# The Statistical Sleuth in R: Chapter 11

Linda Loi	Kate Aloisio	Ruobing Zhang	Nicholas J. Horton <sup>*</sup>
-----------	--------------	---------------	---------------------------------

January 25, 2024

# Contents

1	Intr	roduction	1
<b>2</b>	Alc	ohol metabolism in men and women	<b>2</b>
	2.1	Data coding, summary statistics and graphical display	2
	2.2	Multiple regression	3
	2.3	Refining the Model	5
3	Blo	od brain barrier	7
	3.1	Data coding and summary statistics	$\overline{7}$
	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:

<sup>\*</sup>Department of Mathematics and Statistics, Smith College, 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
                                                  Sex
                                                                   Alcohol
                                  Gastric
Min. : 1.0
               Min. : 0.10
                                                                     : 8
                               Min.
                                      :0.80
                                              Female:18
                                                          Alcoholic
                                              Male :14
 1st Qu.: 8.8
               1st Qu.: 0.60
                               1st Qu.:1.20
                                                          Non-alcoholic:24
Median :16.5
               Median : 1.70
                               Median :1.60
       :16.5
                      : 2.42
                                       :1.86
 Mean
               Mean
                               Mean
 3rd Qu.:24.2
               3rd Qu.: 2.92
                               3rd Qu.:2.20
Max. :32.0
               Max. :12.30
                               Max. :5.20
```

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", "Alcohol
> lm1 = lm(Metabol ~ Gastric+Sex+Alcohol+Gastric*Sex+Sex*Alcohol+Gastric*Alcohol+Gastric*Sex*A
Call:
lm(formula = Metabol ~ Gastric + Sex + Alcohol + Gastric * Sex +
   Sex * Alcohol + Gastric * Alcohol + Gastric * Sex * Alcohol,
   data = case1101)
Residuals:
  Min
         1Q Median
                        ЗQ
                              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
```

#### 2 ALCOHOL METABOLISM IN MEN AND WOMEN

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	
Destinal standard concern 1 OF an OA	democe en	free a dam			

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 resids 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)



ol ~ Gastric + Sex + Alcohol + Gastric \* Sex + Sex \* Al

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+Gastr	ic*Sex+Sex	<pre>x*Alcohol</pre>	l+Gastric*Alco	ohol+Gastric*Sex*A			
Call:								
<pre>lm(formula = Metabol ~ Gastric +</pre>	Sex + Alcohol	+ Gastric	c * Sex -	÷				
Sex * Alcohol + Gastric * Alc	ohol + Gastri	c * Sex *	Alcohol	2				
data = case11012)								
Residuals:								
Min 1Q Median 3Q	Max							
-1.8076 -0.5701 -0.0466 0.4976	1.4002							
Coefficients:								
	Estimate St	d. Error t	t value l	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:AlcoholAlcoholi	c 0.606	2.316	0.26	0.7961				
Residual standard error: 0.941 on	22 degrees o	f freedom						
Multiple R-squared: 0.685, Adjust	ed 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|)
```

5

(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: 0.88	32 on 26 de	egrees of	freedom	
Multiple R-squared	l: 0.673,Ad	ljusted R-s	squared:	0.635	
F-statistic: 17.8	on 3 and 26	6 DF, p-va	alue: 1.7	1e-06	
> anova(lm3, lm2)	# page 322				
Apolugia of Verier	a Table				
Analysis of Varian	ice labie				
Model 1: Metabol ~	Gastric +	Sex + Gast	cric * Se	x	
				stric * Sex + Sex *	<pre>Alcohol +</pre>
Gastric * Alco					
Res.Df RSS Df S					
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)
        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 Dave
```

Dra.	111		LLV	er		1.	Lille	Ireau	nent		Da	.ys
Min.	: 13	334	Min.	: 9	28	Min.	: 0.5	BD:17		Min.		: 9
1st Qu.	: 192	281	1st Qu.	: 162	10	1st Qu	.: 1.1	NS:17		1st	Qu.	:10
Median	: 325	572	Median	: 6439	65	Median	: 3.0			Medi	an	:10
Mean	: 399	965	Mean	: 6687	76	Mean	:23.5			Mean		:10
3rd Qu.	: 506	654	3rd Qu.	:13185	57	3rd Qu	.:24.0			3rd	Qu.	:10
Max.	:123	730	Max.	:17908	63	Max.	:72.0			Max.		:11
Sex		V	leight		Los	3	Tun	nor		Y		
Female:	26	Min.	:184	Min.	:-	-4.90	Min.	: 25	Min	. :	0.0	1
Male :	8	1st G	)u.:225	1st	Qu.:	1.20	1st Qu.	:136	1st	Qu.:	0.0	3
		Media	an :240	Medi	an :	3.95	Median	:166	Med	ian :	0.1	2
		Mean	:242	Mean	. :	3.64	Mean	:183	Mear	ı :	1.5	0
		3rd G	u.:259	3rd	Qu.:	5.97	3rd Qu.	:223	3rd	Qu.:	1.9	5

	Max. :2	98 Max.	:12.80	Max. :484	Max. :8.55	
logliver	lo	gbrain	SAC	logy	logtime	
Min. : 6.	83 Min.	: 7.20	0.5:9	Min. :-4.58	Min. :-0.69	
1st Qu.: 9.	69 1st Q	u.: 9.86	3 :9	1st Qu.:-3.39	1st Qu.:-0.25	
Median :13.	37 Media	n :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 Q	u.: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
                            ЗQ
                                   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
                                    2.10
              0.795999
                         0.378970
                                             0.048
             -0.036987
                                    -0.13
                                             0.902
Days
                         0.295645
SexMale
               0.001295
                         0.373368
                                   0.00
                                             0.997
                                    -0.10
Weight
             -0.000558
                         0.005330
                                             0.918
Loss
             -0.059544
                        0.030422
                                   -1.96
                                             0.064
Tumor
               0.001551
                                    1.26
                                             0.220
                        0.001226
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)
```



Fitted values - SAC + Treat + SAC \* Treat + Days + Sex + Weight + I

### 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: ЗQ Min 1Q Median 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 4.50 0.00010 1.134 SAC3 0.252 SAC24 4.257 0.259 16.43 3.1e-16 SAC72 5.154 0.259 19.89 < 2e-16 0.797 TreatBD 0.183 4.35 0.00016 Residual standard error: 0.533 on 29 degrees of freedom