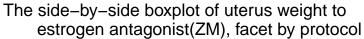
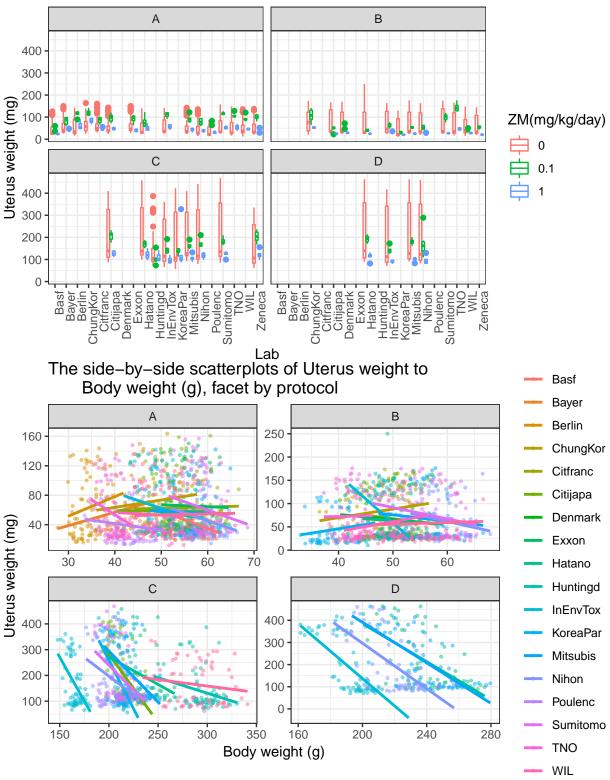
STA721 Final Project

Shuangjie Zhang, Xiyang Hu12/8/2018

- 1. Summary
- 2. Introductions



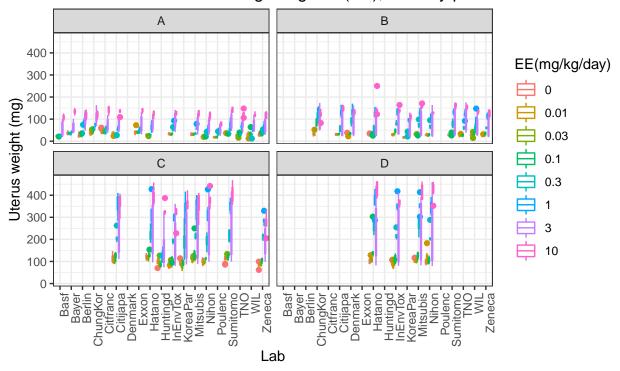


Apeendix

EDA

```
bioassay_lm = bioassay[,-7]
str(bioassay_lm)
## 'data.frame':
                   2677 obs. of 6 variables:
## $ uterus : num 21 22 21 26 24 25 22 26 24 22 ...
## $ weight : num 61.9 55.9 59.1 54.8 57.5 57.6 60.3 59 59.1 61.4 ...
## $ protocol: Factor w/ 4 levels "A", "B", "C", "D": 1 1 1 1 1 1 1 1 1 1 ...
             : Factor w/ 8 levels "0","0.01","0.03",...: 1 1 1 1 1 1 1 1 1 1 1 ...
## $ ZM
              : Factor w/ 3 levels "0", "0.1", "1": 1 1 1 1 1 1 1 1 1 ...
## $ lab
              : Factor w/ 19 levels "Basf", "Bayer", ...: 1 1 1 1 1 1 1 1 1 1 1 ...
table(bioassay_lm$EE, bioassay_lm$ZM)
##
##
           0 0.1
##
                   0
          484
              0
    0.01 234
##
                   0
##
    0.03 239
                   0
##
     0.1 246
     0.3 246 0
##
                   0
##
         246
##
         246 245 246
     3
     10 245
ggplot(data=bioassay,mapping = aes(y = uterus,x = lab,color=EE))+
  geom_boxplot()+theme_bw()+facet_wrap(~ protocol) +
  theme(axis.text.x = element_text(angle = 90, hjust = 1)) +
  labs(x = "Lab", y="Uterus weight (mg)", title="The side-by-side boxplot of uterus weight for differen
       different dose of estrogen agonist(EE), facet by protocol", caption="", colour="EE(mg/kg/day)")
```

The side-by-side boxplot of uterus weight for different labs and different dose of estrogen agonist(EE), facet by protocol



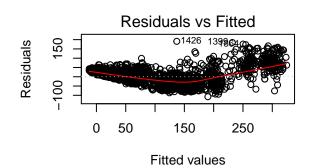
Model Part I

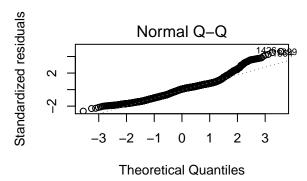
```
lm1 = lm(uterus~., data = bioassay_lm)
summary(lm1)
##
## Call:
## lm(formula = uterus ~ ., data = bioassay_lm)
## Residuals:
##
        Min
                  1Q
                       Median
                                     3Q
                                             Max
                        2.595
  -107.625 -30.150
                                21.979
                                        190.872
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
                             6.55993
                                        2.412 0.015933 *
## (Intercept)
                 15.82251
## weight
                 -0.45365
                             0.05508 -8.237 2.75e-16 ***
## protocolB
                  7.84315
                             2.26416
                                        3.464 0.000541 ***
                207.53588
                             9.48173 21.888
                                              < 2e-16 ***
## protocolC
## protocolD
                221.22623
                            10.07610
                                      21.956
                                               < 2e-16 ***
## EE0.01
                 -0.60177
                             3.31535
                                      -0.182 0.855982
## EE0.03
                  0.26008
                             3.28953
                                        0.079 0.936989
## EE0.1
                             3.25677
                                        2.460 0.013946 *
                  8.01257
## EE0.3
                 47.94479
                             3.25716
                                      14.720
                                               < 2e-16 ***
## EE1
                106.35605
                             3.26542
                                      32.570
                                               < 2e-16 ***
## EE3
                136.45891
                             3.27333
                                      41.688
                                               < 2e-16 ***
## EE10
                             3.28955 45.768 < 2e-16 ***
                150.55730
```

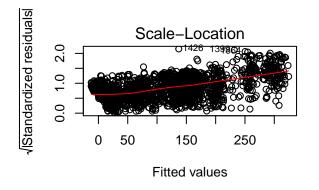
```
## ZMO.1
                -80.51563
                             3.75770 -21.427 < 2e-16 ***
## 7.M1
               -127.18576
                             3.75162 -33.902 < 2e-16 ***
## labBayer
                  2.60266
                             7.88343
                                        0.330 0.741318
## labBerlin
                 14.84134
                             7.55929
                                        1.963 0.049713 *
## labChungKor
                 32.46041
                             6.63073
                                        4.895 1.04e-06 ***
## labCitfranc
                 26.21060
                                        3.518 0.000443 ***
                             7.45111
## labCitijapa
                                        3.399 0.000686 ***
                 21.52689
                             6.33273
## labDenmark
                                        2.430 0.015165 *
                 18.95727
                             7.80137
## labExxon
                 23.72114
                             7.62673
                                        3.110 0.001889 **
## labHatano
                 26.83352
                             6.19974
                                        4.328 1.56e-05 ***
## labHuntingd
                  0.09856
                             8.84741
                                        0.011 0.991112
## labInEnvTox
                             6.38094
                                        0.092 0.927028
                  0.58445
## labKoreaPar
                 -2.51500
                             6.88744
                                      -0.365 0.715023
## labMitsubis
                 24.63683
                             6.19749
                                        3.975 7.22e-05 ***
## labNihon
                             6.20345
                                        2.126 0.033590 *
                 13.18893
## labPoulenc
                 -4.14169
                             7.49225
                                       -0.553 0.580450
## labSumitomo
                             6.32781
                                        4.508 6.83e-06 ***
                 28.52520
## labTNO
                 16.56429
                             6.68045
                                        2.480 0.013218 *
## labWIL
                 10.05022
                             6.63237
                                        1.515 0.129809
## labZeneca
                 17.93047
                             6.42998
                                        2.789 0.005332 **
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 41.56 on 2645 degrees of freedom
## Multiple R-squared: 0.802, Adjusted R-squared: 0.7997
## F-statistic: 345.7 on 31 and 2645 DF, p-value: < 2.2e-16
step(lm1, k=log(2677))
## Start: AIC=20175.57
## uterus ~ weight + protocol + EE + ZM + lab
##
##
              Df Sum of Sq
                                RSS
                                       AIC
## <none>
                            4568714 20176
## - lab
              18
                    304839
                            4873553 20206
## - weight
               1
                    117187
                            4685901 20236
## - protocol
               3
                    855660
                            5424374 20612
## - ZM
               2
                   2030817
                            6599531 21144
## - EE
                   7683826 12252540 22761
##
## Call:
## lm(formula = uterus ~ weight + protocol + EE + ZM + lab, data = bioassay_lm)
## Coefficients:
                                             protocolC
##
  (Intercept)
                     weight
                               protocolB
                                                          protocolD
##
      15.82251
                   -0.45365
                                 7.84315
                                             207.53588
                                                          221.22623
##
        EE0.01
                     EE0.03
                                    EE0.1
                                                 EE0.3
                                                                 EE1
##
      -0.60177
                    0.26008
                                  8.01257
                                              47.94479
                                                          106.35605
##
           EE3
                       EE10
                                    ZMO.1
                                                   ZM1
                                                           labBayer
##
                                            -127.18576
                                                            2.60266
     136.45891
                  150.55730
                               -80.51563
                             labCitfranc
##
     labBerlin labChungKor
                                           labCitijapa
                                                         labDenmark
##
      14.84134
                   32.46041
                                26.21060
                                              21.52689
                                                           18.95727
##
      labExxon
                  labHatano
                             labHuntingd
                                           labInEnvTox
                                                        labKoreaPar
##
      23.72114
                   26.83352
                                  0.09856
                                               0.58445
                                                           -2.51500
```

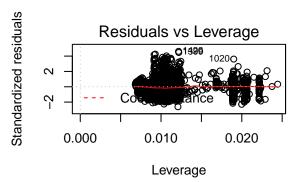
```
## labMitsubis
                    labNihon
                               labPoulenc labSumitomo
                                                               labTNO
                    13.18893
                                               28.52520
##
      24.63683
                                 -4.14169
                                                             16.56429
                   labZeneca
##
        labWIL
##
      10.05022
                    17.93047
library(MASS)
##
## Attaching package: 'MASS'
## The following object is masked from 'package:dplyr':
##
##
       select
box =boxcox(lm1)
     0006-
log-Likelihood
     -11000
     -13000
            -2
                              -1
                                                0
                                                                  1
                                                                                    2
                                                λ
lm2 = lm(formula = log(uterus) ~ log(weight) + protocol + EE + ZM + lab, data = bioassay_lm)
summary(lm2)
##
## Call:
## lm(formula = log(uterus) ~ log(weight) + protocol + EE + ZM +
##
       lab, data = bioassay_lm)
##
## Residuals:
                   1Q
                        Median
                                              Max
## -1.38682 -0.16223 0.01173 0.16583 1.21149
##
## Coefficients:
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) 1.321144
                            0.239602
                                      5.514 3.85e-08 ***
## log(weight) 0.410375
                            0.058372
                                        7.030 2.61e-12 ***
## protocolB
                            0.014890
                                        3.694 0.000225 ***
                0.055004
## protocolC
                0.689389
                            0.083618
                                      8.244 2.58e-16 ***
```

```
## protocolD
               0.675012
                          0.085999
                                     7.849 6.03e-15 ***
## EE0.01
               0.002747
                          0.021776
                                   0.126 0.899614
## EE0.03
               0.012017
                          0.021606
                                    0.556 0.578144
## EEO.1
               0.131728
                          0.021388
                                     6.159 8.44e-10 ***
## EE0.3
               0.551201
                          0.021396
                                   25.762 < 2e-16 ***
## EE1
                          0.021438 49.527
               1.061739
                                           < 2e-16 ***
## EE3
                          0.021477 62.954 < 2e-16 ***
               1.352063
                          0.021571 67.294
## EE10
               1.451617
                                           < 2e-16 ***
                          0.024670 -21.541
## ZMO.1
              -0.531428
                                           < 2e-16 ***
                          0.024622 -46.504 < 2e-16 ***
## ZM1
              -1.145005
## labBayer
               0.457789
                          0.055245
                                     8.286 < 2e-16 ***
## labBerlin
                          0.057515
                                   10.621 < 2e-16 ***
               0.610866
## labChungKor 0.762522
                          0.044567
                                    17.109 < 2e-16 ***
                                   11.425 < 2e-16 ***
## labCitfranc 0.559241
                          0.048950
                          0.041829
                                   10.647
                                           < 2e-16 ***
## labCitijapa 0.445367
## labDenmark
               0.506016
                          0.051920
                                     9.746
                                            < 2e-16 ***
## labExxon
                          0.050097 10.338 < 2e-16 ***
               0.517897
## labHatano
               0.383445
                          0.040728
                                    9.415 < 2e-16 ***
## labHuntingd -0.053061
                          0.053724
                                   -0.988 0.323416
## labInEnvTox 0.455932
                          0.042501 10.728 < 2e-16 ***
## labKoreaPar 0.178793
                          0.046846
                                    3.817 0.000138 ***
## labMitsubis 0.392970
                          0.040688
                                    9.658 < 2e-16 ***
## labNihon
                                    7.545 6.19e-14 ***
               0.307153
                          0.040710
## labPoulenc
               0.097765
                          0.051708
                                    1.891 0.058768 .
## labSumitomo 0.464850
                          0.041551 11.187 < 2e-16 ***
## labTNO
               0.533952
                          0.047525 11.235 < 2e-16 ***
## labWIL
               0.380265
                          0.044646
                                    8.517 < 2e-16 ***
## labZeneca
               0.251125
                          0.041670
                                     6.027 1.91e-09 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.273 on 2645 degrees of freedom
## Multiple R-squared: 0.8937, Adjusted R-squared: 0.8925
## F-statistic: 717.6 on 31 and 2645 DF, p-value: < 2.2e-16
par(mfrow=c(2,2))
plot(lm1)
```









Frequentist Random Effect Model:

```
library(lme4)
```

```
## Loading required package: Matrix
randomeffect = lmer(log(uterus) ~ log(weight) + protocol + EE + ZM + (1+EE+ZM lab), data = bioassay_lm)
## Warning in commonArgs(par, fn, control, environment()): maxfun < 10 *</pre>
## length(par)^2 is not recommended.
## Warning in optwrap(optimizer, devfun, getStart(start, rho$lower, rho$pp), :
## convergence code 1 from bobyqa: bobyqa -- maximum number of function
## evaluations exceeded
## singular fit
summary(randomeffect)
## Linear mixed model fit by REML ['lmerMod']
## Formula: log(uterus) ~ log(weight) + protocol + EE + ZM + (1 + EE + ZM |
##
       lab)
##
      Data: bioassay_lm
##
## REML criterion at convergence: 74.5
##
## Scaled residuals:
##
       Min
                1Q Median
                                3Q
                                        Max
## -5.4143 -0.5782 0.0019 0.5717 5.4534
##
## Random effects:
   Groups
            Name
                         Variance Std.Dev. Corr
```

```
##
   lab
             (Intercept) 0.038505 0.19623
##
             EE0.01
                         0.002410 0.04909 -0.13
                         0.003451 0.05874
                                           0.52 - 0.11
##
             EE0.03
##
             EE0.1
                         0.015031 0.12260
                                            0.39 -0.89 0.55
##
             EE0.3
                         0.094878 0.30802 -0.21 -0.76 -0.02 0.62
##
             EE1
                         0.170539 0.41296 -0.34 -0.58 -0.11 0.41
                                                                    0.94
##
             EE3
                         0.045229 0.21267
                                          -0.45 -0.26 -0.12 0.14
                         0.015601 0.12491 -0.62 0.27 -0.33 -0.41 0.35
##
             EE10
##
             ZMO.1
                         0.108467 0.32934
                                            0.34 0.34 0.88 0.11 -0.28
##
                         0.058896 0.24268
                                            0.35 0.25 0.80 0.15 -0.52
             ZM1
##
   Residual
                         0.053099 0.23043
##
##
##
##
##
##
##
##
    0.94
##
    0.62 0.84
##
   -0.30 -0.16 -0.13
  -0.53 -0.44 -0.40 0.80
##
## Number of obs: 2677, groups: lab, 19
##
## Fixed effects:
##
               Estimate Std. Error t value
## (Intercept) 1.62766
                           0.19575
                                     8.315
                                     8.998
## log(weight) 0.43491
                           0.04834
## protocolB
                                     4.478
                0.05597
                           0.01250
## protocolC
                0.65373
                           0.06946
                                     9.412
## protocolD
                0.64037
                           0.07140
                                     8.969
## EE0.01
                0.02789
                           0.02172
                                     1.284
## EE0.03
                0.02528
                           0.02277
                                     1.110
## EEO.1
                0.11530
                           0.03376
                                     3.416
## EE0.3
                0.46623
                           0.07330
                                     6.361
## EE1
                0.95558
                           0.09684
                                     9.868
## EE3
                1.30993
                           0.05233
                                    25.033
## EE10
                1.43968
                           0.03423
                                    42.062
## ZMO.1
               -0.45946
                           0.07888 -5.825
## ZM1
               -1.07974
                           0.05999 -17.999
##
## Correlation matrix not shown by default, as p = 14 > 12.
## Use print(x, correlation=TRUE) or
##
       vcov(x)
                      if you need it
## convergence code: 1
## singular fit
## maxfun < 10 * length(par)^2 is not recommended.
```

Model Part II

Model Part III