# An Introduction to R

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#### 11 Writing Your Own Functions

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# 1 What Is R? Why Use R?

According to the R website (http://www.R-project.org):

- Language and environment for statistical computing and graphics
- GNU project, similar to the S Language (S-Plus), and environment which was developed at Bell Laboratories (formerly AT&T, now Lucent Technologies) by John Chambers and colleagues
- Available as *free* software
- Runs on several UNIX platforms, Linux platforms, Windows, and MacOS
- *Greatest strength:* Ease with which well-designed publication-quality plots can be produced, including mathematical symbols and formulae where needed
- Open-source and highly extensible; constantly expanding

- Can be extended beyond basic statistics via packages

- Able to define new *functions*, increasing its functionality, and run extensive simulations ("looping")
- Because it is command driven, analysis and results are completely *reproducible* if properly documented
  - Not always true with menu driven analysis packages; much harder to document

### 2 Sourcing / Downloading R

- R Website: http://www.R-project.org
- Windows (95 and later), Linux/UNIX, MacOS X
- To download:
  - Under Download link: CRAN > United States of America
  - http://cran.r-project.org/

### 3 Helpful References

- Documentation section of R website
  - Manuals; FAQs; contributed manuals, tutorials, etc. by users of R (e.g. Frank Harrell); newsletter, help pages, publications
- Biostats department website: http://biostat.mc.vanderbilt.edu
  - Statistical Computing > R and S-Plus Software and Documentation
- Other publications:
  - Simple R Using R for Introductory Statistics by John Verzani (pdf)
  - An Introduction to S and the Hmisc and Design Libraries by Carlos Alzola and Frank E. Harrell (pdf)
  - Data Analysis and Graphics Using R by John Maindonald and John Braun (book)
  - Statistical Tables and Plots Using S and LaTex by Frank E. Harrell (pdf)
  - R Data Import/Export by the R Development Core Team (pdf)
  - R for Beginners by Emmanuel Paradis (pdf)
  - An Introduction to R by W.N. Venables, D.M. Smith, and the R Development Core Team (pdf or book)
  - Introductory Statistics with R by Peter Dalgaard (book)
  - Regression Modelling Strategies by Frank E. Harrell (book)
  - Not R specific, but good sources:
    - \* The Elements of Graphing Data and Visualizing Data by William S. Cleveland (books)

### 4 Interacting with R

- R evaluates and prints out the result of any *expression* that one enters in at the *command line prompt* in the *console window*<sup>1</sup>
- The result, if any, appears on subsequent lines
- Simplest use of R: Using R as a calulator
  - > 2 + 2
  - [1] 4
  - > sqrt(10)
  - [1] 3.162278
- Most frequent use of R: Using R to evaluate *expressions*, which include *functions* and defined *objects*

### 5 R Objects

- All R entities, including *functions* and *data structures*, exist as *objects*<sup>2</sup>
- They can all be operated on as data within *expressions*
- If you type the name of an *object* at the command prompt, the contents of the *object* are printed out (e.g. type q, mean)
  - *Objects* are case sensitive (e.g. Age and age would refer to two different *objects*)
- Managing Project Data in R:
  - By default, R stores all the objects created in your session in a single file: .RData, which is directory specific
  - When running R interactively, R asks whether you want to update
     .RData to contain newly created objects upon termination of the session
  - Since many of the objects are temporary, best to answer n ("No") to this question and not use the .RData mechanism
  - Instead, use the save function to store some of your newly created data frames and selected other objects (i.e. regression fit objects that took significant execution time to create) permanently

 $<sup>^{1}</sup>Data$  Analysis Using R, Maindonald

<sup>&</sup>lt;sup>2</sup>Data Analysis Using R, Maindonald

- Use the save's compress argument to store the resulting file very compactly
- Example: A hypothetical data set containing a sample of 500 subjects. Each had three potential predictors measured: age, sex, and systolic blood pressure, and a diagnosis of a certain disease (present/absent).

```
> library(Hmisc)
  > library(Design)
  > prob1 <- read.table("prob1.csv", header = T, sep = ",")</pre>
  > prob1 <- upData(prob1, labels = c(age = "Age", sex = "Sex", sysbp = "Systolic Blo
        dz = "Disease"), units = c(age = "years"), levels = list(sex = c("Female",
        "Male")))
  +
  Input object size:
                               18688 bytes;
                                                      4 variables
  New object size:
                             19544 bytes;
                                                   4 variables
  > m1 <- lm(age ~ sysbp + sex, data = prob1)</pre>
  > save(m1, prob1, file = "prob1.rda", compress = TRUE)
- To retrieve the saved objects in a future session, use the load function:
  > load("prob1.rda")
```

### 6 Functions

- Almost everything in R is done by calling *functions*
- Most functions have *arguments* that pass values to the function for it to work on or to specify detailed options on how it should do its work<sup>3</sup>
- *Arguments* are given to the function either by name or by their sequential position in the series of arguments
- The 12,000 (and growing) functions in R are organized into *packages*, some of which are loaded when you start R, while others must be loaded explicitly using the library function
  - The *Hmisc* package (i.e. "Harrell Miscellaneous"), which was developed by Frank E. Harrell, contains many functions useful for data analysis, high-level graphics, utility operations, functions for computing sample size and power, importing datasets, imputing missing values, advanced table making, variable clustering, character string manipulation, conversion of S objects to LaTeX code, and recoding variables.<sup>4</sup>
  - The Design library, which was also developed by Frank E. Harrell, is a collection of about 180 functions that assist and streamline regression modeling, testing, estimation, validation, graphics, prediction, and typesetting by storing enhanced model design attributes in the fit.<sup>5</sup>

 $<sup>^3</sup>An$  Introduction to S and the Hmisc and Design Libraries, Harell

<sup>&</sup>lt;sup>4</sup>An Introduction to S and the Hmisc and Design Libraries, Harell

<sup>&</sup>lt;sup>5</sup>The Information on Package 'Design' help page in R (library(help="Design"))

### 7 R Details

#### • Command line prompt: >

- R commands (expressions) are typed following this prompt
- Example:

> mean(c(12, 10, 20, 15, 30))

[1] 17.4

#### • Continuation prompt: +

- Appears in command window when, following a carriage return, the command is still not complete
- Example:
  - > mean(c(5, 4, 3, 7, 9, NA, 10, 19, 18, 14, 2, 15, 22, 35, 49, + NA, NA, 54), na.rm = T)
  - [1] 17.73333
- Comment: #
  - Anything following a # on the command line is taken as "comment" and ignored by R
  - Example: > 175\*(8/5) # convert 175 miles to kms
- Multiple command separator: ;
  - Allows multiple commands to appear on one line
  - *Example:* > exp(1); 200-15
- Assignment: <-
  - Left diamond bracket (<) followed by a minus sign (-)
  - Example: x <- 2 implies "x is assigned to 2"
    - > x <- 2 > x
    - [1] 2
- Arithmetic operators: +, -, \*, /, ^, exp, log, log10
- Comparison operators: <, >, <=, >=, ==, !=
- Logical operators: &, |, !

### 8 Importing Data

- The easiest form of data import into R is a simple text file<sup>6</sup>
  - Often have the data saved as an Excel spreadsheet, SPSS file, or STATA file
  - Export the data as a text file (e.g. a delimited file; either tabdelimited or comma-delimited)
- General Tip: Start R session from within the relevant project directory (folder)
  - R automatically stores all *objects* created in your R session to your workspace (.Rdata file), which is directory specific
  - Allows you to keep defined *objects* separate for each project and not overload your *workspace*
  - Allows you to easily reference your data file for importing
- Example: the Low Birthweight Study data (Hosmer & Lemeshow, Applied Logistic Regression) read in as a tab-delimited file
  - The original file is an Excel spreadsheet (.xls)
  - NOTE: No missing values, so do not need to worry about specifying them
  - Use Excel to save the file as a tab-delimited file (.txt)
    - \* File > Save As: Save As Type: Text (Tab delimited) (\*.txt)
  - In R:

```
> lowbw <- read.table("lowbwt.txt", header = T)</pre>
```

- Example: the Primary Biliary Cirrhosis (PBC) Trial data (Fleming & Harrington, Counting Processes and Survival Analysis) read in as a commadelimited file
  - The original file is an Excel spreadsheet (.xls)
  - Use Excel to save the file as a comma-delimited file (.csv)
    - \* File > Save As: Save As Type: CSV (Comma delimited) (\*.csv)
  - NOTE: Data contains missing values:
    - \* In Excel, replace all blank cells with "NA" (before saving as .csv)
       Edit > Find & Replace: Search for: (blank), Replace with "NA"
    - \* In R (after saving as .csv), specify na.string = "" in read.table function

 $<sup>^{6}\,</sup>An$  Introduction to S and the Hmisc and Design Libraries, Harell; R Data Import/Export, R Development Core Team

- In R:

```
> pbc <- read.table("liver.csv", header = T, na.string = "", sep = ",")</pre>
```

- For STATA files:
  - 1. To read in a STATA file (.dta) directly, use the read.dta function in the *foreign* library (example given in section 12.1)
  - 2. To create an ASCII file from STATA, enter the following commands in STATA (using estriol.dta for illustration):
    - Space-delimited text file: outfile using estriol.dta wide
    - Comma-delimited text file: outfile using estriol.dta wide comma
    - *NOTE:* The option wide ensures one record per line; otherwise the records are wrapped to the next line
- General Tip: Always ensure your data set has been read-in (imported) correctly
  - Use the dim function to check the dimensions (number of rows, number of columns) of your read-in data frame

> dim(lowbw)

- [1] 189 11
- > dim(pbc)
- [1] 310 20
- Use the names function to check the names of the columns of your read-in data frame

> names(lowbw)

```
[1] "id" "low" "age" "lwt" "race" "smoke" "ptl" "ht" "ui"
[10] "ftv" "bwt"
```

> names(pbc)

[1]	"age"	"albumin"	"alkphos"	"ascites"	"bili"	"cholest"
[7]	"edema"	"edmadj"	"hepmeg"	"obstime"	"platelet"	"protime"
[13]	"sex"	"sgot"	"spiders"	"stage"	"status"	"tx"
[19]	"trig"	"urinecu"				

- Use the *Hmisc* contents function to check the following attributes of the variables from your read-in *data frame*: names, labels (if any), units (if any), number of factor levels (if any), factor levels, class, storage mode, and number of NAs
  - > library(Hmisc)
  - > contents(lowbw)

Data frame:lowbw		189 observations	and 11 variables	Maximum # NAs:0
Storageidintegerlowintegerageintegerlwtintegerraceintegersmokeintegerptlintegerhtintegerintintegerftvintegerbwtinteger				
<pre>&gt; contents(pbc)</pre>		040 1		<b>W W W W</b>
Data frame:pbc		310 observations an	d 20 variables	Maximum # NAs:30
Storageagedoublealbumindoublealkphosdoubleascitesintegerbilidoublecholestintegeredemaintegeredmadjdoublehepmegintegerobstimeintegerplateletintegerprotimedoublesexintegersgotdoublespidersintegerstageintegerstatusintegertxintegertrigintegerurinecuinteger	NAs 0 0 0 28 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0			

- General Tip: Make any changes to the read-in data frame (e.g. variable names, labels, or value codes) upfront in order to take advantage of all the new annotations during your analysis
  - Use the *Hmisc* upData function to edit the read-in data frame's contents (e.g. variable names, labels, levels, units, etc.)

- The *Hmisc* upData function accomplishes the following, listed in order in which changes are executed by the function:
  - 1. optionally change names of variables to lower case
  - 2. rename variables
  - 3. adds new variables
  - 4. recomputes existing variables from the original variable and/or from other variables in the data frame
  - 5. changes the storage mode of variables to the most efficient mode (as done with clean.import)
  - 6. drops variables
  - 7. adds, changes, and combines levels of factor variables
  - 8. adds or changes variable labels attributes
  - 9. adds or changes variable units (units of measurement) attributes

```
> library(Hmisc)
> lowbw <- upData(lowbw, labels = c(id = "Subject Identification Code",</pre>
      low = "Low Birthweight?", age = "Mother's Age", lwt = "Mother's Weight at Last Menstru
      race = "Race", smoke = "Did Mother Smoke During Pregnancy?",
      ptl = "Number of Premature Labors", ht = "History of Hypertension?",
      ui = "Uterine Irritability?", ftv = "Number of Physician Visits in 1st Trimester",
      bwt = "Birthweight"), units = c(age = "years", lwt = "lbs",
      bwt = "grams"), levels = list(low = c(">2500g", "<=2500g"),</pre>
      race = c("White", "Black", "Other"), smoke = c("No", "Yes"),
      ht = c("No", "Yes"), ui = c("No", "Yes")))
                           11428 bytes;
                                                 11 variables
Input object size:
New object size:
                         14844 bytes;
                                               11 variables
> contents(lowbw)
Data frame:lowbw
                         189 observations and 11 variables
                                                               Maximum # NAs:0
                                            Labels Units Levels Storage
id
                      Subject Identification Code
                                                                 integer
                                  Low Birthweight?
                                                               2 integer
low
                                      Mother's Age years
                                                                 integer
age
         Mother's Weight at Last Menstrual Period
                                                     lbs
                                                                 integer
lwt
                                                               3 integer
race
                                              Race
               Did Mother Smoke During Pregnancy?
                                                               2 integer
smoke
                       Number of Premature Labors
ptl
                                                                 integer
ht
                         History of Hypertension?
                                                               2 integer
                             Uterine Irritability?
                                                               2 integer
ui
      Number of Physician Visits in 1st Trimester
                                                                 integer
ftv
bwt
                                       Birthweight grams
                                                                 integer
```

+-----+----+ |Variable|Levels 1 +----+ low |>2500g,<=2500g | +----+ | race |White,Black,Other| +----+ | smoke |No,Yes | +----+ No,Yes l ht +----+ | ui No,Yes +----+ > pbc <- upData(pbc, labels = c(age = "Age", albumin = "Serum Albumin",</pre> alkphos = "Serum Alkaline Phosphatase", ascites = "Presense of Ascites", bili = "Serum Bilirubin", cholest = "Serum Cholesterol", + edema = "Presence of Edema", edmadj = "Graded Measurement of Edema", + hepmeg = "Presence of Hepatomegaly", obstime = "Observation Time", + + platelet = "Platelet Count", protime = "Prothrombin Time", sex = "Sex", sgot = "Serum SGOT", spiders = "Presence of Spider Angiomata", + stage = "Stage of Disease", status = "Survival Status", tx = "Treatment Group", + trig = "Serum Triglycerides", urinecu = "Urine Copper"), + + units = c(age = "year", obstime = "day"), levels = list(ascites = c("Absent", "Present"), edema = c("Absent", "Present"), hepmeg = c("Absent", + "Present"), sex = c("Male", "Female"), spiders = c("Absent", + "Present"), stage = c("Best", "Better", "Worse", "Worst"), + status = c("Censored", "Died"), tx = c("Placebo", "Drug"))) + Input object size: 38392 bytes; 20 variables New object size: 43976 bytes; 20 variables > contents(pbc) Data frame:pbc 310 observations and 20 variables Maximum # NAs:30 Labels Units Levels Storage NAs Age year double 0 age Serum Albumin double 0 albumin Serum Alkaline Phosphatase double alkphos 0 ascites Presense of Ascites 2 integer 0 Serum Bilirubin bili double 0 cholest Serum Cholesterol integer 28 Presence of Edema 2 integer edema 0 Graded Measurement of Edema double edmadj 0 Presence of Hepatomegaly hepmeg 2 integer 0 Observation Time integer obstime day 0

platelet	Platelet Count		integer	4
protime	Prothrombin Time		double	0
sex	Sex	2	integer	0
sgot	Serum SGOT		double	0
spiders	Presence of Spider Angiomata	2	integer	0
stage	Stage of Disease	4	integer	0
status	Survival Status	2	integer	0
tx	Treatment Group	2	integer	0
trig	Serum Triglycerides		integer	30
urinecu	Urine Copper		integer	2

+	++
Variable	Levels
ascites	Absent,Present
	Absent,Present
	Absent,Present
sex	Male,Female
spiders	Absent,Present
stage	Best,Better,Worse,Worst
status	Censored,Died
+	Placebo,Drug
+	++

# 9 Data Summaries

There are many functions to produce statistical summaries (including the mean, median, sd, and table functions), but here is an illustration of a few of the more "advanced" ones in the Hmisc library:<sup>7</sup>

### 9.1 The table function

```
> table(pbc$tx)
```

Placebo Drug 153 157

> table(pbc\$tx, pbc\$status)

<sup>&</sup>lt;sup>7</sup>An Introduction to S and the Hmisc and Design Libraries, Harell

Censored Died Placebo 93 60 65 Drug 92 > with(pbc, table(tx, status)) status Censored Died tx 60 Placebo 93 Drug 92 65 9.2 The *Hmisc* describe function > library(Hmisc) > describe(lowbw) lowbw 11 Variables 189 Observations id : Subject Identification Code n missing unique Mean .05 .10 .25 .50 .75 .90 189 0 189 121.1 20.8 30.8 68.0 123.0 176.0 207.2 .95 216.6 lowest : 4 10 11 13 15 , highest: 222 223 224 225 226 \_\_\_\_\_ low : Low Birthweight? n missing unique 189 0 2 >2500g (130, 69%), <=2500g (59, 31%) \_\_\_\_\_ age : Mother's Age [years] .25 .05 .75 .90 n missing unique Mean .10 .50 189 0 24 23.24 16 17 19 23 26 31 .95 32 lowest : 14 15 16 17 18, highest: 33 34 35 36 45 \_\_\_\_\_ \_\_\_\_\_ lwt : Mother's Weight at Last Menstrual Period [lbs] n missing unique Mean .05 .10 .25 .50 .75 .90 189 129.8 94.4 99.6 110.0 121.0 140.0 0 75 170.0 .95 188.2

```
lowest : 80 85 89 90 91 , highest: 215 229 235 241 250
_____
                         ------
race : Race
   n missing unique
  189 0
             3
White (96, 51%), Black (26, 14%), Other (67, 35%)
                              ------
------
smoke : Did Mother Smoke During Pregnancy?
   n missing unique
  189 0
             2
No (115, 61%), Yes (74, 39%)
_____
ptl : Number of Premature Labors
   n missing unique Mean
  189
     0
          4 0.1958
0 (159, 84\%), 1 (24, 13\%), 2 (5, 3\%), 3 (1, 1\%)
-------
                            _____
ht : History of Hypertension?
   n missing unique
  189 0
          2
No (177, 94%), Yes (12, 6%)
_____
ui : Uterine Irritability?
   n missing unique
  189
     0
             2
No (161, 85%), Yes (28, 15%)
_____
ftv : Number of Physician Visits in 1st Trimester
   n missing unique
               Mean
  189
     0 6 0.7937
       0 1 2 3 4 6
Frequency 100 47 30 7 4 1
      53 25 16 4 2 1
%
     _____
bwt : Birthweight [grams]
   n missing unique
                      .05
                                .25
                                     .50
                                          .75
                                                .90
                Mean
                           .10
                 2945 1801
                           2038
                                2414
                                     2977
  189
       0 133
                                          3475
                                               3865
  .95
 3997
```

```
lowest : 709 1021 1135 1330 1474, highest: 4167 4174 4238 4593 4990
_____
> describe(pbc$edmadj)
pbc$edmadj : Graded Measurement of Edema
   n missing unique Mean
  310 0 3 0.1113
0.0 (261, 84%), 0.5 (29, 9%), 1.0 (20, 6%)
> describe(pbc[, c("bili", "stage")])
pbc[, c("bili", "stage")]
2 Variables 310 Observations
_____
bili : Serum Bilirubin
   n missing unique Mean .05 .10 .25 .50 .75 .90
     0 84 3.27 0.500 0.600 0.800 1.350 3.475 7.210
  310
  .95
14.055
lowest : 0.3 0.4 0.5 0.6 0.7 , highest: 21.6 22.5 24.5 25.5 28.0
_____
stage : Stage of Disease
   n missing unique
  310
       0
              4
Best (16, 5%), Better (65, 21%), Worse (120, 39%), Worst (109, 35%)
_____
```

#### 9.3 The *Hmisc* bystats function

```
> library(Hmisc)
> library(Design)
> bystats(pbc$age, pbc$status)
```

Mean of pbc\$age by

```
        N
        Mean

        Censored
        185
        47.73017

        Died
        125
        53.24174

        ALL
        310
        49.95257
```

> summary(age ~ status, data = pbc)

N=310 Age

-----+ | |N |age | +----+ |Survival Status|Censored|185|47.73017| |Died |125|53.24174| 1 ---+----+---+---+----++----++ +----|Overall | |310|49.95257| +----+ > bystats(pbc\$albumin, pbc\$tx, pbc\$status, fun = quantile) quantile of pbc\$albumin by pbc\$tx, pbc\$status 0% 25% 50% 75% 100% Ν Placebo Censored 93 2.90 3.41 3.610 3.830 4.38 Censored 92 2.83 3.40 3.665 3.855 4.64 Drug Placebo Died 60 1.96 3.19 3.435 3.670 4.30 Died 65 2.10 3.05 3.350 3.700 4.40 Drug ALL 310 1.96 3.31 3.555 3.800 4.64 > summary(albumin ~ tx + status, method = "cross", data = pbc, fun = quantile) + UseMethod by tx, status +---+ N I |0% | |25% | |50% | |75% | |100%| +---+ +----+---+ tx |Censored| Died| ALL | +----+----+ |Placebo| 93 | 60 |153 | | 2.90 |1.96 |1.96 | I | 3.41 |3.19 |3.35 | | 3.610 |3.435|3.550| Т | 3.830 |3.670|3.780| L | 4.38 |4.30 |4.38 | +----+ |Drug | 92 | 65 |157 | | 2.83 |2.10 |2.10 |

1	I	3.40	3.05  3.21
1	1	3.665	3.350 3.570
1	I	3.855	3.700 3.830
1	I	4.64	4.40  4.64
+	-+-		-++
ALL	I	185	125  310
1	I	2.83	1.96  1.96
1	I	3.40	3.11  3.31
1	I	3.630	3.430 3.555
1	I	3.850	3.670 3.800
1	I	4.64	4.40  4.64
+	-+-		-++

# 9.4 Summary Statistics

```
> library(Design)
> sublowbw <- lowbw[, -1]
> summ <- summary(~., data = sublowbw)
> latex(summ, size = "smaller", middle.bold = T, digits = 3, file = "")
```

#### > lowbw\$low

```
Low Birthweight?
```

> library(Hmisc)

[1]	>2500g	<=2500g	>2500g						
[10]	<=2500g	<=2500g	>2500g	>2500g	>2500g	>2500g	<=2500g	>2500g	>2500g
[19]	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	<=2500g	<=2500g	>2500g
[28]	>2500g	>2500g	<=2500g	<=2500g	>2500g	>2500g	<=2500g	>2500g	<=2500g
[37]	>2500g	>2500g	<=2500g	<=2500g	>2500g	<=2500g	>2500g	>2500g	>2500g
[46]	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	<=2500g
[55]	>2500g	>2500g	>2500g	<=2500g	>2500g	>2500g	>2500g	<=2500g	>2500g
[64]	<=2500g	>2500g	>2500g	>2500g	>2500g	<=2500g	<=2500g	>2500g	>2500g
[73]	<=2500g	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g
[82]	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	<=2500g	>2500g	>2500g
[91]	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	>2500g	<=2500g	<=2500g
[100]	<=2500g	>2500g	<=2500g	>2500g	>2500g	>2500g	>2500g	<=2500g	>2500g
[109]	>2500g	>2500g	<=2500g	<=2500g	>2500g	>2500g	>2500g	<=2500g	>2500g
[118]	<=2500g	>2500g	>2500g	>2500g	>2500g	<=2500g	>2500g	>2500g	<=2500g
[127]	>2500g	>2500g	>2500g	<=2500g	>2500g	<=2500g	<=2500g	>2500g	>2500g
[136]	>2500g	>2500g	<=2500g	>2500g	<=2500g	>2500g	>2500g	>2500g	>2500g
[145]	>2500g	>2500g	>2500g	<=2500g	>2500g	>2500g	>2500g	>2500g	>2500g
[154]	<=2500g	<=2500g	>2500g	<=2500g	>2500g	>2500g	<=2500g	<=2500g	>2500g
[163]	>2500g	>2500g	<=2500g	>2500g	<=2500g	<=2500g	>2500g	>2500g	<=2500g
[172]	>2500g	>2500g	<=2500g	<=2500g	<=2500g	<=2500g	<=2500g	<=2500g	>2500g
[181]	>2500g	>2500g	<=2500g	<=2500g	>2500g	<=2500g	<=2500g	<=2500g	<=2500g
Level	s: >2500g	g <=2500g	5						

Table 1: Descriptive Statistics (N = 189)

Low Birthweight? : $\leq 2500g$	31% (59)
Mother's Age years	19 $23$ 26
Mother's Weight at Last Menstrual Period 1bs	110 $121$ 140
Race : White	51% (96)
Black	14% (26)
Other	35% (67)
Did Mother Smoke During Pregnancy? : Yes	39% (74)
Number of Premature Labors : 0	84% (159)
1	13% (24)
2	3% (5)
3	1% (1)
History of Hypertension? : Yes	6% (12)
Uterine Irritability? : Yes	15% (28)
Number of Physician Visits in 1st Trimester : 0	53% (100)
1	25% (47)
2	16% (30)
3	4% (7)
4	2% (4)
6	1% (1)
Birthweight grams	2414 <b>2977</b> 3475

 $a\ b\ c$  represent the lower quartile a, the median b, and the upper quartile c for continuous variables.

Numbers after percents are frequencies.

```
> unclass(lowbw$low)
 [1] 1 2 1 1 1 1 1 1 1 2 2 1 1 1 1 2 1 1 1 1 1 1 1 1 2 2 1 1 1 2 2 1 1 2 1 2 1
[38] 1 2 2 1 2 1 1 1 1 1 1 1 1 1 1 1 2 1 1 1 2 1 1 1 2 1 1 1 2 1 2 1 1 1 2 2 1 1 1 2 2 1 1 2 1
[149] 1 1 1 1 1 2 2 1 2 1 1 2 2 1 1 1 2 2 2 1 1 2 1 1 2 2 2 2 2 2 2 2 1 1 1 2 2 1
[186] 2 2 2 2 2
attr(,"levels")
[1] ">2500g" "<=2500g"
attr(,"label")
[1] "Low Birthweight?"
> unclass(lowbw$low) - 1
 [38] 0 1 1 0 1 0 0 0 0 0 0 0 0 0 0 1 0 0 0 1 0 0 0 1 0 1 0 0 0 0 1 1 0 0 1 0
[112] 1 0 0 0 1 0 1 0 0 0 1 0 0 1 0 0 1 0 0 1 0 1 0 0 0 0 1 0 1 0 0 0 0 0 0 0 0 0 0 1
[186] 1 1 1 1
attr(,"levels")
[1] ">2500g" "<=2500g"
attr(,"label")
[1] "Low Birthweight?"
> sublowbw2 <- lowbw[, -c(1, 2)]</pre>
> low.summ <- summary(unclass(lowbw$low) - 1 ~ ., data = sublowbw2)</pre>
> latex(low.summ, middle.bold = T, file = "")
```

# 10 Graphics in R

As with the statistical summaries, there is a large variety of plotting functions in R. The following is an illustration of just a few:

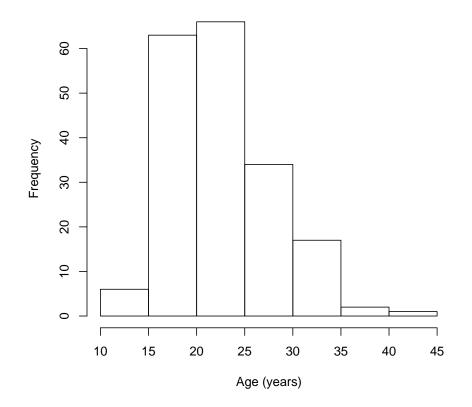
#### 10.1 Histograms

> hist(lowbw\$age, main = "Histogram of Mother's Age", xlab = "Age (years)")

Table 2: Low Birthweight?	N = 189
---------------------------	---------

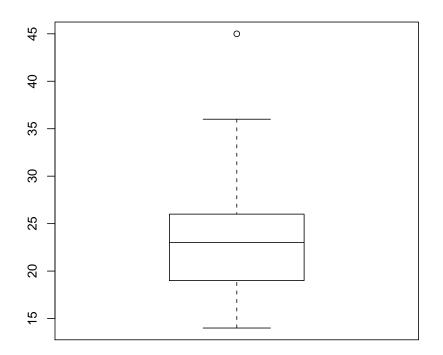
<b>Mother's Age</b> [14,20)		
[14,20)	years	
	51	0.29
[20,24)	56	0.36
[24,27)	36	0.42
[27, 45]	46	0.20
Mother's Weight at Last Menstrual Perio	d lbs	
[ 80,112)	53	0.47
[112,122)	43	0.23
[122,141]	46	0.26
[141,250]	47	0.26
Race		
White	96	0.24
Black	26	0.42
Other	67	0.37
Did Mother Smoke During Pregnancy?		
No	115	0.25
Yes	74	0.41
Number of Premature Labors		
0	159	0.26
1	24	0.67
2	5	0.40
-3	1	0.00
History of Hypertension?		
No	177	0.29
Yes	12	0.58
Uterine Irritability?		
No	161	0.28
Yes	28	0.50
Number of Physician Visits in 1st Trimes		
	100	0.36
1	47	0.23
2	30	0.23
3	7	0.57
4	. 4	0.25
6	1	0.00
Birthweight	grams	
[709,2424]	48	1.00
[2424,2992)	48	0.23
[2992,3487)	46	0.00
[3487,4990]	40	0.00
Overall	1	
	189	0.31

# Histogram of Mother's Age



# 10.2 Boxplots

> boxplot(lowbw\$age, main = "Boxplot of Mother's Age", xlab = "Age (years)")

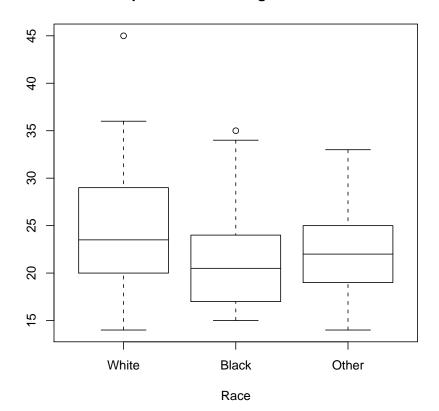


Boxplot of Mother's Age



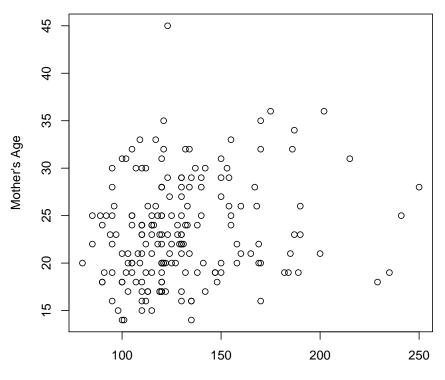
> boxplot(lowbw\$age ~ lowbw\$race, main = "Boxplot of Mother's Age Across Race", + xlab = "Race")

### **Boxplot of Mother's Age Across Race**



10.3 Scatter plots

> plot(lowbw\$age ~ lowbw\$lwt, main = "Mother's Age vs. Weight", + xlab = "Mother's Weight", ylab = "Mother's Age")

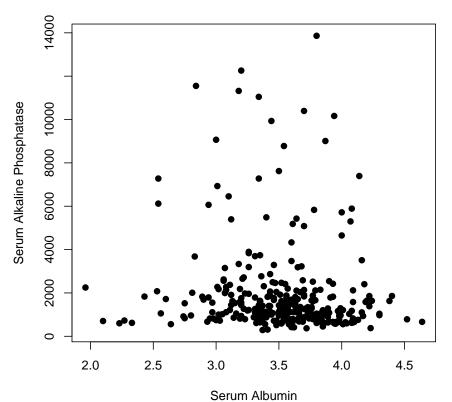


Mother's Age vs. Weight

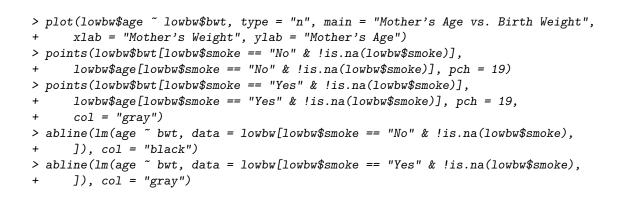


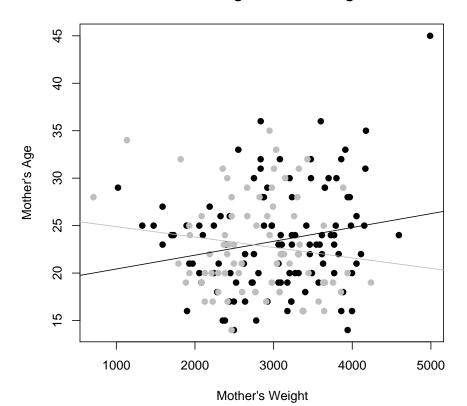
> plot(pbc\$albumin, pbc\$alkphos, main = "Serum Alk. Phos. vs. Serum Albumin", + xlab = "Serum Albumin", ylab = "Serum Alkaline Phosphatase", + pch = 19) > plot(pbc\$albumin, pbc\$alkphos, main = "Serum Alk. Phos. vs. Serum Albumin",

+ xlab = label(pbc\$albumin), ylab = label(pbc\$alkphos), pch = 19)



Serum Alk. Phos. vs. Serum Albumin



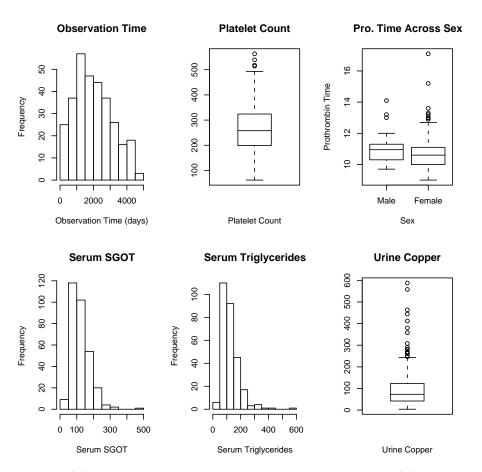


Mother's Age vs. Birth Weight

NOTES: (1) type = "n" in the plot command will produce the axes, the axes labels, and the main title for the plot, but will not plot any of the points; (2) the is.na function indicates which elements are missing (i.e. NA), so !is.na(lowbw\$smoke) will return only the non-missing values of lowbw\$smoke; (3) the abline function adds a straight line to a plot

#### 10.4 Multiple plots per page

```
> par(mfrow = c(2, 3))
> hist(pbc$obstime, main = "Observation Time", xlab = "Observation Time (days)")
> boxplot(pbc$platelet, main = "Platelet Count", xlab = "Platelet Count")
> boxplot(pbc$protime ~ pbc$sex, main = "Pro. Time Across Sex",
+ xlab = "Sex", ylab = "Prothrombin Time")
> hist(pbc$sgot, main = "Serum SGOT", xlab = "Serum SGOT")
> hist(pbc$trig, main = "Serum Triglycerides", xlab = "Serum Triglycerides")
> boxplot(pbc$urinecu, main = "Urine Copper", xlab = "Urine Copper")
> par(mfrow = c(1, 1))
```

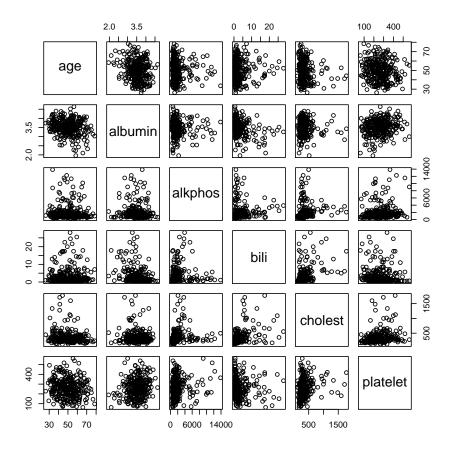


NOTES: (1) the par function is used to set graphical parameters; (2) the mfrow argument allows you to change the number of plots per page by specifying the number of rows, and the number of columns, respectively (by default, there is one plot per page, which is equivalent to one row and one column)

### 10.5 Pairs plots

```
> library(Hmisc)
```

```
> pairs(pbc[Cs(age, albumin, alkphos, bili, cholest, platelet)])
```



### 10.6 Graphs with text

rates.dat contains a dataset for a study of beta-blocker adherence post-AMI.

```
> rates <- read.table("rates.dat", header = T)
> par(oma = c(5, 0, 2, 0))
> plot(rates$day, rates$rate1, type = "1", ylim = c(0, 85), axes = F,
+ xlab = "Days since discharge", ylab = "Percent beta-blocker users (%)")
> axis(1, at = c(0, 30, 90, 180, 270, 365))
> axis(2, at = c(0, 20, 40, 60, 80))
> axis(1, at = rates$day, labels = rates$atrisk1, tick = F, line = 4,
+ cex = 0.8)
> axis(1, at = rates$day, labels = rates$atrisk0, tick = F, line = 6.5,
+ cex = 0.8)
> lines(rates$day, rates$rate0, type = "1")
> box()
```

```
> mtext("No. at-risk: patients discharged on beta-blockers", side = 1,
+ line = 4, adj = 0, cex = 0.8)
> mtext("No. at-risk: patients not discharged on beta-blockers",
+ side = 1, line = 6.5, adj = 0, cex = 0.8)
> mtext("Figure 1. Outpatient adherence to beta-blocker therapy post-AMI",
+ side = 3, cex = 1.2, line = 1)
> text(180, 70, "Discharged on beta-blockers", cex = 0.8)
> text(180, 20, "Not discharged on beta-blockers", cex = 0.8)
```

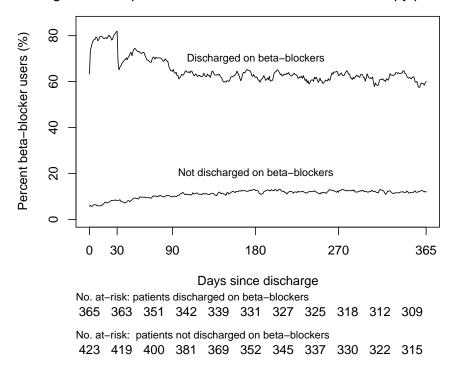


Figure 1. Outpatient adherence to beta-blocker therapy post-AM

NOTES: (1) the oma argument of the par function allows you to change the size of the outer margins of the plot given in lines of text (the order is bottom, left, top, right); (2) the axis function adds and axis to a plot, allowing the speciation of the side, position, label, and other options (corresponds to setting axes = F in the plot command); (3) the lines function adds a line to a plot; (4) the box function draws a box around a plot; (5) the mtext function writes text into the margins of a plot; and (6) the text function writes text inside a plot

#### 10.7 Different page layouts

Unfortunately, the LaTex interface I used to create this pdf file had problems placing the following plot in this pdf file, but we can still run the code in R and view the resulting plot.

```
albhist<-hist(pbc$albumin, plot=FALSE)
obtimehist<-hist(pbc$obstime, plot=FALSE)
def.par <- par(no.readonly = TRUE) # save default, for resetting...
layout(matrix(c(2,0,1,3),2,2,byrow=TRUE), widths=c(3,1),
    heights=c(1,3), respect=TRUE)
plot(pbc$albumin, pbc$obstime, xlab="Serum Albumin",
    ylab="Observation Time", main="Observation Time by Serum Albumin")
barplot(albhist$counts, main="Serum Albumin",
    space=0)
barplot(obtimehist$counts, horiz=TRUE, main="Observation Time",
    space=0)
par(def.par)#- reset to default</pre>
```

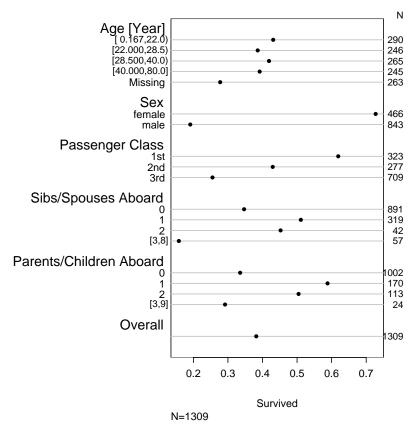
### 10.8 Graphical Data Summary

The titanic3 data set contains information on N = 1309 passengers from the *Titanic*. Chapter 12 in Frank Harrell's *Regression Modelling Strategies* develops a binary logistic regression model to describe the patterns of survival in these passengers, based on passenger age, sex, ticket class, and the number of family members accompanying each passenger.<sup>8</sup>

```
> library(Hmisc)
> library(Design)
> getHdata(titanic3)
> x <- titanic3[Cs(pclass, survived, age, sex, sibsp, parch)]</pre>
> x <- upData(x, labels = c(sex = "Sex", pclass = "Passenger Class",
      sibsp = "Sibs/Spouses Aboard", parch = "Parents/Children Aboard"))
Input object size:
                             54396 bytes;
                                                   6 variables
New object size:
                          54604 bytes;
                                                 6 variables
> dd <- datadist(x)
> options(datadist = "dd")
> titanic.summ <- summary(survived ~ age + sex + pclass + cut2(sibsp,</pre>
      0:3) + cut2(parch, 0:3), data = x)
+
> latex(titanic.summ, file = "")
> par(cex = 0.85)
> plot(titanic.summ, main = "Univariable Summaries of Titanic Survival")
  <sup>8</sup>Regression Modelling Strategies, Harrell
```

Table 3: Survived N=1309

	Ν	survived
Age Year		
[0.167, 22.0)	290	0.43
[22.000, 28.5)	246	0.39
[28.500, 40.0)	265	0.42
[40.000, 80.0]	245	0.39
Missing	263	0.28
Sex		
female	466	0.73
male	843	0.19
Passenger Class		
1st	323	0.62
2nd	277	0.43
3rd	709	0.26
Sibs/Spouses Aboard		
0	891	0.35
1	319	0.51
2	42	0.45
[3,8]	57	0.16
Parents/Children Aboard		
0	1002	0.34
1	170	0.59
2	113	0.50
[3,9]	24	0.29
Overall		
	1309	0.38



#### Univariable Summaries of Titanic Survival

#### 10.9 Summarizing/Describing the Fitted Model

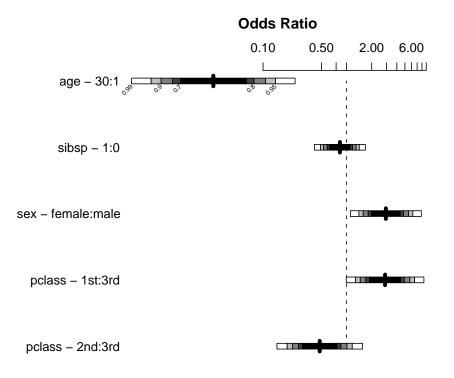
It is always important for the analyst to present and interpret a fitted model, once the proper variables have been modelled and all assumptions have been met. The coefficients in the model may be interpreted by computing, for each variable, the change in log odds for a sensible change in the variable value (e.g. interquartile range).<sup>9</sup>

> library(Hmisc) > library(Design) > dd <- datadist(x) > options(datadist = "dd") > titanic.model <- lrm(survived ~ (sex + pclass + rcs(age, 5))^2 + + rcs(age, 5) \* sibsp, data = x)

<sup>&</sup>lt;sup>9</sup>Regression Modelling Strategies, Harrell

> titanic.model.summ <- summary(titanic.model, age = c(1, 30), + sibsp = 0:1)

> plot(titanic.model.summ, log = T)



Adjusted to:sex=male pclass=3rd age=28 sibsp=0

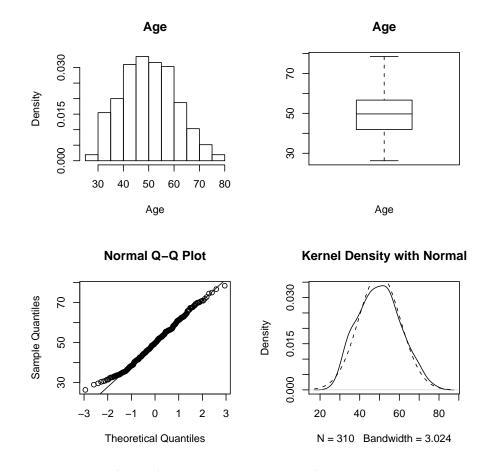
# 11 Writing Your Own Functions

- Perhaps one of the best features of R is its capability of writing your own functions
- Writing your own functions will become very useful when you find yourself executing the same set of commands (e.g. finding the mean and standard deviation, or plotting the same general plot) repeated times
- Some Examples:

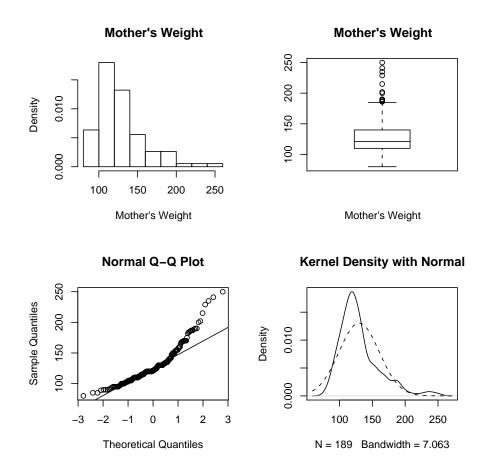
1. A function that prints out the mean and standard deviation of a set of numbers:<sup>10</sup>

```
> mean.and.sd <- function(x) {</pre>
  +
         av < -mean(x)
         stdev <- sd(x)
  +
         c(mean = av, SD = stdev)
  + }
  > mean.and.sd(1:10)
                   SD
      mean
  5.500000 3.027650
  > mean.and.sd(lowbw$age)
        mean
                     SD
  23.238095 5.298678
2. A functions that generates four plots for a continuous variable: (1)
  a histogram, (2) a boxplot, (3) a normal Q-Q plot, and (4) a Kernel
  Density plot:
  > dist.shape <- function(xdata, varname) {</pre>
         par(mfrow = c(2, 2))
  +
         hist(xdata, freq = F, main = varname, xlab = varname)
         boxplot(xdata, main = varname, xlab = varname)
   +
         qqnorm(xdata)
         qqline(xdata)
         dx <- density(xdata)</pre>
         xbar <- mean(xdata)</pre>
         stdev <- sd(xdata)</pre>
         rangex <- range(dx$x)</pre>
         xx <- seq(rangex[1], rangex[2], length = length(dx$y))</pre>
   +
         plot(dx, main = "Kernel Density with Normal")
         lines(xx, dnorm(xx, xbar, stdev), lty = 2)
  +
  + }
  > dist.shape(pbc$age, "Age")
```

 $<sup>^{10}</sup>Data$  Analysis and Graphics Using R, Maindonald and Braun



> dist.shape(lowbw\$lwt, "Mother's Weight")



3. A function that finds high and low outliers (greater than +/-4 S.D. of the mean) of one or more continuous variables (assumes first column of data frame represents an ID):

```
> out <- function(dataframe, colnames, idname) {</pre>
      data <- dataframe[, colnames]</pre>
+
      I <- dim(data)[2]</pre>
+
      for (i in 1:I) {
+
           colm <- data[!is.na(data[[i]]), i]</pre>
+
           colname <- colnames[[i]]</pre>
+
           con <- 4
           bottom <- mean(colm) - con * sd(colm)</pre>
           top <- mean(colm) + con * sd(colm)</pre>
+
+
           nhigh <- length(colm[colm > top])
+
           nlow <- length(colm[colm < bottom])</pre>
+
           if (nlow > 0) {
+
                lowouts <- dataframe[colm < bottom & !is.na(data[[i]]),</pre>
```

```
c(idname, colname)]
+
             cat("LOW OUTLIERS", "\t", "( <", bottom, ")", "\n",</pre>
+
                  "ID", "\t", colname, "\n", file = "outliers.txt",
                  append = T)
+
+
             J <- dim(lowouts)[1]</pre>
             K <- dim(lowouts)[2]</pre>
+
             for (j in 1:J) {
                  for (k in 1:K) {
                    cat(paste(lowouts[j, k]), "\t", file = "outliers.txt",
                      append = T)
                   if (k == K)
                      cat("\n", file = "outliers.txt", append = T)
+
                 }
+
             }
             cat("-----", "\n".
+
                  "\n", file = "outliers.txt", append = T)
+
         }
+
         if (nhigh > 0) {
             highouts <- dataframe[colm > top & !is.na(data[[i]]),
+
+
                  c(idname, colname)]
             cat("HIGH OUTLIERS", "\t", "( >", top, ")", "\n",
+
                  "ID", "\t", colname, "\n", file = "outliers.txt",
                  append = T)
             J <- dim(highouts)[1]</pre>
             K <- dim(highouts)[2]</pre>
             for (j in 1:J) {
                  for (k in 1:K) {
+
                    cat(paste(highouts[j, k]), "\t", file = "outliers.txt",
                      append = T)
                    if (k == K)
                      cat("\n", file = "outliers.txt", append = T)
                 }
+
             }
             cat("-----", "\n",
+
                  "\n", file = "outliers.txt", append = T)
+
         }
+
     }
+
+ }
> out(lowbw, c("age", "lwt", "bwt"), "id")
```

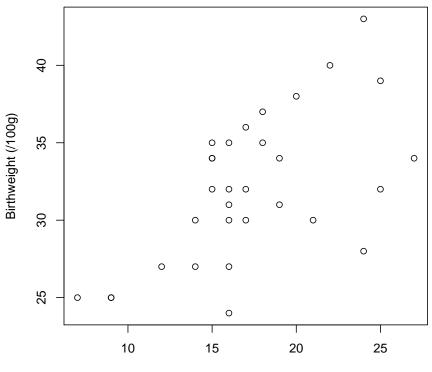
# 12 Statistics with R

### 12.1 Correlation

Example: Estriol and Birthweight

Consider a study investigating birthweight (/100 g) and estriol levels (mg/24hr) in pregnant women (estriol.dta)

```
> library(foreign)
> est <- read.dta("estriol.dta")
> names(est)
[1] "estriol" "birthwt"
> dim(est)
[1] 31 2
> plot(est$estriol, est$birthwt, main = "Birthweight by Estriol",
+ xlab = "Estriol (mg/24hr)", ylab = "Birthweight (/100g)")
```



**Birthweight by Estriol** 

Estriol (mg/24hr)

12.1.1 Pearson Correlation & Testing for Association

```
> cor(est)
         estriol birthwt
estriol 1.0000000 0.6097313
birthwt 0.6097313 1.0000000
> cor.test(est$estriol, est$birthwt)
       Pearson's product-moment correlation
data: est$estriol and est$birthwt
t = 4.1427, df = 29, p-value = 0.0002712
alternative hypothesis: true correlation is not equal to 0
95 percent confidence interval:
0.3257757 0.7927878
sample estimates:
     cor
0.6097313
12.1.2 Spearman Rank Correlation
> library(Hmisc)
> rcorr(est$estriol, est$birthwt, type = "spearman")
    х
         у
x 1.00 0.56
y 0.56 1.00
```

n= 31

# P x y x 0.001 y 0.001 > rcorr(est\$estriol, est\$birthwt, type = "pearson") x y x 1.00 0.61 y 0.61 1.00

```
n= 31
```

Ρ

x y x 3e-04 y 3e-04

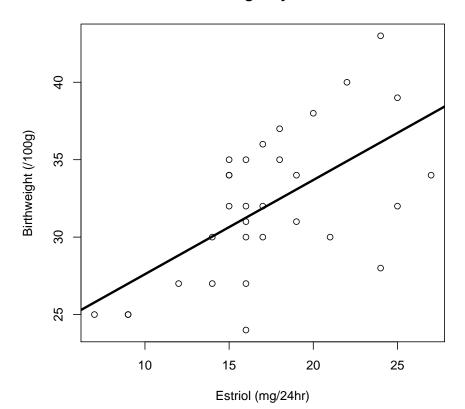
# 12.2 Simple Linear Regression

### Example: Estriol and Birthweight

```
Consider regressing birthweight on estriol.
> est.m <- lm(birthwt ~ estriol, data = est)</pre>
> summary(est.m)
Call:
lm(formula = birthwt ~ estriol, data = est)
Residuals:
    Min
              1Q Median
                                ЗQ
                                         Max
-8.12000 -2.03810 -0.03810 3.35371 6.88000
Coefficients:
            Estimate Std. Error t value Pr(>|t|)
(Intercept) 21.5234 2.6204 8.214 4.68e-09 ***
estriol
            0.6082
                         0.1468 4.143 0.000271 ***
___
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Residual standard error: 3.821 on 29 degrees of freedom
Multiple R-Squared: 0.3718,
                                  Adjusted R-squared: 0.3501
F-statistic: 17.16 on 1 and 29 DF, p-value: 0.0002712
```

### 12.2.1 Interpretation of Output

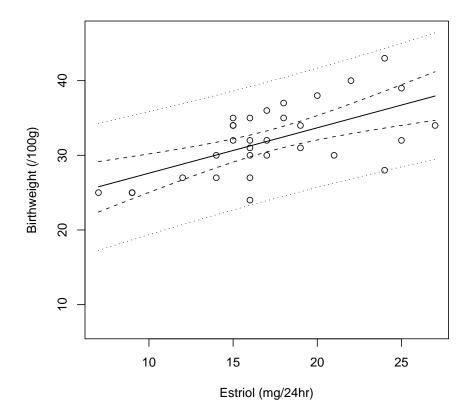
```
> plot(est$estriol, est$birthwt, main = "Birthweight by Estriol",
+ xlab = "Estriol (mg/24hr)", ylab = "Birthweight (/100g)")
> abline(est.m, lwd = 3)
```



**Birthweight by Estriol** 

12.2.2 Plotting 95% Confidence Intervals & Prediction Intervals

```
> pred.frame <- data.frame(estriol = 7:27)
> pp <- predict(est.m, int = "p", newdata = pred.frame)
> pc <- predict(est.m, int = "c", newdata = pred.frame)
> pred.estriol <- pred.frame$estriol
> plot(est$estriol, est$birthwt, ylim = range(est$estriol, pp,
+ na.rm = T), main = "Plot with Confidence and Tolerance Bands",
+ xlab = "Estriol (mg/24hr)", ylab = "Birthweight (/100g)")
> matlines(pred.estriol, pc, lty = c(1, 2, 2), col = "black")
> matlines(pred.estriol, pp, lty = c(1, 3, 3), col = "black")
```



# **Plot with Confidence and Tolerance Bands**

12.2.3 Linear Regression for Categorical Variables

### Example: Low Birthweight Study

We would like to determine whether there is a difference in mean birthweight for those with no previous pre-term labor and those with 1+ pre-term labors.

```
> lowbw$ptl.cat <- ifelse(lowbw$ptl > 0, 1, 0)
> names(lowbw)
 [1] "id"
                "low"
                           "age"
                                      "lwt"
                                                "race"
                                                           "smoke"
                                                                      "ptl"
 [8] "ht"
                "ui"
                           "ftv"
                                      "bwt"
                                                "ptl.cat"
> table(lowbw$ptl)
  0
          2
               3
      1
159
    24
          5
               1
```

```
> table(lowbw$ptl.cat)
 0
    1
159 30
> lowbw.m <- lm(bwt ~ ptl.cat, data = lowbw)</pre>
> summary(lowbw.m)
Call:
lm(formula = bwt ~ ptl.cat, data = lowbw)
Residuals:
Birthweight [grams]
     Min 1Q Median
                                    ЗQ
                                             Max
-1992.572 -495.400 -8.572 586.428 1976.428
Coefficients:
           Estimate Std. Error t value Pr(>|t|)
(Intercept) 3013.57 56.57 53.269 < 2e-16 ***
                       142.00 -3.058 0.00256 **
            -434.17
ptl.cat
___
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Residual standard error: 713.4 on 187 degrees of freedom
Multiple R-Squared: 0.04761,
                                  Adjusted R-squared: 0.04252
F-statistic: 9.349 on 1 and 187 DF, p-value: 0.002558
> t.test(bwt ~ ptl.cat, data = lowbw, var.equal = T, alternative = "two.sided")
       Two Sample t-test
data: bwt by ptl.cat
t = 3.0576, df = 187, p-value = 0.002558
alternative hypothesis: true difference in means is not equal to 0
95 percent confidence interval:
154.0518 714.2929
sample estimates:
mean in group 0 mean in group 1
      3013.572
                      2579.400
> t.test(bwt ~ ptl.cat, data = lowbw, var.equal = T, alternative = "less")
       Two Sample t-test
data: bwt by ptl.cat
t = 3.0576, df = 187, p-value = 0.9987
alternative hypothesis: true difference in means is less than 0
```

```
95 percent confidence interval:
     -Inf 668.8983
sample estimates:
mean in group 0 mean in group 1
       3013.572
                       2579.400
> t.test(bwt ~ ptl.cat, data = lowbw, var.equal = T, alternative = "greater")
        Two Sample t-test
data: bwt by ptl.cat
t = 3.0576, df = 187, p-value = 0.001279
alternative hypothesis: true difference in means is greater than 0
95 percent confidence interval:
 199.4463
               Inf
sample estimates:
mean in group 0 mean in group 1
       3013.572
                       2579.400
```

# 12.3 Simple Logistic Regression

### Example: Estriol and Birthweight

Suppose we wish to relate estricle level to the probability of having a low birthweight infant, where a low birthweight indicator in which birthweight <3000 grams indicates low birthweight.

```
> est$birthwt.ind <- ifelse(est$birthwt < 30, 1, 0)</pre>
> table(est$birthwt.ind)
 0
   1
23 8
> library(Hmisc)
> library(Design)
> dd <- datadist(est)</pre>
> options(datadist = "dd")
> est.birthwt.m <- lrm(birthwt.ind ~ estriol, data = est)</pre>
> summary(est.birthwt.m)
             Effects
                                   Response : birthwt.ind
 Factor
             Low High Diff. Effect S.E. Lower 0.95 Upper 0.95
 estriol
             15 19.5 4.5
                             -1.56 0.7
                                         -2.93
                                                     -0.20
 Odds Ratio 15 19.5 4.5
                              0.21
                                           0.05
                                                      0.82
                                     NA
> anova(est.birthwt.m)
```

Wald S	Stati	stics
--------	-------	-------

Response: birthwt.ind

Factor	Chi-Square	d.f.	Р
estriol	5.03	1	0.0249
TOTAL	5.03	1	0.0249

### 12.4 Simple Proportional Hazards Regression

### Example: Primary Biliary Cirrhosis (PBC) Trial

We would like to investigate the role of d-penicillamine (DPCA) for treating PBC on the patients' survival.

### 12.4.1 Kaplan-Meier Estimates

> library(survival)

First lets just look at patient survival, regardless of treatment.

```
> pbc.surv <- survfit(Surv(obstime, status == "Died"), data = pbc)</pre>
> pbc.surv
Call: survfit(formula = Surv(obstime, status == "Died"), data = pbc)
        n
             events
                         rmean se(rmean)
                                             median
                                                       0.95LCL
                                                                  0.95UCL
      310
                 125
                          2971
                                      102
                                                3358
                                                          3086
                                                                     3853
> summary(pbc.surv)
Call: survfit(formula = Surv(obstime, status == "Died"), data = pbc)
 time n.risk n.event survival std.err lower 95% CI upper 95% CI
   41
         310
                    1
                         0.997 0.00322
                                               0.990
                                                              1.000
   51
         309
                    1
                         0.994 0.00455
                                               0.985
                                                              1.000
   71
         308
                    1
                         0.990 0.00556
                                                0.979
                                                              1.000
   77
         307
                    1
                         0.987 0.00641
                                                0.975
                                                              1.000
  110
         306
                    1
                         0.984 0.00715
                                                0.970
                                                              0.998
  130
         305
                         0.981 0.00782
                                                0.965
                                                              0.996
                    1
  131
         304
                    1
                         0.977 0.00844
                                                0.961
                                                              0.994
  140
         303
                         0.974 0.00901
                                                0.957
                                                              0.992
                    1
  179
         302
                    1
                         0.971 0.00954
                                                0.952
                                                              0.990
  186
         301
                    1
                         0.968 0.01004
                                                0.948
                                                              0.988
                         0.488 0.04022
 3395
          43
                    1
                                               0.415
                                                              0.573
 3428
                         0.476 0.04096
                                                0.402
          41
                    1
                                                              0.563
 3445
          40
                    1
                         0.464 0.04163
                                               0.389
                                                              0.553
```

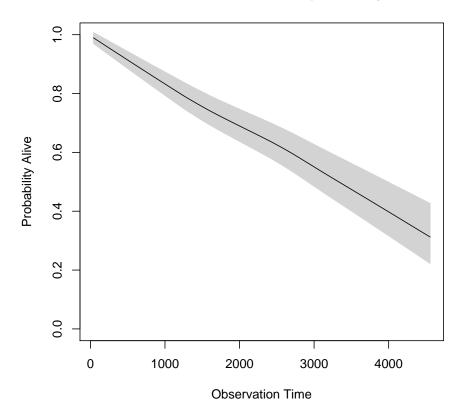
3574	37	1	0.451	0.04235	0.376	0.543
3584	34	1	0.438	0.04314	0.361	0.531
3762	30	1	0.424	0.04410	0.345	0.519
3839	27	1	0.408	0.04517	0.328	0.507
3853	25	1	0.392	0.04622	0.311	0.494
4079	17	1	0.369	0.04891	0.284	0.478
4191	13	1	0.340	0.05272	0.251	0.461
> plot(r	bc.surv,	main =	"Kapla	an-Meier	Survival Estimates",	<pre>xlab = "Observation Time")</pre>

# Kaplan–Meier Survival Estimates

The following plot shows the estimated (smoothed) distribution of time patients stayed alive. The shaded area represents the 95% confidence interval. A similar plot was shown in an article published in the Journal of the American Medical Association (JAMA, June 20, 2001 – Vol 285, No. 23).

```
> plot(pbc.surv$time, pbc.surv$surv, type = "n", ylim = c(0, 1),
+ xlab = "Observation Time", ylab = "Probability Alive", main = "Estimated Distbn of Tim
> polygon(xx, yy, col = "lightgray", border = "lightgray")
```

```
> lines(lowess(pbc.surv$time, pbc.surv$surv))
```

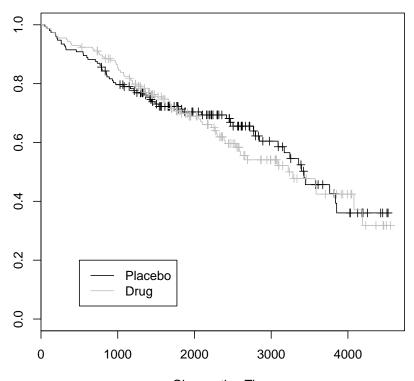


## **Estimated Distbn of Time Participants Stayed Alive**

Now let's look at the role of treatment on patient survival.

```
> pbc.surv.tx <- survfit(Surv(obstime, status == "Died") ~ tx,</pre>
      data = pbc)
> pbc.surv.tx
Call: survfit(formula = Surv(obstime, status == "Died") ~ tx, data = pbc)
             n events rmean se(rmean) median 0.95LCL 0.95UCL
tx=Placebo 153
                   60
                       2987
                                   144
                                         3428
                                                 3090
                                                           Inf
tx=Drug
           157
                   65
                      2947
                                   142
                                         3282
                                                 2583
                                                           Inf
> summary(pbc.surv.tx)
```

Call:	survfit	t(formula	a = Surv(	obstime,	status == "Di	.ed") ~ tx, data	= pbc)		
		tx=P:	lacebo						
time	n.risk	n.event	survival	<pre>std.err</pre>	lower 95% CI	upper 95% CI			
51	153	1	0.993	0.00651	0.981	1.000			
77	152	1	0.987	0.00918	0.969	1.000			
110	151	1	0.980	0.01121	0.959	1.000			
130	150			0.01290		0.999			
186	149	1	0.967	0.01437	0.940	0.996			
•									
•									
•									
3428				0.05977		0.612			
3445				0.06113		0.594			
3762				0.06419	0.317	0.573			
3839				0.06711		0.550			
3853	12	1	0.361	0.06907	0.248	0.525			
		+D-							
+ imo		tx=Di	•	atd orr	1	05% CT			
					lower 95% CI				
41				0.00635	0.981	1.000			
71			0.987			1.000			
131			0.981			1.000			
140			0.975			0.999			
179	153	T	0.968	0.01401	0.941	0.996			
·									
•									
3282	22	1	0.477	0.05492	0.381	0.598			
3574				0.05792		0.580			
3584				0.06028		0.561			
4079				0.07242					
4191				0.07242	0.195	0.518			
1101		-	0.010	0.01010	0.100	0.010			
> plot	t(pbc.sı	urv.tx,	col = c("	black",	"gray"), main	= "Kaplan-Meier	Survival	Estimates	by Tre
+	xlab =	"Observa	ation Time	e")					
<pre>&gt; legend(500, 0.2, legend = c("Placebo", "Drug"), col = c("black",</pre>									
+	"gray")	), lty =	c(1, 1))		-				



# Kaplan–Meier Survival Estimates by Treatment

**Observation Time** 

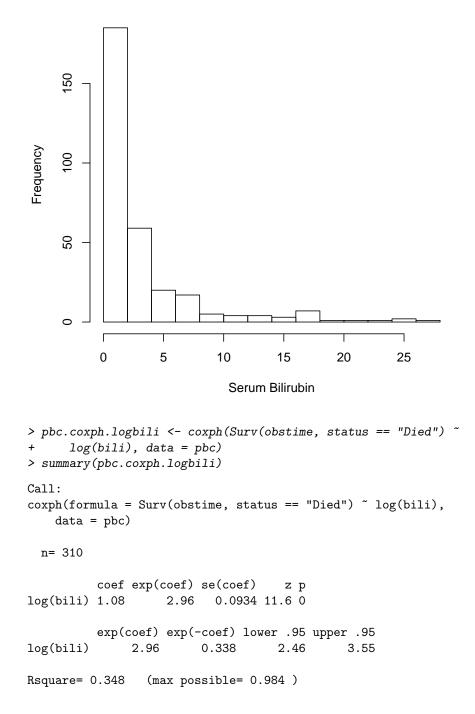
### 12.4.2 Log-Rank Test

```
> library(survival)
> survdiff(Surv(pbc$obstime, pbc$status == "Died") ~ pbc$tx)
Call:
survdiff(formula = Surv(pbc$obstime, pbc$status == "Died") ~
    pbc$tx)
                  N Observed Expected (O-E)<sup>2</sup>/E (O-E)<sup>2</sup>/V
pbc$tx=Placebo 153
                           60
                                  61.8
                                           0.0507
                                                       0.100
pbc$tx=Drug
                157
                           65
                                  63.2
                                           0.0495
                                                       0.100
 Chisq= 0.1 on 1 degrees of freedom, p= 0.751
```

12.4.3 Cox Proportional Hazards Regression

```
> library(survival)
> pbc.coxph.tx <- coxph(Surv(obstime, status == "Died") ~ tx, data = pbc)</pre>
> summary(pbc.coxph.tx)
Call:
coxph(formula = Surv(obstime, status == "Died") ~ tx, data = pbc)
 n= 310
        coef exp(coef) se(coef) z p
txDrug 0.0568 1.06 0.179 0.317 0.75
      exp(coef) exp(-coef) lower .95 upper .95
txDrug
        1.06
                0.945
                           0.745
                                     1.50
Rsquare= 0 (max possible= 0.984 )
Likelihood ratio test= 0.1 on 1 df, p=0.751
Wald test
            = 0.1 on 1 df, p=0.751
Score (logrank) test = 0.1 on 1 df, p=0.751
> hist(pbc$bili, main = "Histogram of Serum Bilirubin", xlab = "Serum Bilirubin")
```

# Histogram of Serum Bilirubin



Likelihood ratio test= 133	s on 1 df, p=0	
Wald test = 135	on 1 df, p=0	
Score (logrank) test = 159	on 1 df, p=0	