Rexample

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EX (Drug)

- Response variable Y
- Dosage level (d)
- Drug product with levels: A, B, C
 - reference level: A
 - Dummy variables: P_B, P_c

Create the dataset

```
Dose = rep(c(0.2, 0.4, 0.8, 1.6), each = 3)
Product = rep(c("A", "B", "C"), 4)
y = c(2, 1.8, 1.3, 4.3, 4.1, 2, 6.5, 4.9, 2.8, 8.9, 5.7, 3.4)
ds = data.frame(Dose = (Dose), Product = factor(Product), Response = y)
xtabs(Response ~ Dose + Product, data = ds)
##
        Product
## Dose
          Α
               В
                   C
##
     0.2 2.0 1.8 1.3
##
     0.4 4.3 4.1 2.0
     0.8 6.5 4.9 2.8
##
     1.6 8.9 5.7 3.4
ds
```

```
Dose Product Response
##
## 1
       0.2
                  Α
                          2.0
## 2
       0.2
                  В
                          1.8
## 3
       0.2
                  С
                          1.3
## 4
       0.4
                          4.3
                  Α
## 5
       0.4
                  В
                          4.1
## 6
       0.4
                  С
                          2.0
## 7
       0.8
                  Α
                          6.5
## 8
       0.8
                  В
                          4.9
## 9
       0.8
                  С
                          2.8
## 10 1.6
                         8.9
                  Α
## 11 1.6
                  В
                         5.7
                  С
## 12 1.6
                          3.4
```

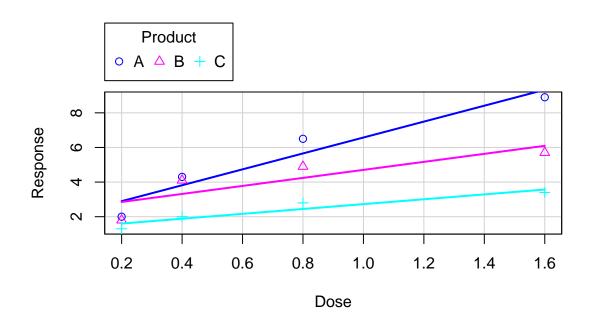
Fit multiple regression with interaction

```
Y = \beta_0 + \beta_1 d + \beta_2 P_B + \beta_3 P_C + \beta_4 dP_B + \beta_5 dP_C + \epsilon
```

```
modelfull = lm(Response ~ Dose * Product, data = ds)
summary(modelfull)
```

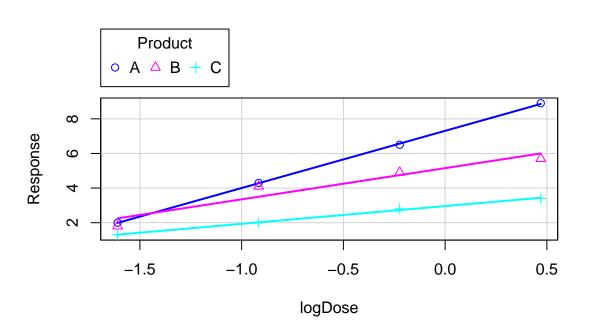
```
##
## Call:
## lm(formula = Response ~ Dose * Product, data = ds)
##
## Residuals:
##
       Min
                  1Q
                       Median
                                    3Q
                                             Max
## -1.05043 -0.40391 -0.02609 0.52739
##
## Coefficients:
                 Estimate Std. Error t value Pr(>|t|)
##
                   1.9783
                              0.7454
                                       2.654
                                              0.03783 *
## (Intercept)
## Dose
                   4.5957
                              0.8085
                                       5.684
                                              0.00128 **
## ProductB
                   0.4087
                              1.0542
                                       0.388
                                              0.71163
                                              0.55635
## ProductC
                  -0.6565
                              1.0542
                                      -0.623
## Dose:ProductB
                  -2.2783
                              1.1434
                                      -1.992
                                              0.09339 .
## Dose:ProductC
                  -3.1913
                              1.1434
                                      -2.791 0.03154 *
##
## Signif. codes:
                  0 '*** 0.001 '** 0.01 '* 0.05 '. ' 0.1 ' ' 1
##
## Residual standard error: 0.867 on 6 degrees of freedom
## Multiple R-squared: 0.9194, Adjusted R-squared: 0.8523
## F-statistic: 13.69 on 5 and 6 DF, p-value: 0.003126
```

```
library(car)
scatterplot(Response ~ Dose | Product, smooth = FALSE, data = ds)
```



Notice that a log transformation on Dose may help fit a more linear relationship

```
ds = data.frame(logDose = log(Dose), Product = factor(Product),
    Response = y)
# Full model
modelfull = lm(Response ~ logDose * Product, data = ds)
summary(modelfull)
##
## Call:
## lm(formula = Response ~ logDose * Product, data = ds)
##
## Residuals:
##
      Min
                1Q Median
                                3Q
                                       Max
## -0.4500 -0.0475 0.0000 0.0475 0.6000
##
## Coefficients:
##
                    Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                     7.3072
                                 0.2103 34.748 3.79e-08 ***
## logDose
                     3.3038
                                 0.2186 15.111 5.30e-06 ***
## ProductB
                     -2.1548
                                 0.2974 -7.245 0.000351 ***
## ProductC
                     -4.3486
                                 0.2974 -14.622 6.42e-06 ***
## logDose:ProductB -1.5004
                                 0.3092 -4.853 0.002844 **
## logDose:ProductC -2.2795
                                 0.3092 -7.372 0.000319 ***
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## Residual standard error: 0.3389 on 6 degrees of freedom
## Multiple R-squared: 0.9877, Adjusted R-squared: 0.9774
## F-statistic: 96.3 on 5 and 6 DF, p-value: 1.207e-05
scatterplot(Response ~ logDose | Product, smooth = FALSE, data = ds)
```



CI for coefficient

Create at least 90% simultaneous CI for β_4 , β_5 and $\beta_4 - \beta_5$ using **Bonferroni method**

coefficients(modelfull)

```
##
        (Intercept)
                              logDose
                                               ProductB
                                                                 ProductC
##
           7.307215
                             3.303772
                                              -2.154805
                                                                -4.348646
## logDose:ProductB logDose:ProductC
##
          -1.500403
                            -2.279458
```

- $b_4 = -1.50$
- $b_5 = -2.28$
- $b_4 b_5 = -3.78$
- Critical value $t_{1-0.1/(2\cdot3),n-p} = t_{1-0.1/6,6}$

```
vmat = vcov(modelfull)
round(vmat, 3)
```

```
##
                    (Intercept) logDose ProductB ProductC logDose:ProductB
## (Intercept)
                          0.044
                                   0.027
                                           -0.044
                                                    -0.044
                                                                      -0.027
## logDose
                          0.027
                                   0.048
                                           -0.027
                                                    -0.027
                                                                      -0.048
## ProductB
                         -0.044
                                 -0.027
                                            0.088
                                                     0.044
                                                                       0.054
## ProductC
                         -0.044 -0.027
                                            0.044
                                                     0.088
                                                                       0.027
## logDose:ProductB
                         -0.027 -0.048
                                            0.054
                                                     0.027
                                                                       0.096
## logDose:ProductC
                         -0.027 -0.048
                                            0.027
                                                     0.054
                                                                       0.048
                    logDose:ProductC
## (Intercept)
                               -0.027
## logDose
                               -0.048
## ProductB
                                0.027
## ProductC
                                0.054
## logDose:ProductB
                                0.048
## logDose:ProductC
                                0.096
```

- $S_{b_4-b_5} = \sqrt{S_{b_4}^2 + S_{b_5}^2 2S_{b_4b_5}}$
- $S_{b_4}^2$ is the (5,5)th element in the matrix $S_{b_5}^2$ is the (6,6)th element in the matrix
- $S_{b_4b_5}$ is the (5,6)th element in the matrix

CI for beta4

```
coefficients(modelfull)[5] + c(-1, 1) * qt(1 - 0.1/6, 6) * sqrt(vmat[5, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6, 0.1/6
                                                                                                          5])
```

[1] -2.3504094 -0.6503963

CI for beta5

```
coefficients(modelfull)[6] + c(-1, 1) * qt(1 - 0.1/6, 6) * sqrt(vmat[6,
6])
```

[1] -3.129465 -1.429452

CI for beta4-beta5

```
coefficients(modelfull)[5] - coefficients(modelfull)[6] + c(-1,
    1) * qt(1 - 0.1/6, 6) * sqrt(vmat[5, 5] + vmat[6, 6] - 2 *
    vmat[5, 6])
```

[1] -0.07095119 1.62906183