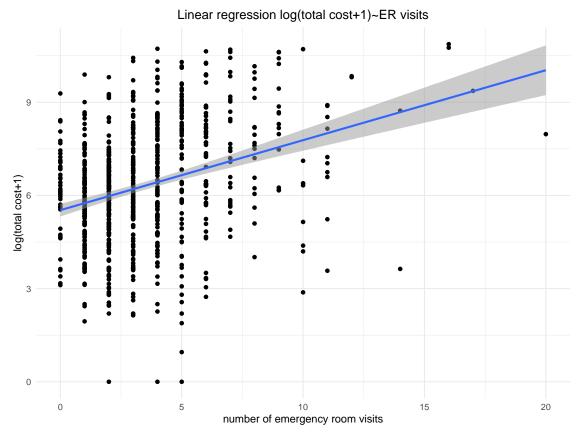
c) Creat "comp_bin" variable.

```
hr_df <-
    hr_df %>%
    mutate(comp_bin = case_when(complications == 0 ~ 0, TRUE ~ 1),
        comp_bin = factor(comp_bin))
```

d) SLR with transformed "total cost" and predictor "ERvisits"

```
##
## Call:
## lm(formula = log_totalcost ~ e_rvisits, data = .)
##
## Residuals:
##
       Min
                1Q Median
                                ЗQ
                                        Max
##
   -6.6532 -1.1230 0.0309 1.2797
                                    4.2964
##
## Coefficients:
               Estimate Std. Error t value Pr(>|t|)
##
```

```
## (Intercept)
                5.52674
                           0.10510
                                    52.584
                                              <2e-16 ***
## e_rvisits
                0.22529
                           0.02432
                                     9.264
                                              <2e-16 ***
##
                     '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Signif. codes:
                   0
##
## Residual standard error: 1.799 on 786 degrees of freedom
## Multiple R-squared: 0.09844,
                                    Adjusted R-squared: 0.09729
## F-statistic: 85.82 on 1 and 786 DF, p-value: < 2.2e-16
```



Therefore, the regression equation is:

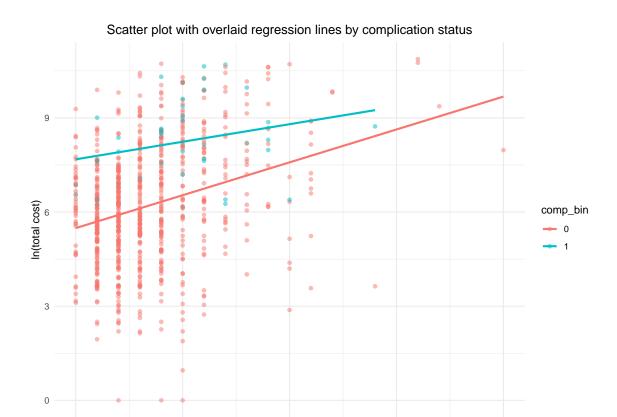
$$\hat{Y} = 5.527 + 0.225X$$

where $\hat{Y} = \ln(\text{total cost} + 1)$, X = number of ER visits

The estimated slope is 0.225 and it's P-value is it's significant, indicates that if the number of emergency room visits increase by 1, the $\log(\text{total cost}+1)$ will increase by 0.225 on avarage. In other word, with the number of emergency room visits increase by 1, the total cost for a patient will increase by 25% on avarage.

e) MLR with "comp_bin" and "ERvisits" as predictors

i) Is "comp_bin" an effect modifier?



As the plot shows, there are non-parallel slopes for different category of "com_bin", therefore, we can add interaction effects to the model to test.

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number of ER visits

```
##
## Call:
## lm(formula = log_totalcost ~ e_rvisits * comp_bin, data = .)
##
## Residuals:
##
     Min
             1Q Median
                           ЗQ
                                 Max
## -6.536 -1.083 0.004 1.200
                               4.398
##
## Coefficients:
                      Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                       5.48849
                                  0.10500 52.271 < 2e-16 ***
## e_rvisits
                       0.20947
                                  0.02490
                                            8.412 < 2e-16 ***
                       2.19096
                                  0.55447
                                            3.951 8.47e-05 ***
## comp_bin1
## e_rvisits:comp_bin1 -0.09753
                                  0.09630 -1.013
                                                     0.311
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 1.759 on 784 degrees of freedom
## Multiple R-squared: 0.1405, Adjusted R-squared: 0.1372
```

Ω

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```
## F-statistic: 42.72 on 3 and 784 DF, p-value: < 2.2e-16
```

As the regression model shows, the interaction term is not significant (P-value 0.311). Therefore, we can conclude that "comp_bin" is not an effect modifier of the relationship between "total cost" and "ERvisits".

ii) Is "comp_bin" a confounder?

```
##
## Call:
## lm(formula = log_totalcost ~ e_rvisits + comp_bin, data = .)
## Residuals:
##
       Min
                10 Median
                                3Q
                                        Max
  -6.5249 -1.0769 -0.0074
                                    4.4024
##
                            1.1847
##
## Coefficients:
##
               Estimate Std. Error t value Pr(>|t|)
## (Intercept)
               5.51020
                           0.10279
                                    53.606 < 2e-16 ***
## e rvisits
                0.20295
                           0.02405
                                     8.437 < 2e-16 ***
## comp_bin1
                1.70573
                           0.27915
                                     6.111 1.56e-09 ***
## ---
## Signif. codes:
                   0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 1.759 on 785 degrees of freedom
## Multiple R-squared: 0.1394, Adjusted R-squared: 0.1372
## F-statistic: 63.57 on 2 and 785 DF, p-value: < 2.2e-16
```

Notice that ER visit remains statistically significantly associated with transformed total cost (p<0.001), but the magnitude of the association is lower after adjustment. (0.203 versus 0.225). The regression coefficient decreases by about 10%.

As rule of thumb, "comp_bin" meets the criteria for confounders. Thus, "comp_bin" is a confounder of the relationship between "total cost" and "ERvisits".

iii) Should "comp_bin" be included along with "ERvisits"?

```
anova(hr_lm1,hr_lm3)
```

```
## Analysis of Variance Table
##
## Model 1: log_totalcost ~ e_rvisits
## Model 2: log_totalcost ~ e_rvisits + comp_bin
## Res.Df RSS Df Sum of Sq F Pr(>F)
## 1 786 2544.8
## 2 785 2429.3 1 115.55 37.339 1.563e-09 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
```

"comp_bin" should be included along with "ERvisits" for two reasons.

First, as mentioned in ii), "comp_bin" is a potential confounder and thus need to be adjusted. Second, I compare these two models using partial ANOVA. Since the P-value < 0.05, at a significane level of 0.05, we reject H_0 , and conclude that the model including "comp_bin" is "superior".

Therefore, "comp_bin" should be included along with "ERvisits".

f)

i) Fit a new MLR

```
hr_1m4 <-
   hr df %>%
   lm(log_totalcost ~ e_rvisits + comp_bin + age + gender + duration, data = .)
summary(hr_lm4)
##
## Call:
## lm(formula = log_totalcost ~ e_rvisits + comp_bin + age + gender +
##
       duration, data = .)
##
## Residuals:
##
      Min
                1Q Median
                                3Q
                                       Max
## -5.4711 -1.0340 -0.1158 0.9493
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) 5.9404610 0.5104064 11.639 < 2e-16 ***
## e_rvisits
               0.1745975
                          0.0225736
                                      7.735 3.20e-14 ***
## comp_bin1
               1.5044946
                          0.2584882
                                       5.820 8.57e-09 ***
                                     -2.380
                                               0.0175 *
               -0.0206475
                           0.0086746
              -0.2067662
                          0.1387002
                                     -1.491
                                               0.1364
## gendermale
## duration
               0.0057150
                          0.0004888 11.691
                                             < 2e-16 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 1.624 on 782 degrees of freedom
## Multiple R-squared: 0.2694, Adjusted R-squared: 0.2647
## F-statistic: 57.68 on 5 and 782 DF, p-value: < 2.2e-16
```

Therefore, the regression equation is:

$$\hat{Y} = 5.940 + 0.175X_1 + 1.504X_2 - 0.021X_3 - 0.207X_4 + 0.006X_5$$

where $\hat{Y} = \ln(\text{total cost} + 1); X1 = \text{number of ER visits}; X2 = \text{having complication}; X3 = \text{age}; X4 = \text{being male}; X5 = \text{duration of treatment}.$

The regression results show that variables of the model, except for gender are all significant associated with of total cost of patients. Holding all other variables constant, if the number of emergency room visits increase by 1, the total cost for a patient will increase by 19% on avarage. Compared with the coefficient from SLR, the coefficient decrease.

ii) Compare the SLR and MLR models.

```
anova(hr_lm1,hr_lm4)
```

```
## Analysis of Variance Table
##
## Model 1: log_totalcost ~ e_rvisits
## Model 2: log_totalcost ~ e_rvisits + comp_bin + age + gender + duration
```

```
## Res.Df RSS Df Sum of Sq F Pr(>F)
## 1 786 2544.8
## 2 782 2062.2 4 482.62 45.753 < 2.2e-16 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.05 '.' 0.1 ' ' 1</pre>
```

I would choose the MLR models to address the investigator's objective. First, by comparing the two models using partial ANOVA, we can reject H_0 and conclude that the MLR model is suprior.

Second, $R^2_{adjusted}$ of the SLR and the MLR model is 0.097 and 0.265 respectively, which indicates that the MLR model can better explain the variation of the dependent variable.