

# The Statistical Sleuth in R:

## Chapter 11

Linda Loi      Kate Aloisio      Ruobing Zhang      Nicholas J. Horton\*

January 24, 2019

### Contents

|          |   |          |
|----------|---|----------|
| <b>1</b> | <b>Introduction</b>   | <b>1</b> |
| <b>2</b> | <b>Alcohol metabolism in men and women</b>                      | <b>2</b> |
| 2.1      | Data coding, summary statistics and graphical display . . . . . | 2        |
| 2.2      | Multiple regression . . . . .                                   | 3        |
| 2.3      | Refining the Model . . . . .                                    | 5        |
| <b>3</b> | <b>Blood brain barrier</b>                                      | <b>7</b> |
| 3.1      | Data coding and summary statistics . . . . .                    | 7        |
| 3.2      | Graphical presentation . . . . .                                | 8        |
| 3.3      | Multiple regression . . . . .                                   | 10       |
| 3.4      | Refining the model . . . . .                                    | 11       |

## 1 Introduction

This document is intended to help describe how to undertake analyses introduced as examples in the Third Edition of the *Statistical Sleuth* (2013) by Fred Ramsey and Dan Schafer. More information about the book can be found at <http://www.proaxis.com/~panorama/home.htm>. This file as well as the associated `knitr` reproducible analysis source file can be found at <http://www.math.smith.edu/~nhorton/sleuth3>.

This work leverages initiatives undertaken by Project MOSAIC (<http://www.mosaic-web.org>), an NSF-funded effort to improve the teaching of statistics, calculus, science and computing in the undergraduate curriculum. In particular, we utilize the `mosaic` package, which was written to simplify the use of R for introductory statistics courses. A short summary of the R needed to teach introductory statistics can be found in the `mosaic` package vignette (<http://cran.r-project.org/web/packages/mosaic/vignettes/MinimalR.pdf>).

To use a package within R, it must be installed (one time), and loaded (each session). The package can be installed using the following command:

---

\*Department of Mathematics and Statistics, Smith College, [nhorton@smith.edu](mailto:nhorton@smith.edu)

```
> install.packages('mosaic') # note the quotation marks
```

Once this is installed, it can be loaded by running the command:

```
> require(mosaic)
```

This needs to be done once per session.

In addition the data files for the *Sleuth* case studies can be accessed by installing the **Sleuth3** package.

```
> install.packages('Sleuth3') # note the quotation marks
```

```
> require(Sleuth3)
```

We also set some options to improve legibility of graphs and output.

```
> trellis.par.set(theme=col.mosaic()) # get a better color scheme for lattice
> options(digits=3, show.signif.stars=FALSE)
```

The specific goal of this document is to demonstrate how to calculate the quantities described in Chapter 11: Model Checking and Refinement using R.

## 2 Alcohol metabolism in men and women

How do men and women metabolise alcohol? This is the question addressed in case study 11.1 in the *Sleuth*.

### 2.1 Data coding, summary statistics and graphical display

We begin by reading the data and summarizing the variables.

```
> summary(case1101)
```

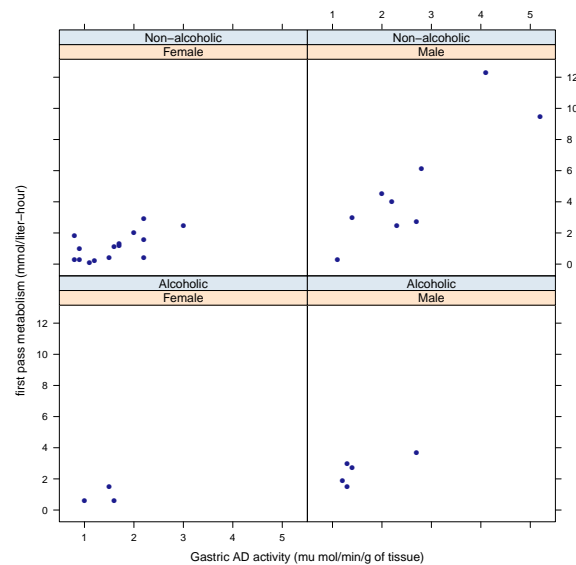
| Subject      | Metabol       | Gastric      | Sex       |
|--------------|---------------|--------------|-----------|
| Min. : 1.0   | Min. : 0.10   | Min. :0.80   | Female:18 |
| 1st Qu.: 8.8 | 1st Qu.: 0.60 | 1st Qu.:1.20 | Male :14  |
| Median :16.5 | Median : 1.70 | Median :1.60 |           |
| Mean :16.5   | Mean : 2.42   | Mean :1.86   |           |
| 3rd Qu.:24.2 | 3rd Qu.: 2.92 | 3rd Qu.:2.20 |           |
| Max. :32.0   | Max. :12.30   | Max. :5.20   |           |

Alcohol  
Alcoholic : 8  
Non-alcoholic:24

A total of 32 volunteers were included in this data. There were 18 females and 14 males, as recorded on Display 11.1 (page 311 of the *Sleuth*).

The following is a graphical display of the variables akin to Display 11.2 (page 312).

```
> xyplot(Metabol ~ Gastric | Sex+Alcohol, data=case1101, auto.key=TRUE,
+ xlab="Gastric AD activity (mu mol/min/g of tissue)",
+ ylab="first pass metabolism (mmol/liter-hour)")
```



## 2.2 Multiple regression

First we can fit a full model for estimating *metabolism* given a subjects *gastric AD activity*, whether they are *alcoholic* and *gender*. This first model is summarized on page 321 (Display 11.9).

```
> case1101 = transform(case1101, Sex = factor(Sex, levels = c("Male", "Female")))
> case1101 = transform(case1101, Alcohol = factor(Alcohol, levels = c("Non-alcoholic", "Alcoholic")))
> lm1 = lm(Metabol ~ Gastric+Sex+Alcohol+Gastric*Sex+Sex*Alcohol+Gastric*Alcohol+Gastric*Sex*Alcohol, data=case1101)
```

Call:

```
lm(formula = Metabol ~ Gastric + Sex + Alcohol + Gastric * Sex +
    Sex * Alcohol + Gastric * Alcohol + Gastric * Sex * Alcohol,
    data = case1101)
```

Residuals:

| Min    | 1Q     | Median | 3Q    | Max   |
|--------|--------|--------|-------|-------|
| -2.429 | -0.619 | -0.047 | 0.515 | 3.652 |

Coefficients:

| Estimate | Std. Error | t value | Pr(> t ) |
|----------|------------|---------|----------|
|          |            |         |          |

```

(Intercept)          -1.660      1.000    -1.66    0.110
Gastric              2.514      0.343     7.32  1.5e-07
SexFemale            1.466      1.333     1.10    0.282
AlcoholAlcoholic    2.552      1.946     1.31    0.202
Gastric:SexFemale   -1.673      0.620    -2.70    0.013
SexFemale:AlcoholAlcoholic -2.252      4.394    -0.51    0.613
Gastric:AlcoholAlcoholic -1.459      1.053    -1.39    0.179
Gastric:SexFemale:AlcoholAlcoholic 1.199      2.998     0.40    0.693

```

```

Residual standard error: 1.25 on 24 degrees of freedom
Multiple R-squared: 0.828, Adjusted R-squared: 0.777
F-statistic: 16.5 on 7 and 24 DF, p-value: 9.35e-08

```

Next we can calculate a number of model diagnostics, including leverage, studentized residuals and Cook's distance (pages 325-327).

```
> require(MASS)
```

```

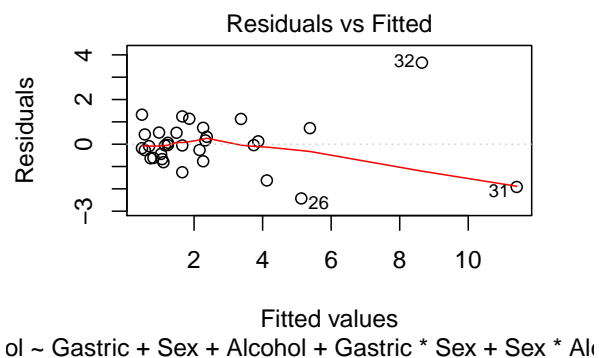
> case1101 = transform(case1101, hat = hatvalues(lm1))
> case1101 = transform(case1101, studres = studres(lm1))
> case1101 = transform(case1101, cooks = cooks.distance(lm1))
> # display a particular row
> case1101[31,]

```

| Subject | Metabol | Gastric | Sex      | Alcohol       | hat   | studres | cooks |
|---------|---------|---------|----------|---------------|-------|---------|-------|
| 31      | 31      | 9.5     | 5.2 Male | Non-alcoholic | 0.601 | -2.72   | 1.1   |

The following is a residual plot for the full model akin to Display 11.7 (page 319).

```
> plot(lm1, which=1)
```



From these diagnostics it appears that observations 31 and 32 may be influential points. Therefore, we next re-fit the full model excluding these two observations. The following results are found in Display 11.9 and discussed on page 321.

```

> case11012 = case1101[-c(31, 32),]
> lm2 = lm(Metabol ~ Gastric+Sex+Alcohol+Gastric*Sex+Sex*Alcohol+Gastric*Alcohol+Gastric*Sex*Alcohol,
           data = case11012)

Call:
lm(formula = Metabol ~ Gastric + Sex + Alcohol + Gastric * Sex +
    Sex * Alcohol + Gastric * Alcohol + Gastric * Sex * Alcohol,
    data = case11012)

Residuals:
    Min       1Q   Median       3Q      Max
-1.8076 -0.5701 -0.0466  0.4976  1.4002

Coefficients:
              Estimate Std. Error t value Pr(>|t|)
(Intercept)    -0.680     1.309   -0.52  0.6088
Gastric         1.921     0.608    3.16  0.0046
SexFemale       0.486     1.467    0.33  0.7436
AlcoholAlcoholic 1.572     1.812    0.87  0.3949
Gastric:SexFemale -1.081    0.721   -1.50  0.1483
SexFemale:AlcoholAlcoholic -1.272    3.467   -0.37  0.7172
Gastric:AlcoholAlcoholic -0.866    0.963   -0.90  0.3784
Gastric:SexFemale:AlcoholAlcoholic 0.606    2.316    0.26  0.7961

Residual standard error: 0.941 on 22 degrees of freedom
Multiple R-squared:  0.685, Adjusted R-squared:  0.585
F-statistic: 6.83 on 7 and 22 DF,  p-value: 0.000226

```

### 2.3 Refining the Model

This section addresses the process of refining the model. We first tested the lack of fit for the removal of `Alcohol` as shown in Display 11.13 (page 329).

```

> lm3 = lm(Metabol ~ Gastric+Sex+Gastric*Sex, data=case11012); summary(lm3)

Call:
lm(formula = Metabol ~ Gastric + Sex + Gastric * Sex, data = case11012)

Residuals:
    Min       1Q   Median       3Q      Max
-1.5962 -0.6025 -0.0408  0.4759  1.6473

Coefficients:
              Estimate Std. Error t value Pr(>|t|)
(Intercept)    -0.680     1.309   -0.52  0.6088
Gastric         1.921     0.608    3.16  0.0046
SexFemale       0.486     1.467    0.33  0.7436
Gastric:SexFemale -1.081    0.721   -1.50  0.1483

```

```
(Intercept)      0.0695      0.8019      0.09      0.9316
Gastric          1.5654      0.4074      3.84      0.0007
SexFemale       -0.2668      0.9932     -0.27      0.7904
Gastric:SexFemale -0.7285      0.5394     -1.35      0.1885

Residual standard error: 0.882 on 26 degrees of freedom
Multiple R-squared:  0.673, Adjusted R-squared:  0.635
F-statistic: 17.8 on 3 and 26 DF,  p-value: 1.71e-06
```

```
> anova(lm3, lm2) # page 322
```

Analysis of Variance Table

Model 1: Metabol ~ Gastric + Sex + Gastric \* Sex

Model 2: Metabol ~ Gastric + Sex + Alcohol + Gastric \* Sex + Sex \* Alcohol +  
Gastric \* Alcohol + Gastric \* Sex \* Alcohol

|   | Res.Df | RSS  | Df | Sum of Sq | F    | Pr(>F) |
|---|--------|------|----|-----------|------|--------|
| 1 | 26     | 20.2 |    |           |      |        |
| 2 | 22     | 19.5 | 4  | 0.74      | 0.21 | 0.93   |

Next we assessed a model without an intercept which is scientifically plausible as summarized in Display 11.14 (page 329).

```
> lm4 = lm(Metabol ~ Gastric+Gastric:Sex - 1, data=case11012); summary(lm4)
```

Call:

```
lm(formula = Metabol ~ Gastric + Gastric:Sex - 1, data = case11012)
```

Residuals:

|  | Min     | 1Q      | Median  | 3Q     | Max    |
|--|---------|---------|---------|--------|--------|
|  | -1.6171 | -0.6075 | -0.0262 | 0.4772 | 1.6230 |

Coefficients: (1 not defined because of singularities)

|                   | Estimate | Std. Error | t value | Pr(> t ) |
|-------------------|----------|------------|---------|----------|
| Gastric           | 0.726    | 0.121      | 5.99    | 1.9e-06  |
| Gastric:SexMale   | 0.873    | 0.174      | 5.02    | 2.6e-05  |
| Gastric:SexFemale | NA       | NA         | NA      | NA       |

Residual standard error: 0.852 on 28 degrees of freedom

Multiple R-squared: 0.877, Adjusted R-squared: 0.868

F-statistic: 99.9 on 2 and 28 DF, p-value: 1.8e-13

```
> anova(lm4, lm3)
```

Analysis of Variance Table

```

Model 1: Metabol ~ Gastric + Gastric:Sex - 1
Model 2: Metabol ~ Gastric + Sex + Gastric * Sex
  Res.Df  RSS Df Sum of Sq    F Pr(>F)
1      28 20.3
2      26 20.2  2      0.094 0.06  0.94

```

Note that the “Summary of Statistical Findings” section (page 312) is based on this final model.

### 3 Blood brain barrier

Neuroscientists working to better understand the blood brain barrier have infused rats with cells to induce brain tumors. This is the topic addressed in case study 11.2 in the *Sleuth*.

#### 3.1 Data coding and summary statistics

We begin by reading the data, performing transformations where needed and summarizing the variables.

```

> names(case1102)

[1] "Brain"      "Liver"      "Time"       "Treatment"  "Days"       "Sex"
[7] "Weight"     "Loss"       "Tumor"

> case1102 = transform(case1102, Y = Brain/Liver)
> case1102 = transform(case1102, logliver = log(Liver))
> case1102 = transform(case1102, logbrain = log(Brain))
> case1102 = transform(case1102, SAC = as.factor(Time))
> case1102 = transform(case1102, logy = log(Brain/Liver))
> case1102 = transform(case1102, logtime = log(Time))
> case1102 = transform(case1102, Treat = relevel(Treatment, ref="NS"))
> summary(case1102)

```

| Brain          | Liver           | Time         | Treatment | Days       |
|----------------|-----------------|--------------|-----------|------------|
| Min. : 1334    | Min. : 928      | Min. : 0.5   | BD:17     | Min. : 9   |
| 1st Qu.: 19281 | 1st Qu.: 16210  | 1st Qu.: 1.1 | NS:17     | 1st Qu.:10 |
| Median : 32572 | Median : 643965 | Median : 3.0 |           | Median :10 |
| Mean : 39965   | Mean : 668776   | Mean :23.5   |           | Mean :10   |
| 3rd Qu.: 50654 | 3rd Qu.:1318557 | 3rd Qu.:24.0 |           | 3rd Qu.:10 |
| Max. :123730   | Max. :1790863   | Max. :72.0   |           | Max. :11   |

| Sex       | Weight      | Loss          | Tumor       | Y            |
|-----------|-------------|---------------|-------------|--------------|
| Female:26 | Min. :184   | Min. : -4.90  | Min. : 25   | Min. :0.01   |
| Male : 8  | 1st Qu.:225 | 1st Qu.: 1.20 | 1st Qu.:136 | 1st Qu.:0.03 |
|           | Median :240 | Median : 3.95 | Median :166 | Median :0.12 |
|           | Mean :242   | Mean : 3.64   | Mean :183   | Mean :1.50   |
|           | 3rd Qu.:259 | 3rd Qu.: 5.97 | 3rd Qu.:223 | 3rd Qu.:1.95 |

```

          Max. :298   Max. :12.80   Max. :484   Max. :8.55
logliver  logbrain   SAC         logy         logtime
Min.   : 6.83   Min.   : 7.20   0.5:9   Min.   : -4.58   Min.   : -0.69
1st Qu.: 9.69   1st Qu.: 9.86   3 :9   1st Qu.: -3.39   1st Qu.: -0.25
Median :13.37   Median :10.39   24 :8   Median : -2.13   Median : 1.10
Mean   :11.61   Mean   :10.23   72 :8   Mean   : -1.39   Mean   : 1.86
3rd Qu.:14.09   3rd Qu.:10.83           3rd Qu.: 0.67   3rd Qu.: 3.18
Max.   :14.40   Max.   :11.73           Max.   : 2.15   Max.   : 4.28
Treat
NS:17
BD:17

```

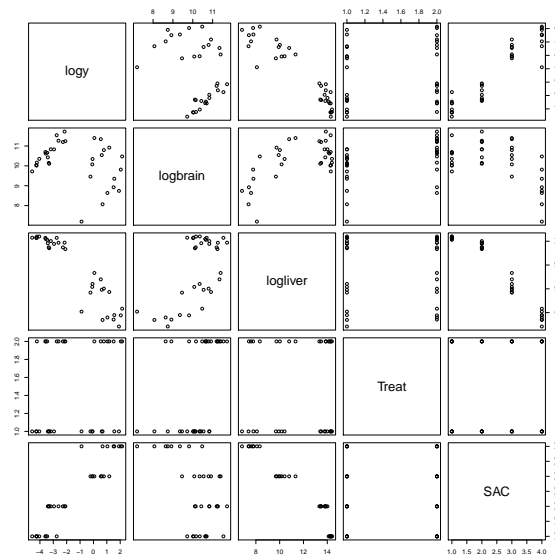
A total of 34 rats were included in this experiment. Each rat was given either the barrier solution ( $n = 17$ ) or a normal saline solution ( $n = 17$ ). Then variables of interest were calculated and are displayed in Display 11.4 (page 314 of the *Sleuth*).

We can graphically relationships between the variables using a pairs plot.

```

> smallds = subset(case1102, select=c("logy", "logbrain", "logliver", "Treat", "SAC"))
> pairs(smallds)

```

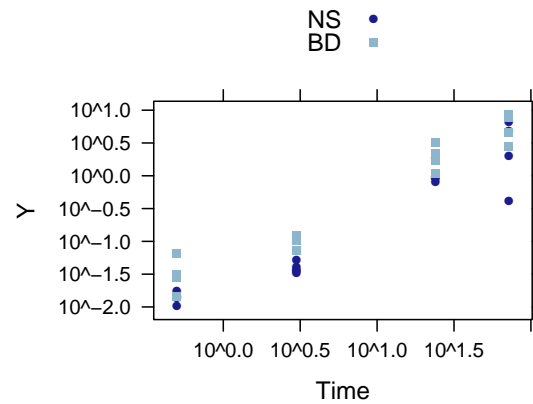


### 3.2 Graphical presentation

The following displays a scatterplot of log ratio ( $Y$ ) as a function of log time, akin to Display 11.5 on page 315.

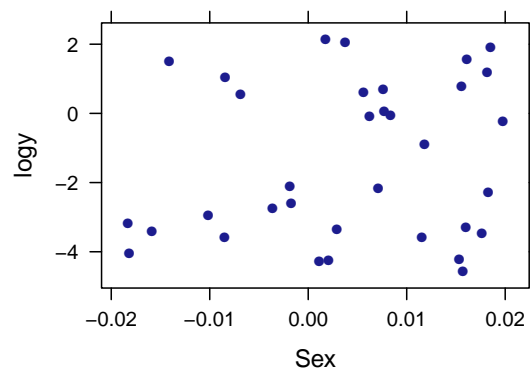


```
> xyplot(Y ~ Time, group=Treat, scales=list(y=list(log=TRUE),
+ x=list(log=TRUE)), auto.key=TRUE, data=case1102)
```

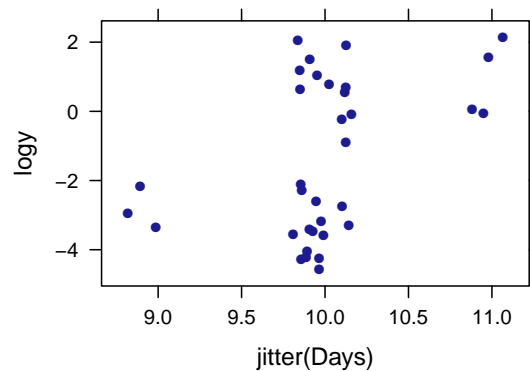


The following graphs are akin to the second and third plots in Display 11.16 on page 333.

```
> case1102=transform(case1102, female = ifelse(Sex=="F", 1, 0))
> xyplot(logy ~ jitter(female), xlab="Sex", data=case1102)
```



```
> xyplot(logy ~ jitter(Days), data=case1102)
```



### 3.3 Multiple regression

We first fit a model that reflects the initial investigation. This is the proposed model from page 317.

```
> lm1 = lm(logy ~ SAC+Treat+SAC*Treat+Days+Sex+
+ Weight+Loss+Tumor, data=case1102); summary(lm1)
```

Call:

```
lm(formula = logy ~ SAC + Treat + SAC * Treat + Days + Sex +
    Weight + Loss + Tumor, data = case1102)
```

Residuals:

| Min     | 1Q      | Median | 3Q     | Max    |
|---------|---------|--------|--------|--------|
| -1.4056 | -0.2559 | 0.0458 | 0.1957 | 1.1583 |

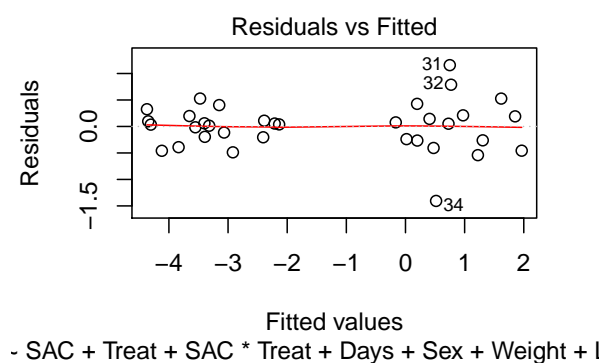
Coefficients:

|               | Estimate  | Std. Error | t value | Pr(> t ) |
|---------------|-----------|------------|---------|----------|
| (Intercept)   | -3.836741 | 3.391046   | -1.13   | 0.271    |
| SAC3          | 1.015463  | 0.399578   | 2.54    | 0.019    |
| SAC24         | 4.337135  | 0.477836   | 9.08    | 1.0e-08  |
| SAC72         | 5.010605  | 0.454953   | 11.01   | 3.5e-10  |
| TreatBD       | 0.795999  | 0.378970   | 2.10    | 0.048    |
| Days          | -0.036987 | 0.295645   | -0.13   | 0.902    |
| SexMale       | 0.001295  | 0.373368   | 0.00    | 0.997    |
| Weight        | -0.000558 | 0.005330   | -0.10   | 0.918    |
| Loss          | -0.059544 | 0.030422   | -1.96   | 0.064    |
| Tumor         | 0.001551  | 0.001226   | 1.26    | 0.220    |
| SAC3:TreatBD  | 0.179831  | 0.551964   | 0.33    | 0.748    |
| SAC24:TreatBD | -0.386047 | 0.585450   | -0.66   | 0.517    |
| SAC72:TreatBD | 0.379104  | 0.569242   | 0.67    | 0.513    |

```
Residual standard error: 0.564 on 21 degrees of freedom
Multiple R-squared: 0.96, Adjusted R-squared: 0.937
F-statistic: 41.9 on 12 and 21 DF, p-value: 6.45e-12
```

We can then display a residual plot to assess the fit of the above model. This is provided in Display 11.6 (page 318).

```
> plot(lm1, which=1)
```



### 3.4 Refining the model

Lastly, we fit a refined model. These results can be found in Display 11.17 (page 334).

```
> lm2 = lm(logy ~ SAC+Treat, data=case1102); summary(lm2)
```

Call:

```
lm(formula = logy ~ SAC + Treat, data = case1102)
```

Residuals:

| Min     | 1Q      | Median  | 3Q     | Max    |
|---------|---------|---------|--------|--------|
| -1.7402 | -0.1755 | -0.0178 | 0.2477 | 1.0551 |

Coefficients:

|             | Estimate | Std. Error | t value | Pr(> t ) |
|-------------|----------|------------|---------|----------|
| (Intercept) | -4.302   | 0.205      | -21.01  | < 2e-16  |
| SAC3        | 1.134    | 0.252      | 4.50    | 0.00010  |
| SAC24       | 4.257    | 0.259      | 16.43   | 3.1e-16  |
| SAC72       | 5.154    | 0.259      | 19.89   | < 2e-16  |
| TreatBD     | 0.797    | 0.183      | 4.35    | 0.00016  |

Residual standard error: 0.533 on 29 degrees of freedom

```
Multiple R-squared: 0.951, Adjusted R-squared: 0.944  
F-statistic: 140 on 4 and 29 DF, p-value: <2e-16
```

```
> anova(lm2, lm1)
```

```
Analysis of Variance Table
```

```
Model 1: logy ~ SAC + Treat
```

```
Model 2: logy ~ SAC + Treat + SAC * Treat + Days + Sex + Weight + Loss +  
Tumor
```

|   | Res.Df | RSS  | Df | Sum of Sq | F    | Pr(>F) |
|---|--------|------|----|-----------|------|--------|
| 1 | 29     | 8.23 |    |           |      |        |
| 2 | 21     | 6.68 | 8  | 1.55      | 0.61 | 0.76   |