Modeling Ordinal Data: Model Building Hands On

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Preparation for Analysis

```
rm(list=ls())
```

Libraries

```
library(MASS)  #' polr() function , stepAIC() function
library(ordinal)  #' clm() function
library(dplyr)  #' data transformation functions
library(boot)  #' logit(), inv.logit() functions
library(ggplot2)  #' plotting functions
library(GGally)  #' scatterplot matrices
```

Functions

Data

```
wd <- "C:/..."; setwd(wd)
getwd()</pre>
```

[1] "C:/Program Files (x86)/ePharm/ePharmRunWorkspace/AnalysisStep726570"

```
dsn <- "AEs-ordinal-data-ver3.csv"
d <- read.csv(dsn, as.is=T)</pre>
d$AEGrade <- as.factor(d$DV)</pre>
d$Sex <- factor(d$SEX,levels=c(1,2), labels=c("Male","Female"))</pre>
d$Smoker <- factor(d$SMOK,levels=c(1,2,3), labels=c("Current", "Former", "Never"))
d$Dose <- as.factor(d$DOSE)</pre>
head(d)
     ID DOSE
                   BIO SEX AGE
                                   WT SMOK
                                              CMAXSS
                                                         CAVGSS
                                                                     BIO2 DV
## 1
     1
          50 20.09565
                         1
                            64 76.66
                                         3 13.215195 13.378387 55.16573
          30 25.37029
                            69 73.50
## 2
                         2
                                         1 15.867221 12.778449 52.19035
                            54 76.00
## 3
      3
          10 15.98545
                         2
                                           6.620023 6.490101 58.65953
## 4
     4
         100 13.36886
                         2
                            50 65.90
                                         1 34.478728 16.450041 37.92031
                                                                           1
     5
## 5
          50 10.21455
                            63 67.00
                                         3 18.182994 11.171032 38.19847
                         1
     6 100 28.84815
                         2 73 74.00
                                         1 26.420695 18.208066 53.85277
##
     AEGrade
                Sex Smoker Dose
## 1
           2
               Male
                       Never
                               50
## 2
           1 Female Current
                               30
## 3
           1 Female Current
                               10
## 4
           1 Female Current
                              100
## 5
               Male
                       Never
                               50
## 6
           O Female Current
summary(d)
```

```
DOSE
                                                               SEX
##
          ID
                                            BIO
                            : 0.0
                                              : 0.9467
                                                                 :1.000
##
    Min.
           : 1.00
                      Min.
                                       Min.
                                                          Min.
    1st Qu.: 50.75
                      1st Qu.: 10.0
                                       1st Qu.:12.0209
                                                          1st Qu.:2.000
##
    Median :100.50
                      Median: 30.0
                                       Median :16.1634
                                                          Median :2.000
           :100.50
                            : 40.9
                                              :16.4975
##
    Mean
                      Mean
                                       Mean
                                                          Mean
                                                                 :1.785
                      3rd Qu.: 50.0
                                       3rd Qu.:20.8966
                                                          3rd Qu.:2.000
##
    3rd Qu.:150.25
    Max.
           :200.00
                             :100.0
                                       Max.
                                              :33.1072
                                                          Max.
                                                                 :2.000
                      Max.
                                            SMOK
##
         AGE
                           WT
                                                            CMAXSS
##
    Min.
           :19.00
                     Min.
                            : 41.50
                                       Min.
                                              :1.000
                                                       Min.
                                                               : 0.000
##
    1st Qu.:48.00
                     1st Qu.: 62.00
                                       1st Qu.:1.000
                                                        1st Qu.: 7.124
    Median :56.00
                     Median: 71.35
                                       Median :1.000
                                                       Median: 14.991
                           : 73.26
                                                               :15.096
##
    Mean
          :53.65
                     Mean
                                       Mean
                                              :1.375
                                                        Mean
##
    3rd Qu.:61.00
                     3rd Qu.: 82.12
                                       3rd Qu.:2.000
                                                        3rd Qu.:20.696
##
    Max.
           :73.00
                     Max.
                            :128.40
                                       Max.
                                              :3.000
                                                        Max.
                                                               :63.131
##
        CAVGSS
                           BIO2
                                              DV
                                                        AEGrade
                                                                    Sex
                                                        0:29
##
    Min.
           : 0.000
                      Min.
                             : 9.199
                                        Min.
                                               :0.00
                                                                Male : 43
##
    1st Qu.: 4.748
                      1st Qu.:30.721
                                        1st Qu.:1.00
                                                        1:57
                                                                Female:157
    Median :10.334
                      Median: 39.101
                                        Median:2.00
                                                        2:78
          : 9.347
                                               :1.71
##
    Mean
                      Mean
                             :40.725
                                        Mean
                                                        3:15
##
    3rd Qu.:12.790
                      3rd Qu.:50.189
                                        3rd Qu.:2.00
                                                        4:21
##
    Max.
           :21.403
                             :76.248
                                               :4.00
                      Max.
                                        Max.
##
        Smoker
                    Dose
##
    Current:148
                  Ω
                      :39
##
    Former: 29
                   10:31
##
    Never: 23
                  30:39
##
                  50:48
##
                   100:43
```

Question 1:

Create at least 1 graph (a) to explore relationships between covariates and (b) between covariates and the response.

Observed Covariate-to-Covariate Relationships

Continuous Variables

ggsave("Cov_Cov_02.png",gg,height=9,width=12)

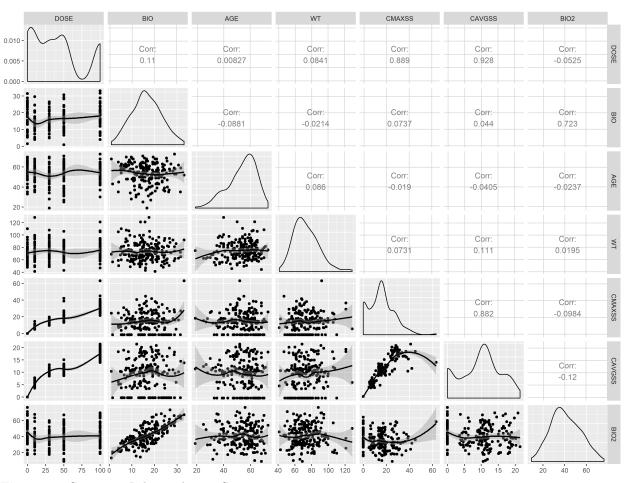


Figure 1. Covariate Relationships - Continuous.

Catergorcal

```
gg <- ggpairs(data=d,columns=c(3,5:6,9,10,13), title = "By Sex",
               upper = list(continuous = "cor"),
               diag = list(continuous= "density"),
               lower = list(continuous = "smooth_loess", mapping=aes(colour = Sex)),
               axisLabels="show")
## Warning in check_and_set_ggpairs_defaults("diag", diag, continuous =
## "densityDiag", : Changing diag$continuous from 'density' to 'densityDiag'
ggsave("Cov_Cov_10.png",gg,height=9,width=12)
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
   By Sex
0.06
0.04 -
                        Corr:
                                        Corr:
                                                        Corr:
                        -0.0881
                                        -0.0214
                                                        0.044
0.02
                                        Corr:
                                                        Corr:
                                                                       Corr:
                                        0.086
                                                       -0.0405
                                                                       -0.0237
                                                        Corr:
                                                                        Corr:
                                                        0.111
                                                                       0.0195
                                                                        Corr:
 50 -
 0 -
 10-
 0 -
 10 -
                                              120
```

Figure 2. Covariate Relationships - By Sex.

```
gg <- ggpairs(data=d,columns=c(3,5:6,9,10,14), title = "By Smoker",
               upper = list(continuous = "cor"),
               diag = list(continuous= "density"),
               lower = list(continuous = "smooth_loess", mapping=aes(colour = Smoker)),
               axisLabels="show")
## Warning in check_and_set_ggpairs_defaults("diag", diag, continuous =
## "densityDiag", : Changing diag$continuous from 'density' to 'densityDiag'
ggsave("Cov_Cov_12.png",gg,height=9,width=12)
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
## `stat_bin()` using `bins = 30`. Pick better value with `binwidth`.
   By Smoker
                                                       CAVGSS
                                                                       BIO2
0.06
0.04
                        Corr
                                        Corr:
                                                       Corr
                                                                       Corr
                        -0.0881
                                       -0 0214
                                                       0.044
                                                                       0.723
0.00
                                        Corr:
                                                       Corr:
                                                       -0.0405
 40
 20
                                                       Corr:
                                                       0.111
                                                                      0.0195
                                                                       Corr
                                                                       -0.12
 -10 -
 80 -
 60 -
```

Figure 3. Covariate Relationships - By Smoker.

80 Current Former

Observed Covariate-to-Response Relationships

Continuous

Frequency & cumulative sum by DOSE

```
d1 <- d %>% group_by(DOSE,DV) %>% summarize(N.AEGrade=nobs_2(DV)) %>%
            mutate(cFreq = cumsum(N.AEGrade)) %>%
            group_by(DOSE) %>% mutate(N.Tot = sum_2(N.AEGrade))
d1$AEGrade <- as.factor(d1$DV)</pre>
d1$p <- d1$N.AEGrade/d1$N.Tot</pre>
d1$cp <- d1$cFreq/d1$N.Tot
d1$Dose <- as.factor(d1$DOSE)</pre>
gg <- ggplot(data=d1, aes(x=AEGrade, y=cp, colour=Dose, group=Dose)) +</pre>
 geom_point() +
  geom_line() +
  theme(axis.text.x = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.text.y = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.title.x = element_text(size=20),
        axis.title.y = element_text(size=20),
        legend.position ="bottom",
        legend.title=element_text(colour="black", size=18, face="bold"),
        legend.text = element_text(colour="black", size=18, face="bold")) +
  xlab("AE Grade") +
  ylab("Cumulative Probability")
ggsave("Cov_Resp_1_02.png",gg,height=9,width=12)
```

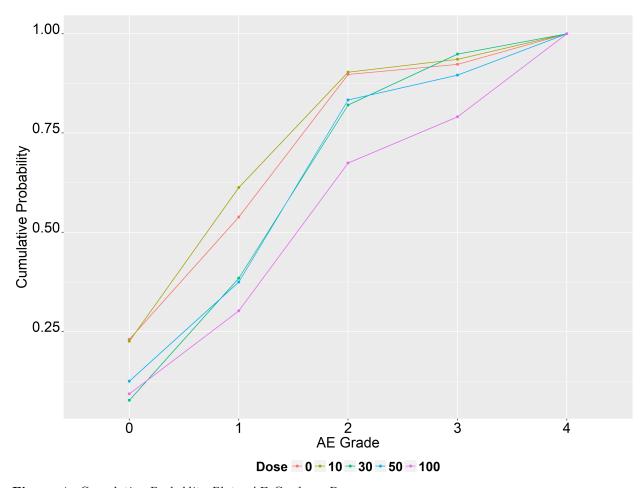


Figure 4. Cumulative Probablity Plot - AE Grade vs Dose.

```
gg <- ggplot(data=d, aes(x=AEGrade, y=DOSE, colour=AEGrade)) +
    geom_boxplot(fill="white", alpha=0.5) +
    # geom_point(pch=0, size=4) +
    geom_jitter(pch=1, size=5, width = 0.05) +
    theme(axis.text.x = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.text.y = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.title.x = element_text(size=20),
        axis.title.y = element_text(size=20),
        legend.position ="none") +
    xlab("AEGrade") +
    ylab("Dose (mg)")
    ggsave("Cov_Resp_1_04.png",gg,height=9,width=12)</pre>
```

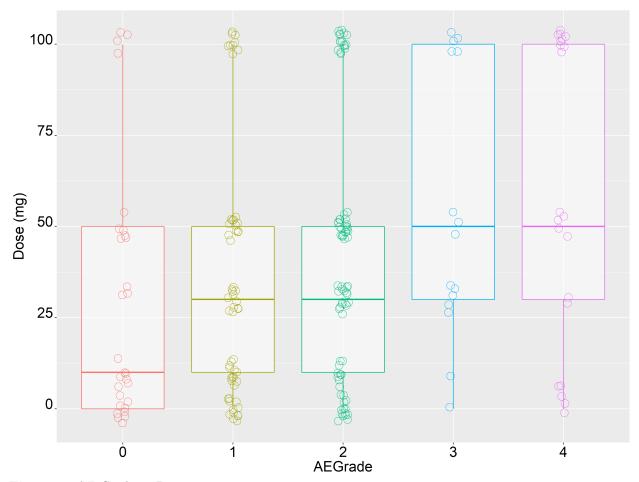


Figure 5. AE Grade vs Dose.

Frequency & cumulative sum by AGE

```
d2 <- d %>% mutate(q20 = quantile(AGE, prob=0.2),
                   q40 = quantile(AGE, prob=0.4),
                   q60 = quantile(AGE, prob=0.6),
                   q80 = quantile(AGE, prob=0.8))
d2$AGEgn <- 1
d2\$AGEgn[d2\$AGE>d2\$q20 & d2\$AGE<=d2\$q40] <- 2
d2$AGEgn[d2$AGE>d2$q40 & d2$AGE<=d2$q60] <- 3
d2$AGEgn[d2$AGE>d2$q60 & d2$AGE<=d2$q80] <- 4
d2$AGEgn[d2$AGE>d2$q8] <- 5
d2 <- d2 %>% group_by(AGEgn) %>% mutate(AGEgmin = min_2(AGE), AGEgm = median_2(AGE),
                                         AGEgmax = max_2(AGE)
d2$AGE_Group <- paste(d2$AGEgmin,d2$AGEgmax,sep=" - ")</pre>
d3 <- d2 %>% group_by(AGE_Group,DV) %>% summarize(N.AEGrade=nobs_2(DV)) %>%
             mutate(cFreq = cumsum(N.AEGrade)) %>%
             group_by(AGE_Group) %>% mutate(N.Tot = sum_2(N.AEGrade))
d3$AEGrade <- as.factor(d3$DV)</pre>
d3$p <- d3$N.AEGrade/d3$N.Tot
```

```
d3$cp <- d3$cFreq/d3$N.Tot
d3$Age <- as.factor(d3$AGE_Group)

gg <- ggplot(data=d3, aes(x=AEGrade, y=cp, colour=Age, group=Age)) +
    geom_point() +
    geom_line() +
    theme(axis.text.x = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.text.y = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.title.x = element_text(size=20),
        axis.title.y = element_text(size=20),
        legend.position ="bottom",
        legend.title=element_text(colour="black", size=18, face="bold"),
        legend.text = element_text(colour="black", size=18, face="bold")) +
        xlab("AE Grade") +
        ylab("Cumulative Probability")
ggsave("Cov_Resp_1_08.png",gg,height=9,width=12)</pre>
```

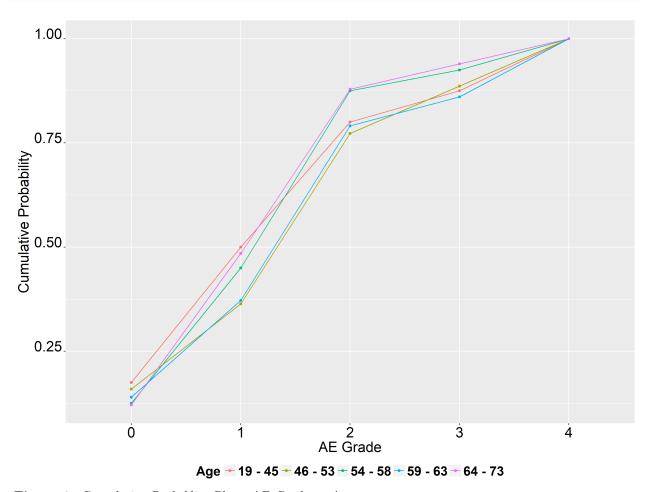


Figure 6. Cumulative Probablity Plot - AE Grade vs Age.

```
gg <- ggplot(data=d, aes(x=AEGrade, y=AGE, colour=AEGrade)) +
  geom_boxplot(fill="white", alpha=0.5) +
  # geom_point(pch=0, size=4) +
  geom_jitter(pch=1, size=5, width = 0.05) +
  theme(axis.text.x = element_text(angle=0, colour="black", vjust=0, size=20),</pre>
```

```
axis.text.y = element_text(angle=0, colour="black", vjust=0, size=20),
    axis.title.x = element_text(size=20),
    axis.title.y = element_text(size=20),
    legend.position ="nonem") +
    xlab("AEGrade") +
    ylab("Age (yrs)")
ggsave("Cov_Resp_1_10.png",gg,height=9,width=12)
```

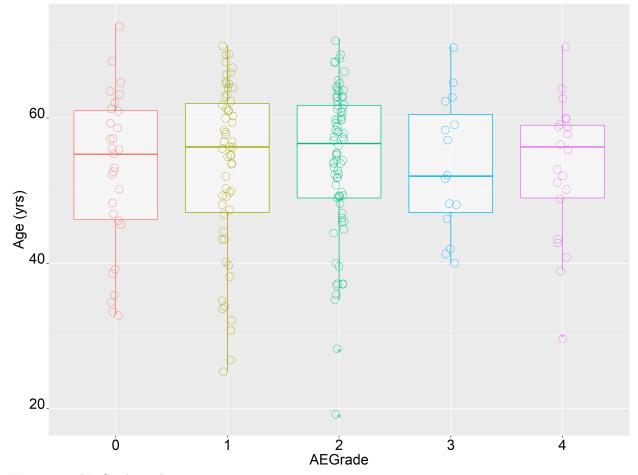


Figure 7. AE Grade vs Age.

Categorical

Frequency & cumulative sum by SEX

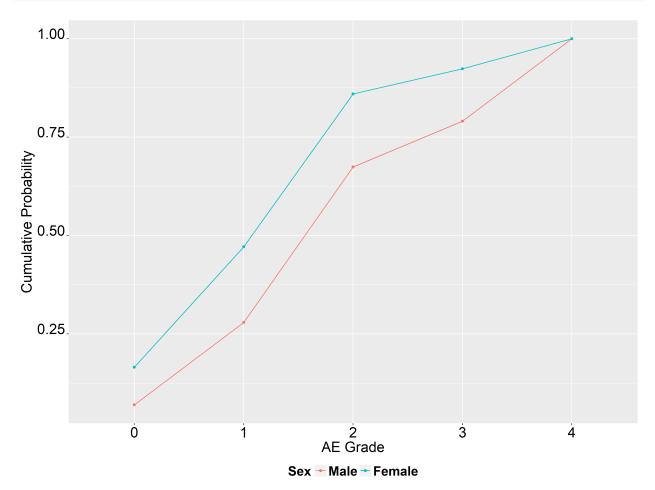


Figure 8. Cumulative Probablity Plot - AE Grade vs Sex.

```
gg <- ggplot(data=d, aes(x=Sex, y=DV, colour=AEGrade)) +
   geom_point() +
   # geom_point(pch=0, size=4) +
   geom_jitter(pch=1, size=5, width = 0.05) +
   theme(axis.text.x = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.text.y = element_text(angle=0, colour="black", vjust=0, size=20),
        axis.title.x = element_text(size=20),
        axis.title.y = element_text(size=20),</pre>
```

```
legend.position ="none") +
xlab("Sex") +
ylab("AE Grade")
ggsave("Cov_Resp_2_03.png",gg,height=9,width=12)
```

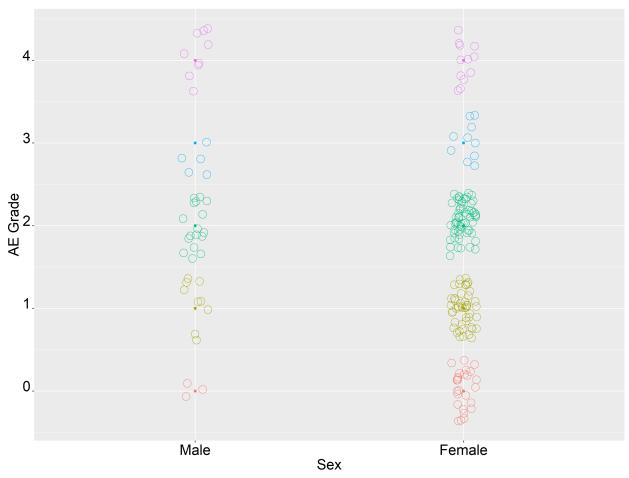


Figure 9. AE Grade vs Sex.

Frequency & cumulative sum by SMOKER

```
axis.title.x = element_text(size=20),
    axis.title.y = element_text(size=20),
    legend.position ="bottom",
    legend.title=element_text(colour="black", size=18, face="bold"),
    legend.text = element_text(colour="black", size=18, face="bold")) +
    xlab("AE Grade") +
    ylab("Cumulative Probability")
ggsave("Cov_Resp_2_04.png",gg,height=9,width=12)
```

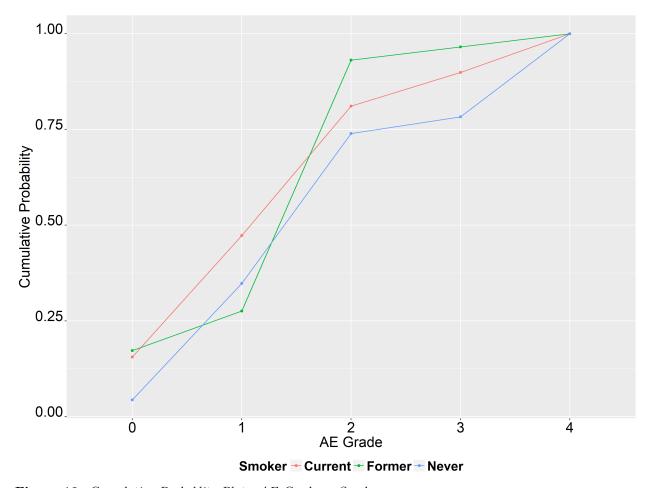


Figure 10. Cumulative Probablity Plot - AE Grade vs Smoker.

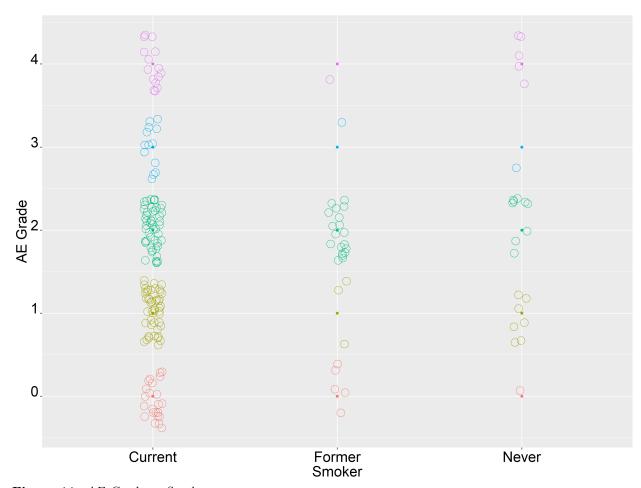


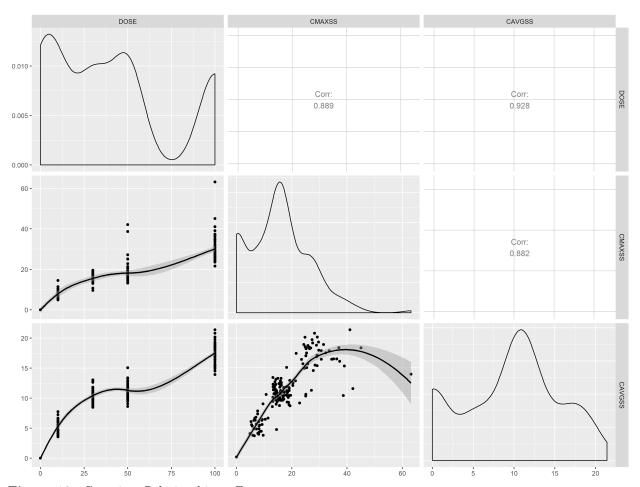
Figure 11. AE Grade vs Smoker.

Question 2:

Do a scatter plot for the 3 exposure measures. Which measure would you choose for further covariate model building?

```
## Warning in check_and_set_ggpairs_defaults("diag", diag, continuous =
## "densityDiag", : Changing diag$continuous from 'density' to 'densityDiag'
```





 ${\bf Figure~12.~} {\it Covariate~Relationships~-Exposure.}$

Question 3

With CMAXSS as the exposure measure in the base model, use stepAIC() to do a complete forward selection followed by a complete backward selection to select a final model.

Models

 ${\bf Exposure} = {\bf CMAXSS}$

d\$EXP <- d\$CMAXSS

Null Model

```
mod0 <- clm(AEGrade ~ 1 , data = d, link = "logit", threshold="flex", Hess=T)
summary(mod0, digits = 3)</pre>
```

```
## formula: AEGrade ~ 1
## data:
           d
##
## link threshold nobs logLik AIC niter max.grad cond.H
## logit flexible 200 -287.18 582.36 5(0) 2.88e-08 8.7e+00
##
## Threshold coefficients:
      Estimate Std. Error z value
##
## 0|1 -1.7744
                 0.2008 -8.835
## 1|2 -0.2819
                   0.1428 -1.973
## 2|3 1.5163
                   0.1841 8.239
## 3|4
       2.1429
                   0.2307 9.290
inv.logit(mod0$coef[1:4])
## 0|1 1|2 2|3
                      3|4
## 0.145 0.430 0.820 0.895
Base Model
mod1 <- clm(AEGrade~ EXP , data = d, link = "logit", threshold="flex", Hess=T)</pre>
summary(mod1, digits = 3)
## formula: AEGrade ~ EXP
## data:
           d
##
## link threshold nobs logLik AIC niter max.grad cond.H
## logit flexible 200 -277.92 565.85 5(0) 4.73e-07 4.9e+03
##
## Coefficients:
      Estimate Std. Error z value Pr(>|z|)
##
## EXP 0.05240 0.01239 4.229 2.34e-05 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Threshold coefficients:
      Estimate Std. Error z value
## 0|1 -1.0934
                 0.2537 -4.311
## 1|2
       0.4735
                   0.2289
                           2.069
                           8.291
## 2|3
       2.3994
                   0.2894
## 3|4
       3.0612
                   0.3293
                           9.295
exp(mod1$beta)
##
       EXP
## 1.053793
inv.logit(mod1$coef[1:4])
        0 | 1
                  1|2
                           2|3
## 0.2509704 0.6162227 0.9167782 0.9552623
```

Full Model

AEGrade ~ EXP

```
mod2 <- clm(AEGrade ~ EXP + BIO + SEX + AGE + WT + SMOK + BIO2 , data = d,
           link = "logit", threshold="flex", Hess=T)
summary(mod2)
## formula: AEGrade ~ EXP + BIO + SEX + AGE + WT + SMOK + BIO2
## data:
##
## link threshold nobs logLik AIC
                                    niter max.grad cond.H
## logit flexible 200 -265.75 553.49 6(0) 1.53e-12 5.6e+06
##
## Coefficients:
         Estimate Std. Error z value Pr(>|z|)
       ## EXP
## BIO -0.0515692 0.0302779 -1.703
                                    0.0885 .
       -0.7612586 0.3661127 -2.079
## SEX
                                     0.0376 *
## AGE -0.0007764 0.0127251 -0.061
                                     0.9514
## WT
        0.0133934 0.0098390 1.361
                                     0.1734
## SMOK 0.1082145 0.2027697 0.534
                                     0.5936
## BIO2 -0.0108457 0.0141631 -0.766
                                    0.4438
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Threshold coefficients:
##
      Estimate Std. Error z value
## 0|1
       -2.767
                   1.494 - 1.852
## 1|2
       -1.093
                   1.485 -0.736
## 2|3
        1.012
                   1.484 0.682
## 3|4
        1.719
                   1.492 1.152
exp(mod2$beta)
##
        EXP
                 BIO
                           SEX
                                    AGE
                                               WT
                                                       SMOK
                                                                BI02
## 1.0559286 0.9497379 0.4670782 0.9992239 1.0134835 1.1142867 0.9892129
inv.logit(mod2$coef[1:4])
                                        3|4
         0 | 1
                   1 | 2
                              2|3
## 0.05913922 0.25098031 0.73339518 0.84794492
Forward & Backward Selection Separately
Forward Selection: Start with Base model
stepAIC(mod1, scope=list(lower=~EXP, upper=~EXP + Sex + AGE + WT + Smoker + BIO + BIO2),
       direction="forward", trace=1, k=2)
## Start: AIC=565.85
```

```
##
##
           Df
                 AIC
## + BIO
           1 555.35
## + Sex
            1 557.44
## + BIO2
            1 559.39
## + WT
            1 561.29
## <none>
              565.85
## + Smoker 2 567.13
## + AGE
            1 567.36
##
## Step: AIC=555.35
## AEGrade ~ EXP + BIO
##
           Df
                 AIC
## + Sex
           1 548.10
## + WT
            1 551.52
## <none>
              555.35
## + BIO2
            1 557.12
## + AGE
            1 557.22
## + Smoker 2 557.39
##
## Step: AIC=548.1
## AEGrade ~ EXP + BIO + Sex
##
                 AIC
           Df
## <none>
            548.10
## + WT
            1 548.34
## + BIO2
            1 549.60
## + AGE
           1 550.10
## + Smoker 2 551.50
## formula: AEGrade ~ EXP + BIO + Sex
## data:
           d
##
## link threshold nobs logLik AIC niter max.grad cond.H
## logit flexible 200 -267.05 548.10 6(0) 2.95e-13 3.0e+04
##
## Coefficients:
##
        EXP
                  BIO SexFemale
##
     0.0570
             -0.0699 -0.9756
##
## Threshold coefficients:
##
      0|1
             1 | 2
                      2|3
                              314
## -3.0663 -1.4059 0.6785 1.3793
Model from forward selection
mod3 <- clm(AEGrade ~ EXP + BIO + Sex, data = d, link = "logit", threshold="flex", Hess=T)
summary(mod3)
## formula: AEGrade ~ EXP + BIO + Sex
## data:
##
```

```
## link threshold nobs logLik AIC
                                     niter max.grad cond.H
## logit flexible 200 -267.05 548.10 6(0) 2.95e-13 3.0e+04
##
## Coefficients:
##
            Estimate Std. Error z value Pr(>|z|)
                        0.01252 4.554 5.28e-06 ***
## EXP
             0.05700
                        0.02097 -3.334 0.000857 ***
## BIO
            -0.06990
                        0.32393 -3.012 0.002597 **
## SexFemale -0.97560
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Threshold coefficients:
      Estimate Std. Error z value
## 0|1 -3.0663
                   0.5081 - 6.035
## 1|2 -1.4059
                   0.4722 - 2.977
## 2|3
       0.6785
                   0.4641
                            1.462
## 3|4
       1.3793
                   0.4834
                            2.853
Backward Selection: Start with final forward model
summary(stepAIC(mod3, scope=list(upper=~EXP + BIO + Sex), direction="backward", trace=1, k=2))
## Start: AIC=548.1
## AEGrade ~ EXP + BIO + Sex
##
##
               AIC
         Df
## <none>
            548.10
## - Sex
          1 555.35
## - BIO
         1 557.44
## - EXP
          1 567.56
## formula: AEGrade ~ EXP + BIO + Sex
## data:
##
                                     niter max.grad cond.H
## link threshold nobs logLik AIC
## logit flexible 200 -267.05 548.10 6(0) 2.95e-13 3.0e+04
## Coefficients:
            Estimate Std. Error z value Pr(>|z|)
                        0.01252 4.554 5.28e-06 ***
## FXP
             0.05700
## BIO
            -0.06990
                        0.02097 -3.334 0.000857 ***
                        0.32393 -3.012 0.002597 **
## SexFemale -0.97560
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Threshold coefficients:
      Estimate Std. Error z value
## 0|1 -3.0663
                   0.5081 -6.035
## 1|2 -1.4059
                   0.4722 - 2.977
## 2|3
       0.6785
                   0.4641
                          1.462
## 314
        1.3793
                   0.4834
                            2.853
```

Model 3 is the Final Model

Question 4:

Comparison of Models Using Relative Likelihood

Base model vs Final model

```
## [1] 0.0001398416

Full model vs Final model

AICr(mod3,mod2)

## [1] 0.06754238
```

BONUS Question:

Using clm() with exposure, is there a way to use AIC to help decide which exposure measure to use for the AE grades data?

Compare AIC and relative likelihood for a base model for each exposure measure. DOSE as exposure

```
mod4 <- clm(AEGrade ~ DOSE, data = d, link = "logit", threshold="flex", Hess=T)
summary(mod4)</pre>
```

```
## formula: AEGrade ~ DOSE
## data:
##
## link threshold nobs logLik AIC
                                      niter max.grad cond.H
## logit flexible 200 -281.23 572.47 5(0) 5.24e-07 3.2e+04
##
## Coefficients:
##
       Estimate Std. Error z value Pr(>|z|)
## DOSE 0.01291
                  0.00379
                             3.406 0.00066 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Threshold coefficients:
##
      Estimate Std. Error z value
## 0|1 -1.3198
                   0.2380 -5.545
        0.2203
## 1|2
                   0.2049
                          1.075
## 2|3
        2.1010
                   0.2597
                            8.089
## 3|4
        2.7494
                   0.3000
                            9.164
```

CAVGSS as exposure

```
mod5 <- clm(AEGrade ~ CAVGSS, data = d, link = "logit", threshold="flex", Hess=T)
summary(mod5)</pre>
```

```
## formula: AEGrade ~ CAVGSS
## data:
           d
##
## link threshold nobs logLik AIC niter max.grad cond.H
## logit flexible 200 -281.19 572.39 5(0) 1.67e-07 2.0e+03
##
## Coefficients:
         Estimate Std. Error z value Pr(>|z|)
##
## CAVGSS 0.07571 0.02213 3.421 0.000623 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Threshold coefficients:
      Estimate Std. Error z value
##
## 0|1 -1.1375
                   0.2696 -4.220
## 1|2
       0.4078
                   0.2471
                           1.650
## 2|3
       2.2845
                   0.2982
                           7.660
## 3|4
       2.9304
                   0.3343 8.767
CMAXSS as exposure
mod6 <- clm(AEGrade ~ CMAXSS, data = d, link = "logit", threshold="flex", Hess=T)
summary(mod6)
## formula: AEGrade ~ CMAXSS
## data:
##
## link threshold nobs logLik AIC niter max.grad cond.H
## logit flexible 200 -277.92 565.85 5(0) 4.73e-07 4.9e+03
##
## Coefficients:
         Estimate Std. Error z value Pr(>|z|)
## CMAXSS 0.05240 0.01239 4.229 2.34e-05 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Threshold coefficients:
##
      Estimate Std. Error z value
## 0|1 -1.0934
                   0.2537 - 4.311
## 1|2
       0.4735
                   0.2289
                           2.069
## 2|3
       2.3994
                   0.2894
                           8.291
        3.0612
## 314
                   0.3293
                           9.295
Relative Likelihood, CAVGSS vs DOSE
AICr (mod5, mod4)
```

Relative Likelihood, CMAXSS vs DOSE

[1] 0.9607894

AICr(mod6, mod4)

[1] 0.03651617

Relative Likelihood, CMAXSS vs CAVGSS

AICr(mod6,mod5)

[1] 0.03800643

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