# HealthyR: R for healthcare data analysis

Never trust a data scientist - they are always plotting.

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#### Preface

Version 0.3.1

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#### Why read this book

We are drowning in information but starved for knowledge. John Naisbitt  $\,$ 

In this age of information, the manipulation, analysis and interpretation of data has become paramount. Nowhere more so than in the delivery of healthcare. From the understanding of disease and the development of new treatments, to the diagnosis and management of individual patients, the use of data and technology is now an integral part of the business of healthcare.

Those working in healthcare interact daily with data, often without realising it. The conversion of this avalanche of information to xvi Preface

useful knowledge is essential for high quality patient care. An important part of this information revolution is the opportunity for everybody to become involved in data analysis. This democratisation of data analysis is driven in part by the open source software movement – no longer do we require expensive specialised software to do this.

The statistical programming language, R, is firmly at the heart of this!

This book will take an individual with little or no experience in data analysis all the way through to performing sophisticated analyses. We emphasise the importance of understanding the underlying data with liberal use of plotting, rather than relying on opaque and possibly poorly understand statistical tests. There are numerous examples included that can be adapted for your own data, together with our own R packages with easy-to-use functions.

We have a lot of fun teaching this course and focus on making the material as accessible as possible. We banish equations in favour of code and use examples rather than lengthy explanations. We are grateful to the many individuals and students who have helped refine these and welcome suggestions and bug reports via https://github.com/SurgicalInformatics.

Ewen Harrison and Riinu Ots

August 2019

#### Structure of the book

Chapters ?? introduces a new topic, and ...

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#### Software information and conventions

I used the **knitr** package (Xie, 2015) and the **bookdown** package (Xie, 2018) to compile my book. My R session information is shown below:

```
xfun::session_info()
## R version 3.4.4 (2018-03-15)
## Platform: x86_64-pc-linux-gnu (64-bit)
## Running under: Ubuntu 16.04.5 LTS
##
## Locale:
     LC_CTYPE=en_GB.UTF-8
##
     LC NUMERIC=C
##
     LC TIME=en GB.UTF-8
##
     LC_COLLATE=en_GB.UTF-8
##
     LC_MONETARY=en_GB.UTF-8
##
     LC_MESSAGES=en_GB.UTF-8
##
     LC_PAPER=en_GB.UTF-8
##
     LC NAME=C
##
     LC_ADDRESS=C
##
     LC TELEPHONE=C
##
     LC MEASUREMENT=en GB.UTF-8
##
     LC_IDENTIFICATION=C
##
## Package version:
##
     base64enc_0.1.3
                       bookdown_0.7
                                         compiler_3.4.4
##
     digest_0.6.20
                       evaluate_0.13
                                         glue_1.3.1
##
     graphics_3.4.4
                       grDevices_3.4.4
                                        highr_0.8
##
     htmltools 0.3.6
                       jsonlite 1.6
                                        knitr 1.22
##
     magrittr_1.5
                       markdown 0.9
                                        methods_3.4.4
##
     mime_0.6
                      Rcpp_1.0.1
                                        rmarkdown_1.12.4
##
     stats_3.4.4
                       stringi_1.4.3
                                        stringr_1.4.0
##
     tinytex 0.11
                       tools 3.4.4
                                        utils 3.4.4
```

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```
## xfun_0.6 yaml_2.2.0
```

Package names are in bold text (e.g., **rmarkdown**), and inline code and filenames are formatted in a typewriter font (e.g., knitr::knit('foo.Rmd')). Function names are followed by parentheses (e.g., bookdown::render\_book()).

#### Acknowledgments

A lot of people helped me when I was writing the book.

Frida Gomam on the Mars

#### Installation

• Download R

```
https://www.r-project.org/
```

• Install RStudio

```
https://www.rstudio.com/products/rstudio/
```

• Install packages (copy these lines into the Console in RStudio):

```
install.packages("tidyverse")
install.packages("gapminder")
install.packages("gmodels")
install.packages("Hmisc")
```

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```
install.packages("devtools")

devtools::install_github("ewenharrison/finalfit")

install.packages("pROC")

install.packages("survminer")
```

When working with data, don't copy or type code directly into the Console. We will only be using the Console for viewing output, warnings, and errors (and installing packages as in the previous section). All code should be in a script and executed (=Run) using Control+Enter (line or section) or Control+Shift+Enter (whole script). Make sure you are always working in a project (the right-top corner of your RStudio interface should say "HealthyR").



# Part I Data wrangling and

visualisation

### Your first R plots

In this session, we will create five beautiful and colourful barplots in less than an hour. Do not worry about understanding every single word or symbol (e.g. the pipe - %>%) in the R code you are about to see. The purpose of this session is merely to

- gain familiarity with the RStudio interface:
  - to know what a script looks like,
  - what is the Environment tab,
  - where do your plots appear.

#### 1.1 Data

Load the example dataset which is already saved as an R-Data file (recognisable by the file extension .rda or .RData):

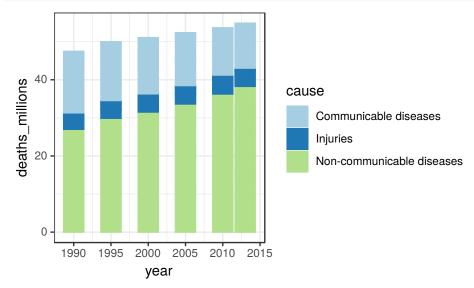
```
library(ggplot2)
source("1_source_theme.R")
load("global_burden_disease_long.rda")
```

After loading the datasets, investigate your Environment tab (top-right). You will see two things listed: mydata and mydata2013, which is a subset of mydata.

Click on the name mydata and it will pop up next to where your

script is. Clicking on the blue button is not as useful (in this session), but it doesn't do any harm either. Try it.

#### 1.2 First plot



ggplot() stands for grammar of graphics plot - a user friendly yet flexible alternative to plot().

aes() stands for aesthetics - things we can see.

geom () stands for geometric.

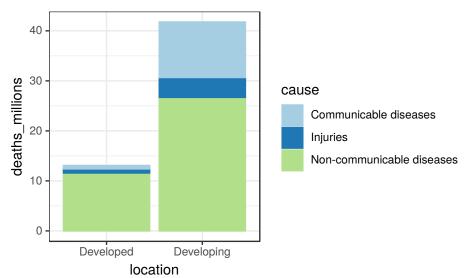
1.2 First plot 5

#### 1.2.1 Question

Why are there two closing brackets - )) - after the last aesthetic (colour)?

#### 1.2.2 Exercise

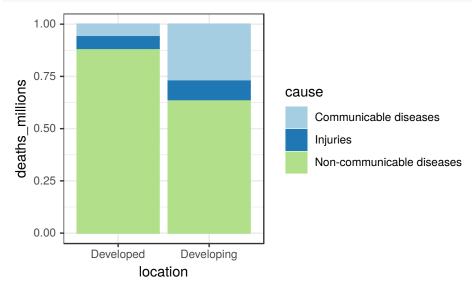
Plot the number of deaths in Developed and Developing countries for the year 2013:



#### 1.3 Comparing bars of different height

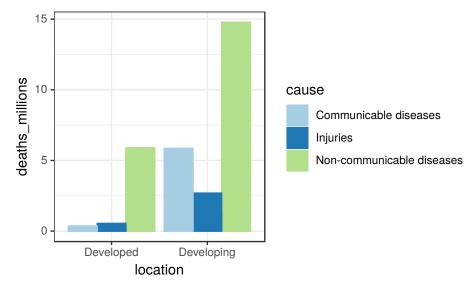
#### 1.3.1 Stretch each bar to 100%

position="fill" stretches the bars to show relative contributions:



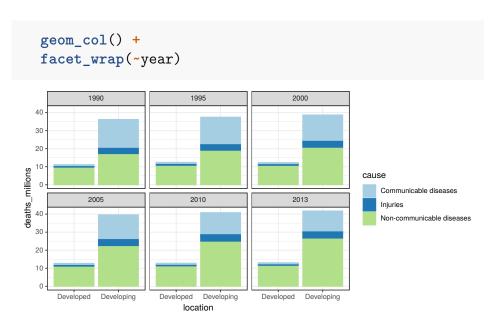
#### 1.3.2 Plot each bar next to each other

position="dodge" puts the different causes next to each rather
(the default is position="stack"):



#### 1.4 Facets (panels)

Going back to the dataframe with all years (1990 – 2015), add facet\_wrap(~year) to plot all years at once:

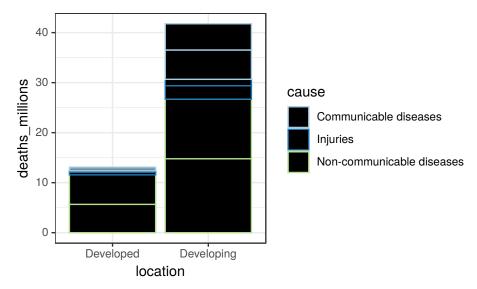


#### 1.5 Extra: using aethetics outside of the aes()

#### 1.5.1 Setting a constant fill

Using the mydata2013 example again, what does the addition of fill = "black" in this code do? Note that putting the ggplot(aes()) code all on one line does not affect the result.

```
mydata2013 %>%
   ggplot(aes(x = location, y = deaths_millions, fill = cause, colour = cause
   geom_col(fill = "black")
```



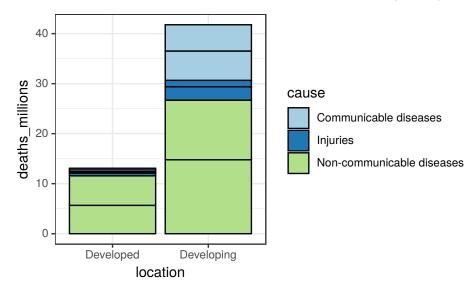
Setting aesthetics (x, y, fill, colour, etc.) outside of aes() sets them to a constant value. R can recognise of a lot of colour names, e.g., try "cornflowerblue", "firebrick", or just "red", "green", "blue", etc. For a full list, search Google for "Colours in R". R also knows HEX codes, e.g. fill = "#fec3fc" is pink.

#### 1.5.2 Exercise

What is the difference between colour and fill in the context of a barplot?

Hint: Use colour = "black" instead of fill = "black" to investigate what ggplot() thinks a colour is.

```
mydata2013 %>%
   ggplot(aes(x = location, y = deaths_millions, fill = cause, colour = cause
   geom_col(colour = "black")
```



#### 1.5.3 Exercise

Why are some of the words in our code quoted (e.g. fill = "black") whereas others are not (e.g. x = location)?

#### 1.6 Two geoms for barplots: geom\_bar() or geom\_col()

Both geom bar() and geom col() create barplots. If you:

- Want to visualise the count of different lines in a dataset use geom\_bar()
  - For example, if you are using a patient-level dataset (each line is a patient record): mydata %>% ggplot(aes(x = sex)) + geom\_bar()
- Your dataset is already summarised use geom\_col()
  - For example, in the GBD dataset we use here, each line already includes a summarised value (deaths\_millions)

If you have used R before you might have come across geom\_bar(stat = "identity") which is the same as geom\_col().

1.7 Solutions 11

#### 1.7 Solutions

1.2.1: There is a double closing bracket because aes() is wrapped inside ggplot() - ggplot(aes()).

#### 1.2.2:

#### 1.5.2:

On a barplot, the colour aesthetic outlines the fill. In a later session we will see, however, that for points and lines, colour is the main aesthetic to define.

#### 1.5.3:

Words in quotes are generally something set to a constant value (e.g. make all outlines black, rather than colour them based on the cause they are representing). Unquoted words are generally variables (or functions). If the word "function" just threw you, Google "Jesse Maegan: What the h\*ck is a function"

#### R Basics

The aim of this module is to familiarise you with how R works. We will read in data and start basic manipulations. We will be working with a shorter version of the Global Burden of Disease dataset that we met earlier.

#### 2.1 Getting help

RStudio has a built in Help tab. To use the Help tab, click your cursor on something in your code (e.g. read\_csv()) and press F1. This will show you the definition and some examples. However, the Help tab is only useful if you already know what you are looking for but can't remember exactly how it works. For finding help on things you have not used before, it is best to Google it. R has about 2 million users so someone somewhere has had the same question or problem.

14 2 R Basics

**TABLE 2.1:** Example of a table (=tibble once read into R), including missing values - denoted NA (Not applicable/Not available).

id	sex	var1	var2	var3
1	Male	4	NA	2
2	Female	1	4	1
3	Female	2	5	NA
4	Male	3	NA	NA

#### 2.2 Objects and functions

The two fundamental concepts to understand about statistical programming are objects and functions. As usual, in this book, we prefer introducing new concepts using specific examples first. And then define things in general terms after examples.

The most common data object you will be working with is a table - so something with rows and columns. It should be regular, e.g., the made-up example in Table 2.1.

Now, regular does not mean it can't have missing values. Missing values are denoted NA that stands for either Not available or Not applicable. In same contexts, these things can have a different meaning, for example:

Since var2 is NA for all Male subjects, it may mean "Not applicable", i.e. something that can only be measured in females. Whereas in var3, NA is more likely to mean "Not available" so real missing data, e.g. lost to follow-up. We will come back to handling missing values later, but let's return to introducing objects and functions.

A table can live anywhere: on paper, in a Spreadsheet, in an SQL database, or it can live in R session's Environment. And yes, R sessions are as fun as they sound, almost as fun as, e.g., music sessions. We usually initiate and interface R using RStudio, but everything

we talk about here (objects, functions, sessions, environment) also work when RStudio is not available, but R is. This can be the case if you are working on a supercomputer that can only serve the R Console, and not an RStudio IDE (reminder from first chapter: Integrated Development Environment). So, regualarly shaped data in rows and columns is called table when it lives outside R. Once you read it into R (import it), we call it a tibble. When you are in one of your very cool R sessions and read in a table, it goes into this session's Environment. There used to be an older version of tables in R - they are called data frames. In most cases, data frames and tibbles work interchangeably (and both are R objects), but tibbles are newer and better. Another great alterntative to base R's data frames are data tables. In this book, and for most of our day-to-day work these days, we only use tibbles.

So, the tibble mydata example of an object that lives in the Environment of your R Session. A function that you may want to apply on numeric data is mean(). R functions always have brackets after their name. This is for two reasons. First, to easily differentiate them from objects - which don't have brackets after their name. Second, and more important, we can put arguments in these brackets. Arguments can also be thought of as input, and in data analysis, the most common input for a function is data: we need to give the R function mean() some data to average. It does not make sense (nor will it work) to feed it the whole tibble that has multiple columns, including patient IDs and a categorical variable (sex). To quickly extract a single column, we use the \$ symbol, like this:

#### mydata\$var1

#### ## [1] 4 1 2 3

You can ignore the ## [1] at the beginning of the extracted values this is something that becomes more useful when printing multiple lines of data, the number in the square brackets keeps count on how many values we are seeing. 16 2 R Basics

We can then use mydata\$var1 as the first argument of mean() by putting it inside its brackets:

```
mean(mydata$var1)
```

#### ## [1] 2.5

Which tells us that the mean of var1 (4, 1, 2, 3) is 2.5. In this example, mydata\$var1 is the first and only argument to mean(). But what happens if we try to calculate the average value of var2 (NA, 4, 5, NA)?

```
mean(mydata$var2)
```

#### ## [1] NA

We get an NA ("Not applicable"). We would expect to see an NA if we tried to, for example, calculate the average of sex:

```
mean(mydata$sex)
```

## Warning in mean.default(mydata\$sex): argument is not numeric or logical:
## returning NA

#### ## [1] NA

In fact, R then gives as us a pretty clear Error (our first Error in this book!) saying it can't compute the mean of an argument that is not numeric or logical. The sentence actually reads pretty fun, as if R was saying it was not logical to calculate the mean of something that is not numeric. But what R is actually saying that it is happy to calculate the mean of two types of variables: numerics or logicals which is a data type with just two potential values TRUE/FALSE. We will come back to data types shortly.

So mean(mydata\$var2) does not return an Error, but it also doesn't return the mean of the numeric values included in this column. That is because the column includes missing values (NAs),

and R does not want to average over NAs implicitly. It is being cautious - what if you didn't know there were missing values for some patients? If you wanted to compare the means of var1 and var2 without any further filtering, you would be comparing samples of different size. Which might be fine if the sample sizes are sufficiently representative and the values are missing at random. Therefore, if you decide to ignore the NAs and want to calculate the mean anyway, you can do so by adding another argument to mean():

```
mean(mydata$var2, na.rm = TRUE)
```

#### ## [1] 4.5

Adding na.rm = TRUE tells are that you are happy for it to calculate the mean of any existing values (but to remove - na.rm - the NA values). This 'removal' only means excluded from the calculation, it does not affect the actual tibble (mydata) holding the columns and rows. R is case sensitive, so it has to be na.rm, not NA.rm etc. There is no need to try to memorise how the arguments of functions are exactly spelled - this is what the Help tab (press F1 when the cursor is on the name of the function) can remind you of. Information about functions (Help) is built into R, so an internet connection is not required for this.

Make sure to always separate arguments with commas, or R will give you an error of Error: unexpected symbol.

Finally, some functions do not need any arguments to work. A good example is the Sys.time() which returns the current time and date. This is very useful when using R to generate and update reports automatically. Including this on the output document means you as well as your colleagues can always be clear on when the results were last updated.

#### Sys.time()

## ## [1] "2019-07-18 16:36:43 BST"

To summarise, objects and functions work hand in hand. Objects are both an input as well as the output of a function (what the function returns). The data values input into a function are usually its first argument, further arguments can be used to specify a functions behaviour. When we say "the function returns", we are referring to its output. The returned object can be different to its input object. In our mean() examples, the input object was a column (mydata\$var1: 4, 1, 2, 3), whereas the output was a single value: 2.5.

## 2.3 Working with Objects

It's sometimes difficult to appreciate how coding works without trying it first. These exercises will show you how R works.

We'll first create an object and call it a, we will give the object a a value of 1. In R the equals = sign tells R to give the object on the left of the sign the value of whatever is on the right of the sign.

#### a = 1

In your environment panel, you should see a appear under the Values section.

Now, lets create b and give it a value of 2.

## b = 2

Lets now add a and b together to create the object c

```
c = a + b
# Print the value of c to the Console
c # should return the number 3
```

## ## [1] 3

All of R is just an extension of this: applying more complex functions (calculations) across more complex objects.

It's important to appreciate that objects can be more than just single numbers too. They can be entire spreadsheets, which in R are known as tibbles (or in base R: data frames).

Note that many people use <- instead of =. They mean the same thing in R. = and <- save what is on the right into the name on the left. There is also a left-to-right operator: ->.

## 2.3.1 Exercise

Create 3 new variables, d, e, f with values 6, 7, 8 using the different assignment operators.

```
d = 6
e <- 7
8 -> f
```

## 2.4 Loading data

Before we load a new dataset, we should clear our experiments from the previous section. Restart R by pressing Ctrl+Shift+F10 or Select Section -> Restart R from the menu above.

Now the environment is clear, lets load in the data:

```
library(tidyverse) # Tidyverse is the package which contains some of the co
mydata = read_csv("global_burden_disease_short.csv")
```

But how can we look at the data we just loaded? How do we know which variables it contains? Hint: the Environment tab.

#### 2.4.1 Exercise

Answer these question about your data:

- 1. At present, how many variables are there?
- 2. How many deaths were there from communicable diseases in 1990? Hint: clicking on columns when Viewing a tibble orders it.

## 2.4.2 Other ways to investigate objects

In most cases, you can rely on the Environment tab to see how many variables you have. If, however, the dataset you are using is too big to easily navigate within, you might need to use names(mydata), head(mydata), or str(mydata).

Furthermore, we can select a single column using the dollar sign: \$.

So if we type:

#### mydata\$deaths

```
## [1] 16149409 26993493 4325788 15449045 29897069 4639869 14775502
```

## [8] 31521934 4776852 13890709 33637815 4833919 12431802 36259550

## [15] 4970846 11809640 38267197 4786929

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R will give us all the data for that variable.

## 2.4.3 Exercise



Ceci n'est pas un pipe.

Image source: https://cran.r-project.org/web/packages/magrittr/vignettes/magrittr.html

Re-write names (mydata) and head (mydata) using the pipe (%>%). Use the keyboard shortcut Ctrl+Shift+M to insert it.

#### 2.4.4 Exercise

How many unique values does the cause variable have? Hint: mydata\$cause piped into unique() piped into length().

## 2.5 Operators

Operators are symbols in R Code that tell R how to handle different pieces of data or objects.

Here are the main operators:

Some of these perform a test on data. A good example of this is the '==' operator.

This tells R to compare two things and ask if they are equal. If they are equal R will return 'TRUE', if not R will return 'FALSE'.

On your R cheat sheet, you can see what the others do. Here is a reminder:

Symbol	What does	Example	Example result
= or <-	assigns	x = 2	the value of x is now 2
==	Equal?	x == 2	TRUE
! =	Not equal?	x != 1	TRUE
<	Less than	x < 2	FALSE
>	Greater than	x > 1	TRUE
<=	Less than or equal to	x <= 2	TRUE
>=	Greater than or equal to	x >= 1	TRUE
%>%	sends data into a function	x %>% print()	2
::	indicates package	dplyr::count()	count() fn. from the dplyr p
->	assigns	2 -> x	the value of x is now 2
&	AND	x > 1 & x < 3	TRUE
	OR	x > 3   x == 3	TRUE
%in%	is value in list	x %in% c(1,2,3)	TRUE
\$	select a column	mydata\$year	1990,1996,
c()	combines values	c(1, 2)	1, 2
#	comment	#Riinu changed this	ignored by R

For example, if we wanted to select the years in the Global Burden of disease study after 2000 (and including 2000) we could type the following:

```
mydata %>%
filter(year >= 2000)
```

To save this as a new object we would then write:

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```
mydata_out = mydata %>%
  filter(year >= 2000)

# Or we could write

mydata %>%
  filter(year >= 2000) -> mydata_out
```

How would you change the above code to only include years greater than 2000 (so not including 2000 itself too)? Hint: look at the table of operators above (also in your HealthyR QuickStart Sheet).

#### 2.5.1 Exercise

Modify the above example to filter for only year 2000, not all years greater than 2000. Save it into a variable called mydata\_year2000.

#### 2.5.2 Exercise

Let's practice this and combine multiple selections together.

This '|' means OR and '&' means AND.

From mydata, select the lines where year is either 1990 or 2013 and cause is "Communicable diseases":

```
new_data_selection = mydata %>%
  filter( (year == 1990 | year == 2013) & cause == "Communicable diseases")
# Or we can get rid of the extra brackets around the years
# by moving cause into a new filter on a new line:

new_data_selection = mydata %>%
  filter(year == 1990 | year == 2013) %>%
  filter(cause == "Communicable diseases")
```

## 2.6 Types of variables

consider structuring as per here: https://finalfit.org/ articles/data\_prep.html

Like many other types of statistical software, R needs to know the variable type of each column. The main types are:

#### 2.6.1 Characters

Characters (sometimes referred to as *strings* or *character strings*) in R are letters, words, or even whole sentences (an example of this may be free text comments). We can specify these using the as.character() function. Characters are displayed in-between "" (or '').

#### 2.6.2 Factors

Factors are fussy characters. Factors are fussy because they have something called levels. Levels are all the unique values this variable could take - e.g. like when we looked at mydata\$cause %>% unique(). Using factors rather than just characters can be useful because:

- The values factor levels can take is fixed. For example, if the levels of your column called sex are "Male" and "Female" and you try to add a new patient where sex is called just "F" you will get a warning from R. If sex was a character column rather than a factor R would have no problem with this and you would end up with "Male", "Female", and "F" in your column.
- Levels have an order. When we plotted the different causes of death in the last session, R ordered them alphabetically (because cause was a character rather than a factor). But if you want to use a non-alphabetical order, e.g. "Communicable diseases"-"Non-communicable diseases"-"Injuries", we need make cause

into a factor. Making a character column into a factor enables us to define and change the order of the levels. Furthermore, there are useful tools such as fct\_inorder or fct\_infreq that can order factor levels for us.

These can be huge benefits, especially as a lot of medical data analyses include comparing different risks to a reference level. Nevertheless, the fussiness of factors can sometimes be unhelpful or even frustrating. For example, if you really did want to add a new level to your gender column (e.g., "Prefer not to say") you will either have to convert the column to a character, add it, and convert it back to a factor, or use fct\_expand to add the level and then add your new line.

#### 2.6.2.1 Exercise

Temporarily type fct\_inorder anywhere in your script, then press F1. Read the **Description** in the Help tab and discuss with your neighbour how fct\_inorder and fct\_infreq would order your factor levels.

#### 2.6.3 Numbers

Self-explanatory! These are numbers. In R, we specify these using the as.numeric() function. Numbers without decimal places are sometimes called integers. Click on the blue arrow in front of mydata in the Environment tab and see that year is an int (integer) whereas deaths is a num (numeric).

## 2.6.4 Specifying variable types

```
as.character(mydata$cause)
as.numeric(mydata$year)
```

```
factor(mydata$year)

#Lets save the cause as a factor

mydata$cause = factor(mydata$cause)

#Now lets print it out

mydata$cause
```

#### 2.6.5 Exercise

Change the order of the levels in mydata\$cause so that "Non-communicable diseases" come before "Injuries". Hint: use F1 to investigate examples of how fct\_relevel() works.

## 2.7 Importing data

For historical reasons, R's default functions (e.g. read.csv() or data.frame()) convert all characters to factors automatically (for more on this see forcats.tidyverse.org<sup>1</sup>. But it is usually more convenient to deal with characters and convert some of the columns to factors when necessary.

Base R:

```
mydata = read.csv("global_burden_disease_short.csv", stringsAsFactors = FALS
```

The tidyverse version, read\_csv(), has stringsAsFactors set to

<sup>1</sup>http://forcats.tidyverse.org

FALSE by default (and it is a lot faster than read.csv() when reading in large datasets).

Tidyverse:

```
mydata = read_csv("global_burden_disease_short.csv")
```

```
## Parsed with column specification:
## cols(
## cause = col_character(),
## year = col_double(),
## deaths = col_double()
## )
```

You can use the "Import Dataset" button in the Environment tab to get the code for importing data from Excel, SPSS, SAS, or Stata.

## 2.8 Adding columns to dataframes

If we wanted to add in a new column or variable to our data, we can simply use the dollar sign '\$' to create a new variable inside a pre-existing piece of data:

```
mydata$new = 1
mydata$new2 = 1:18
```

Run these lines and click on mydata in the Environment tab to check this worked as expected.

Conversely, if we want to delete a specific variable or column we can use the 'NULL' function, or alternatively ask R to select() the data without the new variable included.

```
mydata$new = NULL

mydata = mydata %>%
    select(-new2)
```

We can make new variables using calculations based on variables in the data too.

The mutate function is useful here. All you have to specify within the mutate function is the name of the variable (this can be new or pre-existing) and where the new data should come from.

There are two equivalent ways of defining new columns based on a calculation with a previous column:

mutate formally introduced in later chapter. Need to think how best to present this in book.

Throughout this course we will be using both of these ways to create or modify columns. The first option (using the \$) can look neater when changing a single variable, but when combining multiple ones you will end up repeating mydata\$. mutate() removes the duplication, but it does add a new line and brackets.

## 2.9 Rounding numbers

We can use round() to round the new variables to create integers.

#### 2.9.1 Exercise

Round the new column deaths\_millions to no decimals:

```
## [1] 16 27 4 15 30 5 15 32 5 14 34 5 12 36 5 12 38 5
```

- How would you round it to 2 decimals? Hint: use F1 to investigate round().
- What do ceiling() and floor() do? Hint: sometimes you want to round a number up or down.

## 2.10 The combine function: c()

The combine function combines several values: c()

The combine function can be used with numbers or characters (like words or letters):

```
examplelist = c("Red", "Yellow", "Green", "Blue")
# Ask R to print it by executing it on its own line
examplelist
```

```
## [1] "Red" "Yellow" "Green" "Blue"
```

## 2.10.1 Exercise

There are 18 lines (observations) in mydata. Create a new variable using c() with 18 values (numbers, words, whichever you like, e.g. like we created examplelist). Then add it as new column to mydata\$newlist. Advanced version: do this using a combination of rep() and c().

## 2.11 The paste() function

The paste() function is used to paste several words or numbers into one character variable/sentence.

In the paste function we need to specify what we would like to combine, and what should separate the components. By default, the separation is a space, but we can change this using the sep = option within the paste function.

So, for example if we wanted to make a sentence:

```
#
#paste("Edinburgh", "is", "Great")
# Lets add in full stops

paste("Edinburgh", "is", "Great", sep = ".")
```

## [1] "Edinburgh.is.Great"

```
# separator needs to go in "" as it is a character

# If we really like Edinburgh

#paste("Edinburgh", "is", "Great", sep = "!")

# If we want to make it one word

#paste("Edinburgh", "is", "Great", sep = "") # no separator (still need the
```

We can also join two different variables together using paste():

```
paste("Year is", mydata$year)

## [1] "Year is 1990" "Year is 1990" "Year is 1990" "Year is 1995"
## [5] "Year is 1995" "Year is 1995" "Year is 2000" "Year is 2000"
## [9] "Year is 2000" "Year is 2005" "Year is 2005" "Year is 2005"
## [13] "Year is 2010" "Year is 2010" "Year is 2010" "Year is 2013"

## [17] "Year is 2013" "Year is 2013"

2.11.1 Exercise

Fix this code:
Hint: Think about characters and quotes!

paste(Today is, Sys.Date())
```

## 2.12 Combining two dataframes

For combining dataframes based on shared variables we use the joins: left\_join(), right\_join(), inner\_join(), or full\_join(). Let's split some of the variables in mydata between two new dataframes: first\_data and second\_data. For demonstrating the difference between the different joins, we will only include a subset (first 6 rows) of the dataset in second data:

```
first_data = select(mydata, year, cause, deaths_millions)
second_data = select(mydata, year, cause, deaths_millions) %>% slice(1:6)

# change the order of rows in first_data to demosntrate the join does not not first_data = arrange(first_data, deaths_millions)
```

```
combined_left = left_join(first_data, second_data)
combined_right = right_join(first_data, second_data)
combined_inner = inner_join(first_data, second_data)
combined_full = full_join(first_data, second_data)
```

Those who have used R before, or those who come across older scripts will have seen merge() instead of the joins. merge() works similarly to joins, but instead of having the four options defined clearly at the front, you would have had to use the all = FALSE, all.x = all, all.y = all arguments.



#### 2.12.1 Exercise

Investigate the four new dataframes called **combined\_** using the Environment tab and discuss how the different joins (left, right, inner, full) work.

## 2.13 The summary() function

In R, the summary() function provides a quick way of summarising both data or the results of statistical tests.

Lets get a quick summary of all the variables inside the Global Burden of Disease dataset. It will work for whole datasets and single variables too.

```
mydata %>% summary()
##
     cause
                                 deaths
                                              years_from_1990
                      year
## Length:18
                    Min.
                         :1990
                                 Min.
                                        : 4325788 Min. : 0.00
   Class:character 1st Qu.:1995
                                    1st Qu.: 4868151
                                                       1st Qu.: 5.00
   Mode :character Median :2002 Median :14333106
                                                     Median: 12.50
##
                      :2002 Mean :17189854 Mean
                                                       :12.17
##
                 3rd Qu.:2010
                               3rd Qu.:29171175
                                                  3rd Qu.:20.00
##
                       :2013 Max.
                                     :38267197
                                                       :23.00
                 Max.
                                               Max.
##
    deaths_millions
##
    Min.
           : 4.00
##
    1st Qu.: 5.00
##
    Median :14.50
##
           :17.22
    Mean
##
    3rd Qu.:29.25
            :38.00
    Max.
This even works on statistical tests (we will learn more about these
later):
# lm stands for linear model
lm(deaths ~ year, data = mydata) %>% summary()
##
## Call:
## lm(formula = deaths ~ year, data = mydata)
##
## Residuals:
##
                     1Q
                           Median
                                          3Q
                                                    Max
                         -2889909
                                    12818624
                                              19999627
## -13480641 -11791203
##
## Coefficients:
                  Estimate Std. Error t value Pr(>|t|)
## (Intercept) -181988644
                            736014812
                                        -0.247
                                                   0.808
## year
                                         0.271
                     99482
                                367606
                                                   0.790
##
```

```
## Residual standard error: 12590000 on 16 degrees of freedom
## Multiple R-squared: 0.004556, Adjusted R-squared: -0.05766
## F-statistic: 0.07324 on 1 and 16 DF, p-value: 0.7901
```

#### 2.13.1 When pipe sends data to the wrong place

Note that our usual way of doing things with the pipe would not work here:

```
mydata %>%
  lm(deaths ~ year) %>%
  summary()
```

This is because the pipe tries to send data into the first place of the function (first argument), but lm() wants the formula (deaths ~ year) first, then the dataframe. We can bypass this using data = . to tell the pipe where to put mydata:

```
mydata %>%
  lm(deaths ~ year, data = .) %>%
  summary()
```

#### 2.13.2 Exercise

Try adding a new variable called death\_over\_10m which indicates whether there were more than 10 million deaths for a cause. The new variable should take the form 'Yes' or 'No'.

Then make it into a factor.

Then use summary() to find out about it!

```
mydata = mydata %>%
  mutate(death_over_10m = ifelse(deaths >= 10000000, "Yes", "No")) # Using a
```

```
mydata$death_over_10m = as.factor(mydata$death_over_10m)
mydata$death_over_10m %>% summary()

## No Yes
## 6 12
```

## 2.14 Extra: Creating a dataframe from scratch

It is rare that you will need to create a data frame by hand as most of the time you will be reading in a data from a .csv or similar. But in some cases (e.g. when creating special labels for a plot) it might be useful, so this is how to create one:

## Warning: `data\_frame()` is deprecated, use `tibble()`.
## This warning is displayed once per session.

```
# same as

newdata = data_frame(
  patient_id = paste0("ID", 1:10), #note the commas
  sex = rep(c("Female", "Male"), 5),
  age = 18:27
)
```

If we used data.frame() instead of data\_frame(), all our char-

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acter variables (patient\_id, sex) would become factors automatically. This might make sense for sex, but it doesn't for patient\_id.

## 2.14.1 Exercise

Create a new dataframe called my\_dataframe that looks like this:

Hint: Use the functions pasteO(), seq() and rep()

```
## # A tibble: 10 x 3
##
      patient_id
                    age sex
##
      <chr>>
                  <dbl> <chr>
##
   1 ID11
                     15 Male
##
   2 ID12
                     20 Male
   3 ID13
                     25 Male
##
##
   4 ID14
                     30 Male
##
   5 ID15
                     35 Male
##
   6 ID16
                     40 Female
   7 ID17
                     45 Female
##
   8 ID18
                     50 Female
##
                     55 Female
##
   9 ID19
## 10 ID20
                     60 Female
```

## 2.15 Solutions

## 2.5.3

```
mydata %>% names()
mydata %>% head()
mydata %>% str()
```

## 2.5.4

```
mydata$cause %>% unique() %>% length()
## [1] 3
2.6.2
mydata_year2000 = mydata %>%
  filter(year == 2000)
2.7.5
mydata$cause %>% fct_relevel("Injuries", after = 1)
2.10.1
mydata$deaths_millions = round(mydata$deaths_millions)
mydata$deaths_millions = mydata$deaths_millions %>% round()
2.11.1
examplelist = c("Red", "Yellow", "Green", "Blue",
                "Red", "Yellow", "Green", "Blue",
                "Red", "Yellow", "Green", "Blue",
                "Red", "Yellow", "Green", "Blue",
                "Green", "Blue")
#Let's see what we've made by using print
mydata$newlist = examplelist
# using rep()
```

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```
examplelist2 = rep(c("Green", "Red"), 9)
```

## 2.12.1

```
paste("Today is", Sys.Date())
```

## 2.15.1

# Summarising data

In this session we will get to know our three best friends for summarising data: group\_by(), summarise(), and mutate().

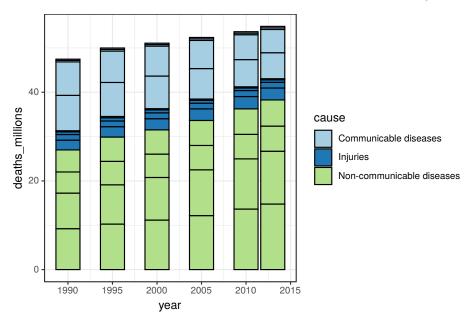
## 3.1 Data

In Session 2, we used a very condensed version of the Global Burden of Disease data. We are now going back to a longer one and we will learn how to summarise it ourselves.

```
source("healthyr_theme.R")
load("global_burden_disease_long.rda")
```

We were already using this longer dataset in Session 1, but with colour=cause to hide the fact that the total deaths in each year was made up of 12 groups of data (as the black lines on the bars indicate):

```
mydata %>%
    ggplot(aes(x = year, y = deaths_millions, fill = cause))+
    geom_col(colour = "black")
```



```
mydata %>%
  filter(year == 1990)
```

```
##
      location
                                     sex year deaths_millions
                             cause
## 1 Developing Non-communicable diseases Male 1990
                                                          9.2277141
## 2 Developing Non-communicable diseases Female 1990
                                                          8.0242455
## 3
     Developed Non-communicable diseases Male 1990
                                                         4.7692902
     Developed Non-communicable diseases Female 1990
                                                          4.9722431
## 5 Developing
                            Injuries Male 1990
                                                    2.2039625
## 6 Developing
                            Injuries Female 1990
                                                     1.2698308
## 7
     Developed
                           Injuries Male 1990
                                                    0.5941184
                           Injuries Female 1990
## 8
     Developed
                                                    0.2578759
## 9 Developing
                   Communicable diseases Male 1990
                                                        7.9819728
## 10 Developing
                   Communicable diseases Female 1990
                                                         7.5416376
## 11 Developed
                   Communicable diseases Male 1990
                                                        0.3387820
                   Communicable diseases Female 1990
## 12 Developed
                                                         0.2870169
```

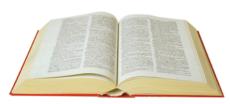
# 3.2 Tidyverse packages: ggplot2, dplyr, tidyr, etc.

Most of the functions introduced in this session come from the tidyverse family (http://tidyverse.org/), rather than Base R. Including library(tidyverse) in your script loads a list of packages: ggplot2, dplyr, tidry, forcats, etc.

# R LINGUA: LIBRARY VS PAG

REAL LIFE





library <

book <

 $\mathbf{C}$ 

library <

package

I went to the library to use the English dictionary (it was I then ordered a specialised book ("General Surgery") to re

I used R to calculate the means and medians of my of then loaded a specialised package ("survival") to calculate the

library(tidyverse)

## 3.3 Basic functions for summarising data

You can always pick a column and ask R to give you the sum(), mean(), min(), max(), etc. for it:

```
mydata$deaths_millions %>% sum()
```

## [1] 309.4174

```
mydata$deaths_millions %>% mean()
```

## ## [1] 4.297463

But if you want to get the total number of deaths for each year (or cause, or sex, whichever grouping variables you have in your dataset) you can use group\_by() and summarise() that make subgroup analysis very convenient and efficient.

## 3.4 Subgroup analysis: group\_by() and summarise()

The group\_by() function tells R that you are about to perform subgroup analysis on your data. It retains information about your groupings and calculations are applied on each group separately. To go back to summarising the whole dataset again use ungroup(). Note that summarise() is different to the summary() function we used in Session 2.

With summarise(), we can calculate the total number of deaths per year:

```
mydata %>%
    group_by(year) %>%
    summarise(total_per_year = sum(deaths_millions)) ->
    summary_data1

mydata %>%
    group_by(year, cause) %>%
    summarise(total_per_cause = sum(deaths_millions)) ->
    summary_data2
```

- summary\_data1 includes the total number of deaths per year.
- summary\_data2 includes the number of deaths per cause per year.

year	total_per_year
1990	47
1995	50
2000	51
2005	52
2010	54
2013	55

year	cause	total_per_cause
1990	Communicable diseases	16
1990	Injuries	4
1990	Non-communicable diseases	27
1995	Communicable diseases	15
1995	Injuries	5
1995	Non-communicable diseases	30

<sup>...</sup> remaining years omitted from printing.

## 3.4.1 Exercise

Compare the sizes - number of rows (observations) and number of columns (variables) - of mydata, summary\_data1, and summary\_data2 (in the Environment tab).

- Convince yourself that for 1990, deaths by the three causes (summary\_data2) add up to total deaths per year (summary\_data1).
- summary\_data2 has exactly 3 times as many rows as summary\_data1. Why?
- mydata has 5 variables, whereas the summarised dataframes have 2 and 3. Which variables got dropped? Why?

## 3.4.2 Exercise

For each cause, calculate its percentage to total deaths in each year.

Hint: Use full\_join() on summary\_data1 and summary\_data2. Solution:

```
alldata = full_join(summary_data1, summary_data2)
```

```
## Joining, by = "year"
```

```
alldata$percentage = 100*alldata$total_per_cause/alldata$total_per_year %>%
```

round() defaults to 0 digits. If you want to round to a specified number of decimal places, use, e.g., round(digits = 2).

TABLE 3.1: alldata

year	total_per_year	cause	$total\_per\_cause$	percentage
1990	47	Communicable diseases	16	34
1990	47	Injuries	4	9
1990	47	Non-communicable diseases	27	57
1995	50	Communicable diseases	15	31
1995	50	Injuries	5	9
1995	50	Non-communicable diseases	30	60

## 3.5 mutate()

Mutate works similarly to summarise() (as in it respects groupings set with group\_by()), but it adds a new column into the original data. summarise(), on the other hand, condenses the data into a minimal table that only includes the variables specifically asked for.

## 3.5.1 Exercise

Investigate these examples to learn how summarise() and mutate() differ.

```
summarise_example = mydata %>%
    summarise(total_deaths = sum(deaths_millions))

mutate_example = mydata %>%
    mutate(total_deaths = sum(deaths_millions))
```

```
mutate_example %>%
slice(1:5) %>%
```

3.5 mutate() 49

**TABLE 3.2:** summarise example

total_	_deaths
3	809

**TABLE 3.3:** mutate\_example

location	cause	sex	year	deaths_millions	total_deaths
Developing	Non-communicable diseases	Male	1990	9	309
Developing	Non-communicable diseases	Female	1990	8	309
Developed	Non-communicable diseases	Male	1990	5	309
Developed	Non-communicable diseases	Female	1990	5	309
Developing	Non-communicable diseases	Male	1995	10	309

You should see that mutate() adds the same total number (309) to every line in the dataframe.

## 3.5.2 Optional advanced exercise

Based on what we just observed on how mutate() adds a value to each row, can you think of a way to redo Exercise 3.4.2 without using a join? Hint: instead of creating summary\_data1 (total deaths per year) as a separate dataframe which we then merge with summary\_data2 (total deaths for all causes per year), we can use mutate() to add total\_per\_year to each row.

```
mydata %>%
   group_by(year, cause) %>%
```

```
summarise(total_per_cause = sum(deaths_millions)) %>%
group_by(year) %>%
mutate(total_per_year = sum(total_per_cause)) %>%
mutate(percentage = 100*total_per_cause/total_per_year) -> alldata
```

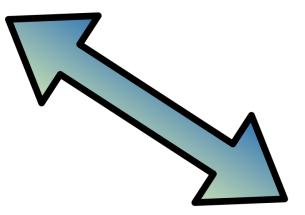
3.6 mutate()

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# 3.6 Wide vs long: spread() and gather()

Developed countries:	1990	1995	2000	2005	2
Communicable diseases	0.6	0.7	0.7	0.7	
Injuries	0.9	1	0.9	0.9	
Non-communicable diseases	9.7	10.8	10.7	11.1	1

1990	1995	2000	2005	20
15.5	14.8	14.1	13.2	1
3.5	3.6	3.8	3.9	catio
17.3	19.1	20.8	22 5	evelo
	15.5 3.5	15.5 14.8 3.5 3.6	15.5 14.8 14.1 3.5 3.6 3.8	15.5 14.8 14.1 13.2 3.5 3.6 3.8 3.9 lo



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#### 3.6.1 Wide format

Although having data in the long format is very convenient for R, for publication tables, it makes sense to spread some of the values out into columns:

```
alldata %>%
    mutate(percentage = paste0(round(percentage, 2), "%")) %>%
    select(year, cause, percentage) %>%
    spread(cause, percentage)
## # A tibble: 6 x 4
## # Groups:
               year [6]
    year `Communicable diseases` Injuries `Non-communicable diseases`
##
     <int> <chr>
                                    <chr>
                                              <chr>
## 1 1990 34.02%
                                    9.11%
                                              56.87%
## 2
     1995 30.91%
                                    9.28%
                                             59.81%
                                             61.72%
## 3
     2000 28.93%
                                    9.35%
## 4
     2005 26.53%
                                    9.23%
                                             64.24%
## 5
                                              67.57%
     2010 23.17%
                                    9.26%
## 6
     2013 21.53%
                                    8.73%
                                             69.75%
```

• select() pick the variables you want to keep. Try running the lines until spread() to see how it works.

#### 3.6.2 Exercise

Calculate the percentage of male and female deaths for each year. Spread it to a human readable form:

#### Hints:

- create summary\_data3 that includes a variable called total\_per\_sex
- merge summary\_data1 and summary\_data3 into a new data frame
- calculate the percentage of total\_per\_sex to total\_per\_year
- round, add % labels

• spread

Solution:

```
mydata %>%
    group_by(year) %>%
    summarise(total_per_year = sum(deaths_millions)) ->
    summary_data1
mydata %>%
    group_by(year, sex) %>%
    summarise(total_per_sex = sum(deaths_millions)) ->
    summary_data3
alldata = full_join(summary_data1, summary_data3)
## Joining, by = "year"
result_spread = alldata %>%
  mutate(percentage = round(100*total_per_sex/total_per_year, 0)) %>%
  mutate(percentage = paste0(percentage, "%")) %>%
  select(year, sex, percentage) %>%
  spread(sex, percentage)
result_spread
## # A tibble: 6 x 3
      year Female Male
     <int> <chr>
##
                  <chr>
## 1 1990 47%
                  53%
## 2 1995 47%
                  53%
## 3 2000 46%
                  54%
## 4 2005 46%
                  54%
## 5
      2010 46%
                  54%
## 6
      2013 45%
                  55%
And save it into a csv file using write_csv():
```

```
write_csv(result_spread, "gbd_genders_summarised.csv")
```

You can open a csv file with Excel and copy the table into Word or PowerPoint for presenting.

#### 3.6.3 Long format

The opposite of spread() is gather():

- The first argument is a name for the column that will include columns gathered from the wide columns (in this example, Male and Female are gathered into sex).
- The second argument is a name for the column that will include the values from the wide-format columns (the values from Male and Female are gathered into percentage).
- Any columns that already are condensed (e.g. year was in one column, not spread out like in the pre-course example) must be included with a negative (i.e. -year).

```
result_spread %>%
  gather(sex, percentage, -year)
```

```
## # A tibble: 12 x 3
##
       year sex
                   percentage
##
      <int> <chr>
                   <chr>
##
      1990 Female 47%
##
      1995 Female 47%
      2000 Female 46%
##
   4 2005 Female 46%
##
   5 2010 Female 46%
##
   6 2013 Female 45%
##
      1990 Male
                   53%
##
##
      1995 Male
                   53%
##
       2000 Male
                   54%
## 10
       2005 Male
                   54%
```

```
## 11 2010 Male 54%
## 12 2013 Male 55%
```

#### 3.6.4 Exercise

Test what happens when you

• Change the order of sex and percentage:

```
result_spread %>%
  gather(percentage, sex, -year)
```

Turns out in the above example, percentage and sex were just label you assigned to the gathered columns. It could be anything, e.g.:

```
result_spread %>%
  gather(`look-I-gathered-sex`, `values-Are-Here`, -year)
```

• What happens if we omit -year:

```
result_spread %>%
  gather(sex, percentage)
```

-year was telling R we don't want the year column to be gathered together with Male and Female, we want to keep it as it is.

# 3.7 Sorting: arrange()

To reorder data ascendingly or descendingly, use arrange():

```
mydata %>%
   group_by(year) %>%
   summarise(total = sum(deaths_millions)) %>%
   arrange(-year) # reorder after summarise()
```

# 3.8 Factor handling

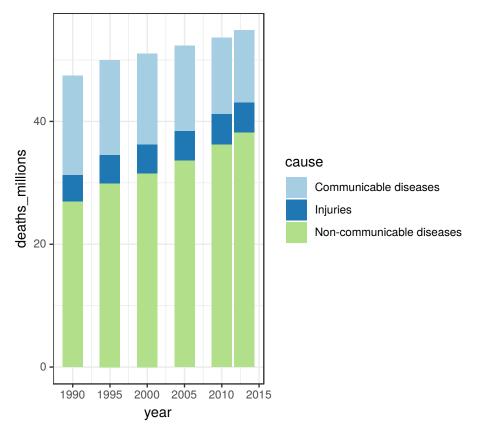
We talked about the pros and cons of working with factors in Session 2. Overall, they are extremely useful for the type of analyses done in medical research.

## 3.8.1 Exercise

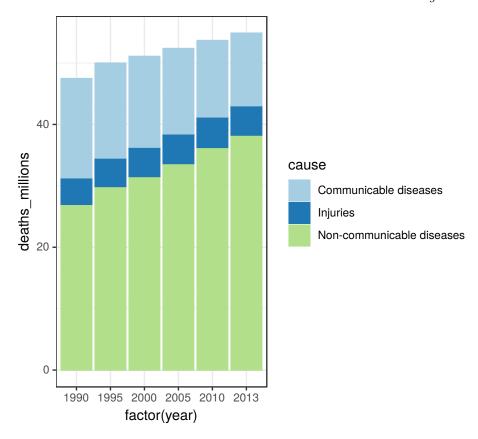
Explain how and why these two plots are different.

```
mydata %>%
    ggplot(aes(x = year, y = deaths_millions, fill = cause))+
    geom_col()
```

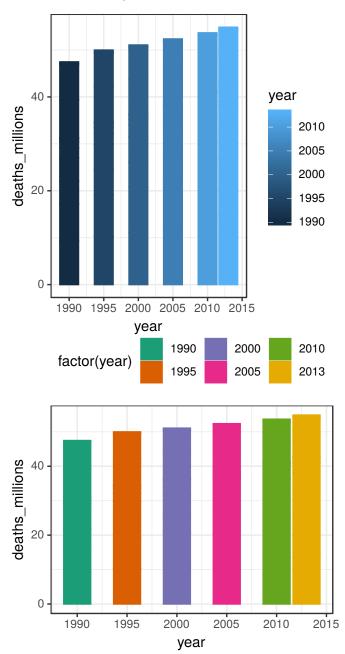




```
mydata %>%
    ggplot(aes(x = factor(year), y = deaths_millions, fill = cause, colour =
    geom_col()
```



What about these?



These illustrate why it might sometimes be useful to use numbers as factors - on the second one we have used fill = factor(year)

as the fill, so each year gets a distinct colour, rather than a gradual palette.

### 3.8.2 fct\_collapse() - grouping levels together

```
mydata$cause %>%
    fct_collapse("Non-communicable and injuries" = c("Non-communicable disea
    mydata$cause2

mydata$cause %>% levels()

## [1] "Communicable diseases" "Injuries"

## [3] "Non-communicable diseases"

mydata$cause2 %>% levels()

## [1] "Communicable diseases" "Non-communicable and injuries"
```

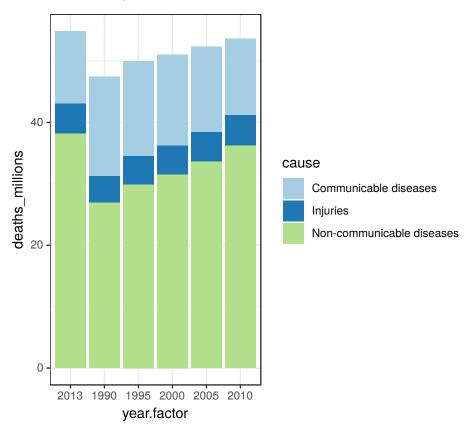
#### 3.8.3 fct\_relevel() - change the order of levels

Another reason to sometimes make a numeric variable into a factor is that we can then reorder it for the plot:

```
mydata$year %>%
  factor() %>%
  fct_relevel("2013") -> #brings 2013 to the front
  mydata$year.factor

source("1_source_theme.R")

mydata %>%
  ggplot(aes(x=year.factor, y=deaths_millions, fill=cause))+
  geom_col()
```



# 3.8.4 fct\_recode() - rename levels

```
mydata$cause %>%
    levels() # levels() lists the factor levels of a column

## [1] "Communicable diseases" "Injuries"
## [3] "Non-communicable diseases"

mydata$cause %>%
    fct_recode("Deaths from injury" = "Injuries") %>%
    levels()
```

```
## [1] "Communicable diseases" "Deaths from injury"
## [3] "Non-communicable diseases"
```

### 3.8.5 Converting factors to numbers

MUST REMEMBER: factor needs to become as.character() before converting to numeric or date! Factors are actually stored as labelled integers (so like number codes), only the function as.character() will turn a factor back into a collated format which can then be converted into a number or date.

#### 3.8.6 Exercise

mydata\$year.factor %>%

Investigate the two examples converting the year.factor variable back to a number.

```
as.character() %>%
as.numeric()
```

#### 3.9 Long Exercise

This exercise includes multiple steps, combining all of the above.

First, create a new script called "2\_long\_exercise.R". Then Restart your R session, add library(tidyverse) and load "global\_burden\_disease\_long.rda".

- Calculate the total number of deaths in Developed and Developing countries. Hint: use group\_by(location) and summarise(new-column-name = sum(variable-to-sum)).
- Calculate the total number of deaths in Developed and Developing countries and for men and women. Hint: this is as easy as adding, sex to group\_by().
- Filter for 1990.
- spread() the location column.

```
## # A tibble: 2 x 3
## sex Developed Developing
## <fct> <dbl> <dbl>
## 1 Female 5.52 16.8
## 2 Male 5.70 19.4
```

## 3.10 Extra: formatting a table for publication

Creating a publication table with both the total numbers and percentages (in brackets) + using formatC() to retain trailing zeros:

```
# Let's use alldata from Exercise 5.2:

mydata %>%
    group_by(year, cause) %>%
    summarise(total_per_cause = sum(deaths_millions)) %>%
```

```
group_by(year) %>%
    mutate(total_per_year = sum(total_per_cause)) %>%
    mutate(percentage = 100*total per cause/total per year) -> alldata
alldata %>%
    mutate(total_percentage =
                    paste0(round(total_per_cause, 1) %>% formatC(1, format
                           "(", round(percentage, 1) %>% formatC(1, format
                    ) %>%
    select(year, cause, total_percentage) %>%
    spread(cause, total_percentage)
## # A tibble: 6 x 4
## # Groups: year [6]
    year `Communicable diseases` Injuries `Non-communicable diseases`
    <int> <chr>
##
                                   <chr>
                                               <chr>>
## 1 1990 16.1 (34.0%)
                                4.3 (9.1%) 27.0 (56.9%)
## 2 1995 15.4 (30.9%)
                                4.6 (9.3%) 29.9 (59.8%)
## 3 2000 14.8 (28.9%)
                                4.8 (9.4%) 31.5 (61.7%)
## 4 2005 13.9 (26.5%)
                                4.8 (9.2%) 33.6 (64.2%)
## 5 2010 12.4 (23.2%)
                                5.0 (9.3%) 36.3 (67.6%)
## 6 2013 11.8 (21.5%)
                                4.8 (8.7%) 38.3 (69.7%)
```

## 3.11 Solution: Long Exercise

```
mydata %>%
  filter(year == 1990) %>%
  group_by(location, sex) %>%
```

```
summarise(total_deaths = sum(deaths_millions)) %>%
spread(location, total_deaths)
```

# Different types of plots

## 4.1 Data

##

##

##

Max.

3rd Qu.:1.959e+07

:1.319e+09

We will be using the gapminder dataset:

```
library(tidyverse)
library(gapminder)
mydata = gapminder
summary(mydata)
##
                                                 lifeExp
         country
                      continent
                                      year
## Afghanistan: 12
                     Africa:624 Min.
                                         :1952
                                                Min.
             : 12
## Albania
                    Americas:300
                                  1st Qu.:1966
                                                1st Qu.:48.20
             : 12
## Algeria
                    Asia
                           :396 Median :1980
                                              Median:60.71
   Angola
             : 12
                   Europe :360
                                 Mean
                                      :1980
                                              Mean
                                                     :59.47
   Argentina: 12 Oceania: 24
                                  3rd Qu.:1993
                                                3rd Qu.:70.85
   Australia: 12
                               Max.
                                     :2007
                                           Max.
                                                  :82.60
    (Other)
               :1632
##
##
                          gdpPercap
         pop
                                    241.2
##
    Min.
           :6.001e+04
                        Min.
    1st Qu.:2.794e+06
                        1st Qu.:
                                   1202.1
##
    Median :7.024e+06
                                   3531.8
                        Median :
    Mean
           :2.960e+07
                        Mean
                                   7215.3
```

3rd Qu.:

Max.

9325.5

:113523.1

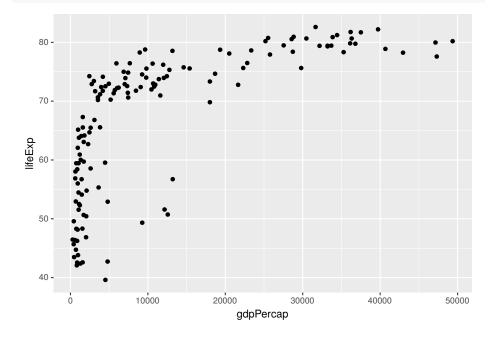
```
mydata$year %>% unique()
```

## [1] 1952 1957 1962 1967 1972 1977 1982 1987 1992 1997 2002 2007

# 4.2 Scatter plots/bubble plots - geom\_point()

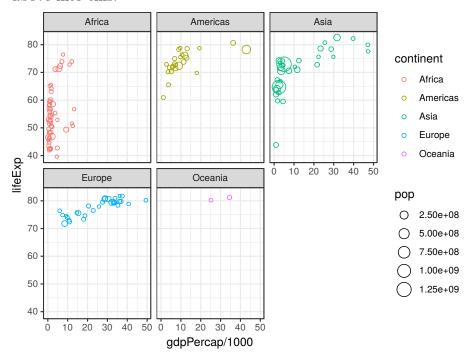
Plot life expectancy against GDP per capita (x = gdpPercap, y=lifeExp) at year 2007:

```
mydata %>%
  filter(year == 2007) %>%
  ggplot(aes(x = gdpPercap, y=lifeExp)) +
  geom_point()
```



#### 4.2.1 Exercise

Follow the step-by-step instructions to transform the grey plot just above into this:

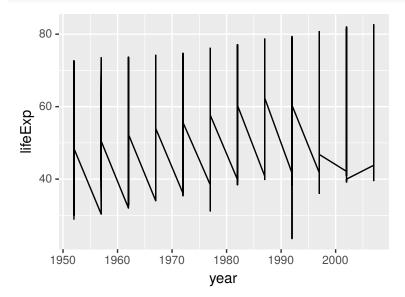


- Add points: geom point()
  - Change point type: shape = 1 (or any number from your Quickstart Sheet) inside the geom\_point()
- Colour each country point by its continent: colour=continent to aes()
- Size each country point by its population: size=pop to aes()
- Put the country points of each continent on a separate panel: + facet\_wrap(~continent)
- Make the background white: + theme\_bw()

# 4.3 Line chart/timeplot - geom\_line()

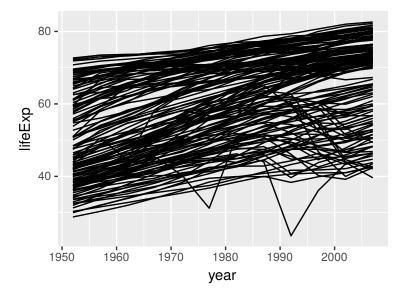
Plot life expectancy against year (x = year, y=lifeExp), add  $geom_line()$ :

```
mydata %>%
  ggplot(aes(x = year, y=lifeExp)) +
  geom_line()
```



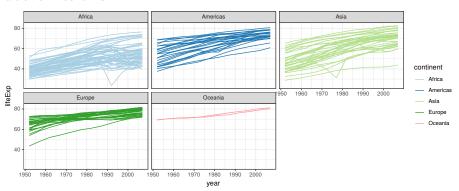
The reason you now see this weird zig-zag is that, using the above code, R does not know you want a connected line for each country. Specify how you want data points grouped to lines: group = country in aes():

```
mydata %>%
  ggplot(aes(x = year, y=lifeExp, group = country)) +
  geom_line()
```



#### 4.3.1 Exercise

Follow the step-by-step instructions to transform the grey plot just above into this:



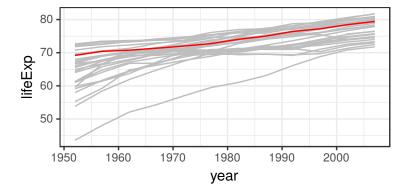
- Colour lines by continents: colour=continent to aes()
- Similarly to what we did in <code>geom\_point()</code>, you can even size the line thicknesses by each country's population: <code>size=pop</code> to <code>aes()</code>
- Continents on separate panels: + facet\_wrap(~continent)
- Make the background white: + theme\_bw()

Use a nicer colour scheme: + scale\_colour\_brewer(palette = "Paired")

## 4.3.2 Advanced example

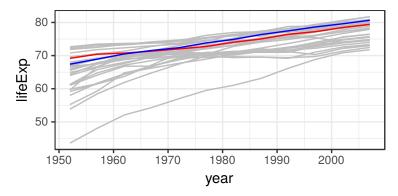
For European countries only (filter(continent == "Europe") %>%), plot life expectancy over time in grey colour for all countries, then add United Kingdom as a red line:

```
mydata %>%
  filter(continent == "Europe") %>% #Europe only
  ggplot(aes(x = year, y=lifeExp, group = country)) +
  geom_line(colour = "grey") +
  theme_bw() +
  geom_line(data = filter(mydata, country == "United Kingdom"), colour = "red")
```



#### 4.3.3 Advanced Exercise

As previous, but add a line for France in blue:

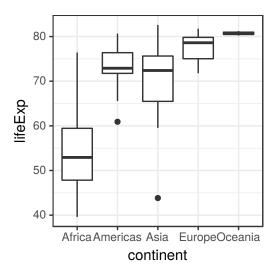


# 4.4 Box-plot - geom\_boxplot()

Plot the distribution of life expectancies within each continent at year 2007:

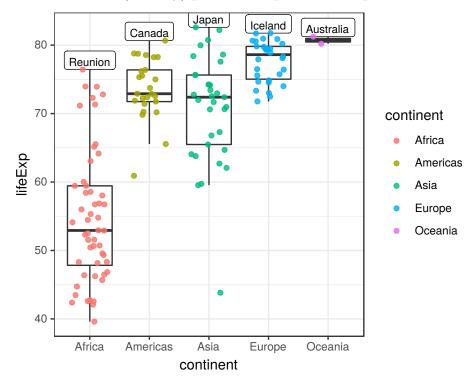
- filter(year == 2007) %>%
- x = continent, y = lifeExp
- + geom\_boxplot()

```
mydata %>%
  filter(year == 2007) %>%
  ggplot(aes(x = continent, y = lifeExp)) +
  geom_boxplot() +
  theme_bw()
```



# 4.4.1 Exercise

Add individual (country) points on top of the box plot:



Hint: Use geom\_jitter() instead of geom\_point() to reduce overlap by spreading the points horizontally. Include the width=0.3 option to reduce the width of the jitter.

# **Optional:**

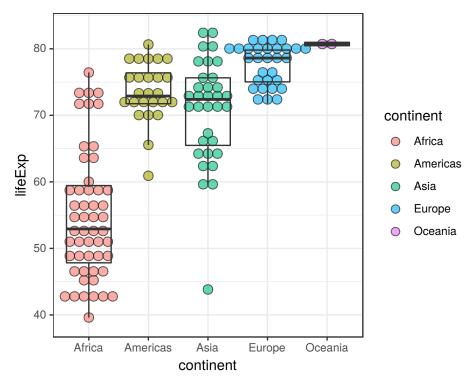
Include text labels for the highest life expectancy country of each continent.

**Hint 1** Create a separate dataframe called label\_data with the maximum countries for each continent:

```
label_data = mydata %>%
  filter(year == max(year)) %>% # same as year == 2007
  group_by(continent) %>%
  filter(lifeExp == max(lifeExp) )
```

Hint 2 Add geom\_label() with appropriate aes():

```
+ geom_label(data = label_data, aes(label=country), vjust = 0)
4.4.2 Dot-plot - geom_dotplot()
geom_dotplot(aes(fill=continent), binaxis = 'y',
stackdir = 'center', alpha=0.6)
```



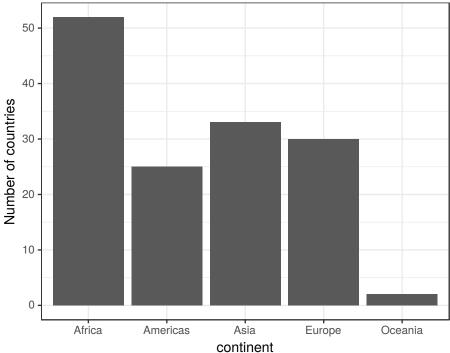
# 4.5 Barplot - geom\_bar() and geom\_col()

In the first module, we plotted barplots from already summarised data (using the geom\_col), but geom\_bar() is perfectly happy to count up data for you. For example, we can plot the number of countries in each continent without summarising the data beforehand:

```
mydata %>%
  filter(year == 2007) %>%
  ggplot(aes(x = continent)) +
  geom_bar() +
```

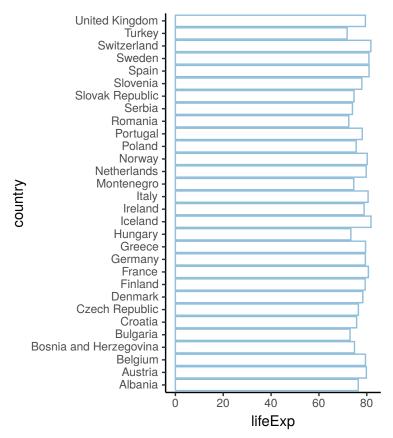
79





## 4.5.1 Exercise

Create this barplot of life expectancies in European countries (year 2007). Hint: coord\_flip() makes the bars horizontal, fill = NA makes them empty, have a look at your QuickStar sheet for different themes.



# 4.6 All other types of plots

These are just some of the main ones, see this gallery for more options: http://www.r-graph-gallery.com/portfolio/ggplot2-package/

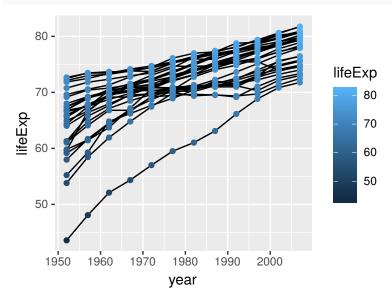
And the ggplot() documentation: http://docs.ggplot2.org/

Remember that you can always combine different types of plots - i.e. add lines or points on bars, etc.

# 4.7 Specifying aes() variables

The aes() variables wrapped inside ggplot() will be taken into account by all geoms. If you put aes(colour = lifeExp) inside geom\_point(), only points will be coloured:

```
mydata %>%
  filter(continent == "Europe") %>%
  ggplot(aes(x = year, y = lifeExp, group = country)) +
  geom_line() +
  geom_point(aes(colour = lifeExp))
```



# 4.8 Extra: Optional exercises

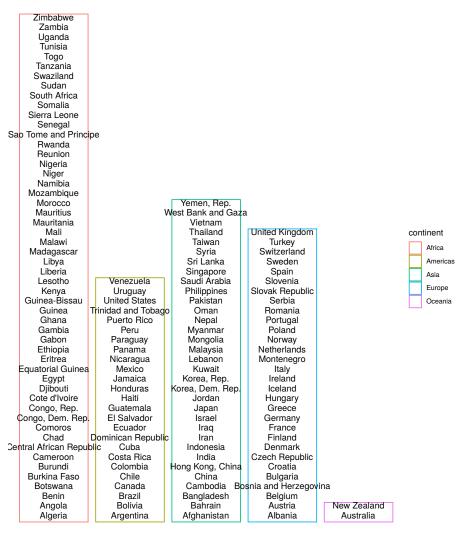
#### 4.8.1 Exercise

Make this:

```
mydata$dummy = 1  # create a column called "dummy" that includes number 1 f

mydata2007 = mydata %>%
    filter(year==max(year)) %>%
    group_by(continent) %>%
    mutate(country_number = cumsum(dummy)) # create a column called "country"
    # is a cumulative sum of the number of countries before it - basically in

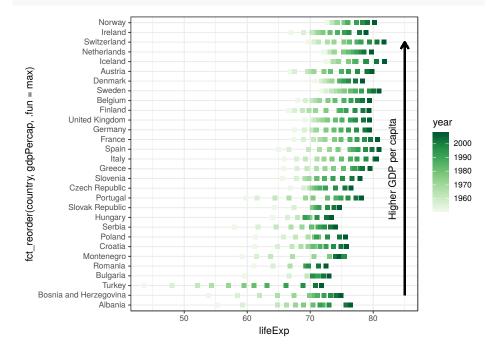
mydata2007 %>%
    ggplot(aes(x = continent)) +
    geom_bar(aes(colour=continent), fill = NA) +
    geom_text(aes(y = country_number, label=country), size=4, vjust=1, colour=theme_void()
```



#### 4.8.2 Exercise

```
Make this:
```

```
Hints: coord_flip(), scale_color_gradient(...),
geom_segment(...), annotate("text", ...)
```



4.9 Solutions 85

#### 4.9 Solutions

#### 4.2.1

#### 4.3.1

```
mydata %>%
  ggplot( aes(x = year, y=lifeExp, group = country, colour=continent)) +
  geom_line() +
  facet_wrap(~continent) +
  theme_bw() +
  scale_colour_brewer(palette = "Paired")

which
```

```
Add + geom_line(data = filter(mydata, country == "France"), colour = "blue")
```

## 4.4.1

```
mydata %>%
  filter(year == 2007) %>%
  ggplot(aes(x = continent, y = lifeExp)) +
  geom_boxplot(outlier.shape = NA) +
```

```
geom_jitter(aes(colour=continent), width=0.3, alpha=0.8) + #width defaults
theme_bw()
```

```
mydata %>%
  filter(year == 2007) %>%
  ggplot(aes(x = continent, y = lifeExp)) +
  geom_boxplot(outlier.shape = NA) +
  geom_jitter(aes(colour=continent), width=0.3, alpha=0.8)
  theme_bw()
```

#### 4.5.1

```
mydata %>%
  filter(year == 2007) %>%
  filter(continent == "Europe") %>%
  ggplot(aes(x = country, y = lifeExp)) +
  geom_col(colour = "#91bfdb", fill = NA) +
  coord_flip() +
  theme_classic()
```

# Fine tuning plots

# 5.1 Data and initial plot

We can save a ggplot() object into a variable (usually called p but can be any name). This then appears in the Environment tab. To plot it it needs to be recalled on a separate line. Saving a plot into a variable allows us to modify it later (e.g., p + theme\_bw()).

```
library(gapminder)
library(tidyverse)

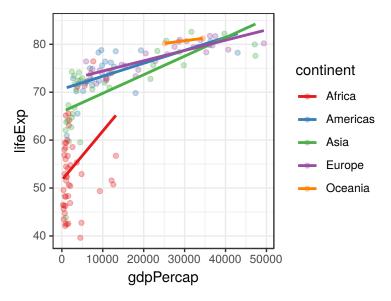
mydata = gapminder

mydata$year %>% unique()
```

## [1] 1952 1957 1962 1967 1972 1977 1982 1987 1992 1997 2002 2007

```
p = mydata %>%
  filter(year == 2007) %>%
  group_by(continent, year) %>%
  ggplot(aes(y = lifeExp, x = gdpPercap, colour = continent)) +
  geom_point(alpha = 0.3) +
  theme_bw() +
  geom_smooth(method = "lm", se = FALSE) +
  scale_colour_brewer(palette = "Set1")

p
```

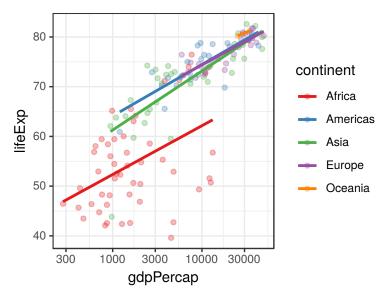


# 5.2 Scales

# 5.2.1 Logarithmic

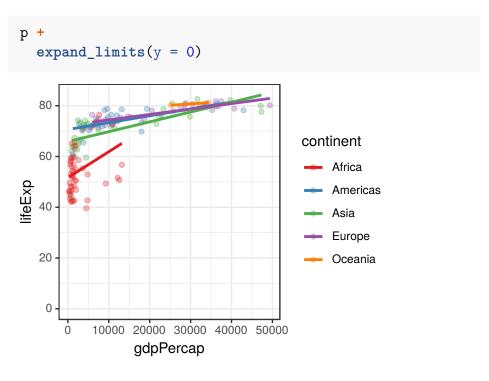
```
p +
scale_x_log10()
```

5.2 Scales 89



# 5.2.2 Expand limits

Specify the value you want to be included:



Or two:

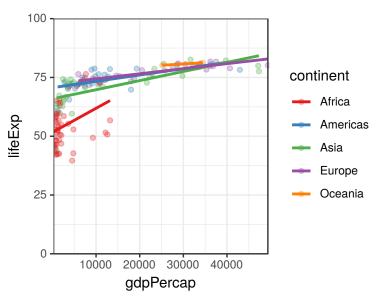
```
p +
expand_limits(y = c(0, 100))

continent
Africa
Americas
Asia
Europe
Oceania
```

By default, ggplot() adds some padding around the included area (see how the scale doesn't start from 0, but slightly before). You can remove this padding with the expand option:

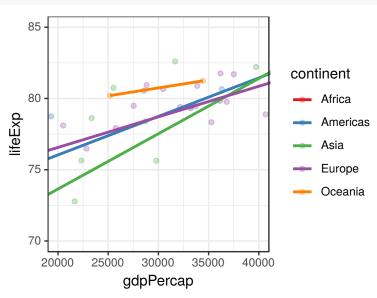
```
p +
  expand_limits(y = c(0, 100)) +
  coord_cartesian(expand = FALSE)
```

5.2 Scales 91



## 5.2.3 Zoom in

```
p +
  coord_cartesian(ylim = c(70, 85), xlim = c(20000, 40000))
```



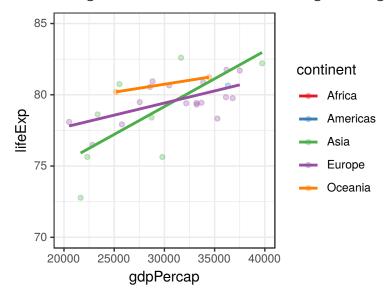
#### 5.2.4 Exercise

How is this one different to the previous?

```
p +
   scale_y_continuous(limits = c(70, 85)) +
   scale_x_continuous(limits = c(20000, 40000))
```

## Warning: Removed 114 rows containing non-finite values (stat\_smooth).

## Warning: Removed 114 rows containing missing values (geom\_point).

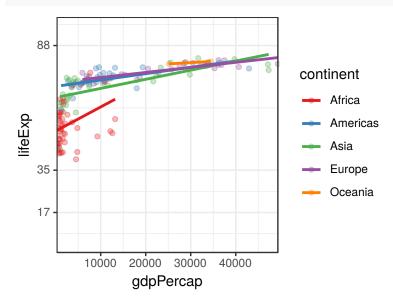


Answer: the first one zooms in, still retaining information about the excluded points when calculating the linear regression lines. The second one removes the data (as the warnings say), calculating the linear regression lines only for the visible points.

#### 5.2.5 Axis ticks

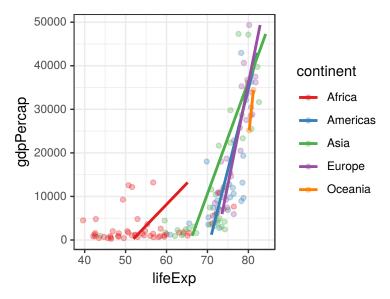
5.2 Scales 93

```
p +
  coord_cartesian(ylim = c(0, 100), expand = 0) +
  scale_y_continuous(breaks = c(17, 35, 88))
```



# 5.2.6 Swap the axes

```
p +
coord_flip()
```

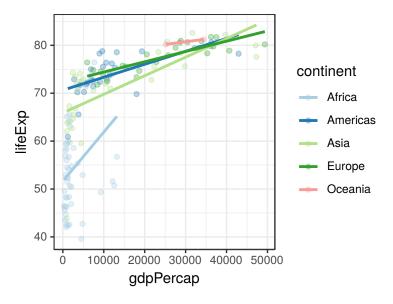


# 5.3 Colours

# 5.3.1 Using the Brewer palettes:

```
p +
    scale_color_brewer(palette = "Paired")
```

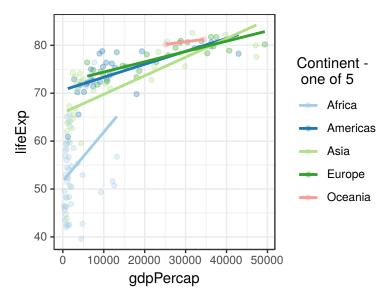
5.3 Colours 95



# 5.3.2 Legend title

scale\_colour\_brewer() is also a conventient place to change the legend title:

```
p +
    scale_color_brewer("Continent - \n one of 5", palette = "Paired")
```



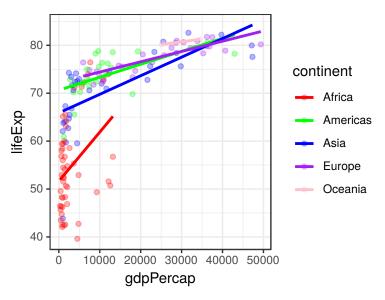
Note the \n inside the new legend title - new line.

# 5.3.3 Choosing colours manually

Use words:

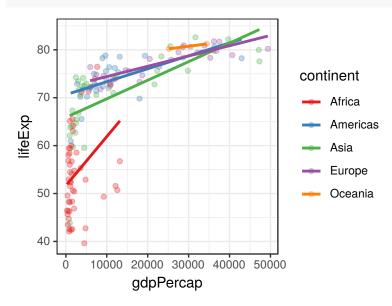
```
p +
    scale_color_manual(values = c("red", "green", "blue", "purple", "pink"))
```

5.3 Colours 97



Or HEX codes (either from http://colorbrewer2.org/ or any other resource):

```
p +
    scale_color_manual(values = c("#e41a1c", "#377eb8", "#4daf4a", "#984ea3",
```

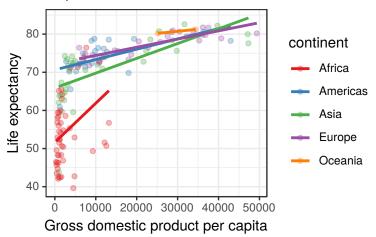


Note that  $\t\t http://colorbrewer2.org/$  also has options for Colourblind safe and Print friendly.

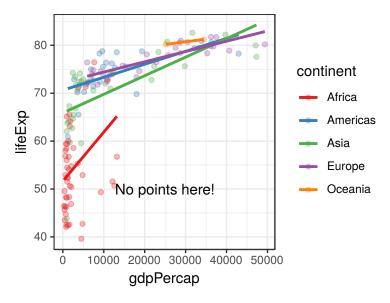
## 5.4 Titles and labels

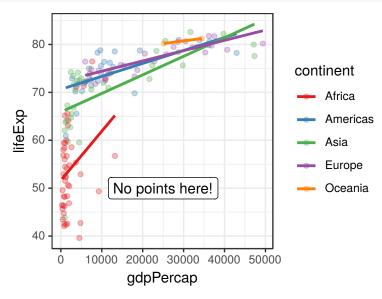
## Health and economics

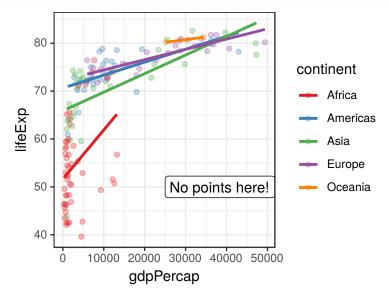
## Gapminder dataset, 2007



#### 5.4.1 Annotation





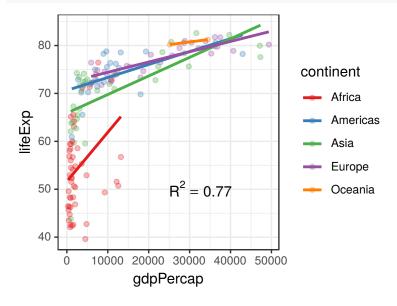


hjust stand for horizontal justification. It's default value is 0.5 (see how the label was centered at 25,000 - our chosen x location), 0 means the label goes to the right from 25,000, 1 would make it end at 25,000.

## 5.4.2 Annotation with a superscript and a variable

```
fit_glance = data.frame(r.squared = 0.7693465)

plot_rsquared = paste0(
   "R^2 == ",
```

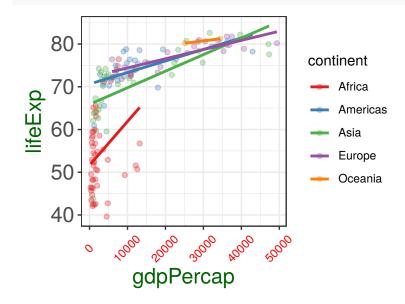


## 5.5 Text size

```
p +
   theme(axis.text.y = element_text(size = 16),
        axis.text.x = element_text(colour = "red", angle = 45, vjust = 0.5),
```

5.5 Text size 103

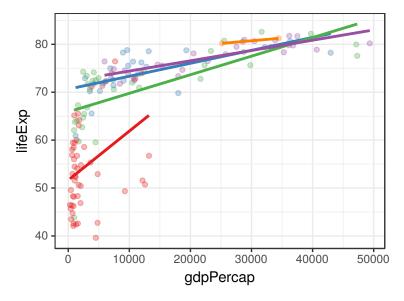
```
axis.title = element_text(size = 16, colour = "darkgreen")
)
```



## 5.5.1 Legend position

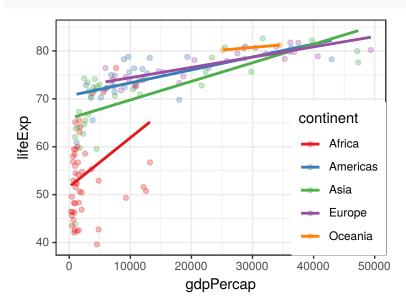
Use the following words: "right", "left", "top", "bottom", or "none" to remove the legend.

```
p +
   theme(legend.position = "none")
```



Or use relative coordinates (0-1) to give it an -y location:

```
p +
   theme(legend.position = c(1,0),
        legend.justification = c(1,0)) #bottom-right corner
```



```
p +
  theme(legend.position = "top") +
  guides(colour = guide_legend(ncol = 2))
                         Africa
                                    Europe
                         - Americas -- Oceania
             continent •
                          Asia
   80
lifeExp 60
   50
              10000
                      20000
                               30000
                                        40000
                                                50000
```

gdpPercap

# 5.6 Saving your plot

```
ggsave(p, file = "my_saved_plot.png", width = 5, height = 4)
```

# Part II Data analysis

In the second part of this book, we focus specifically on the business of data analysis. That is, formulating clear questions and seeking to answer them using available datasets.

Again, we emphasise the importance of understanding the underlying data through visualisation, rather than relying on statistical tests or, heaven forbid, the p-value alone.

There are five chapters. Testing for continuous outcome variables (6) leads naturally into Linear regression (7). We would expect the majority of actual analysis done by readers to be using the methods in chapter 7 rather than 6. Similarly, Testing for categorical outcome variables (8) leads naturally to Logistic regression (9), where we would expect the majority of work to focus. Chapters 6 and 8 however do provide helpful reminders of how to prepare data for these analyses and shouldn't be skipped. Time-to-event data introduces survival analysis and includes sections on the manipulation of dates.

# Tests for continuous outcome variables

Continuous data can be measured. Categorical data can be counted.

#### 6.1 Continuous data

Continuous data is everywhere in healthcare. From physiological measures in patients such as systolic blood pressure or pulmonary function tests, through to populations measures like life expectancy or disease incidence, the analysis of continuous outcome measures is common and important.

Our goal in most health data questions, is to draw a conclusion on a comparison between groups. For instance, understanding differences life expectancy between the year 2002 and 2007 or between the Africa and Europe, is usually more useful than simply describing the average life expectancy across the entire world across all of time.

The basis for comparisons between continuous measures is the distribution of the data. That word, as many which have a statistical flavour, brings on the sweats in a lot of people. It needn't. By distribution, we are simply referring to the shape of the data.

## 6.2 The Question

The examples in this chapter all use the data introduced previously from the amazing Gapminder project<sup>1</sup>. We will start by looking at the life expectancy of populations over time and in different geographical regions.

#### 6.3 Get the data

```
# Load packages
library(tidyverse)
library(finalfit)
library(gapminder)

# Create object mydata from object gapminder
mydata = gapminder
```

#### 6.4 Check the data

It is vital that data is carefully inspected when first read. The three functions below provide a clear summary allowing errors or miscoding to be quickly identified. It is particularity important to ensure that any missing data is identified. If you don't do this you will regret it! There are many times when an analysis has got to a

<sup>1</sup>https://www.gapminder.org/

## lifeExp

## pop

12.9

23.6

48.2

106157896.7 60011.0 2793664.0 7023595.5 19585221.8

60.7

70.8

relatively advanced stage before research realised the dataset was incomplete.

```
glimpse(mydata)
                         # each variable on a single line, variable type, fi
## Observations: 1,704
## Variables: 6
## $ country <fct> Afghanistan, Afghanistan, Afghanistan, Afghanistan, ...
## $ continent <fct> Asia, Asia, Asia, Asia, Asia, Asia, Asia, Asia, Asia...
             <int> 1952, 1957, 1962, 1967, 1972, 1977, 1982, 1987, 1992...
## $ year
## $ lifeExp <dbl> 28.801, 30.332, 31.997, 34.020, 36.088, 38.438, 39.8...
            <int> 8425333, 9240934, 10267083, 11537966, 13079460, 1488...
## $ gdpPercap <dbl> 779.4453, 820.8530, 853.1007, 836.1971, 739.9811, 78...
missing_glimpse(mydata) # missing data for each variable
##
            label var_type
                             n missing_n missing_percent
## country
              country
                         <fct> 1704
                                           0
                                                      0.0
## continent continent
                          <fct> 1704
                                           0
                                                      0.0
## year
                year
                        <int> 1704
                                          0
                                                      0.0
## lifeExp
                                                      0.0
              lifeExp
                         <dbl> 1704
                                           0
## pop
                        <int> 1704
                                                      0.0
                 pop
## gdpPercap gdpPercap
                         <dbl> 1704
                                                      0.0
                                           0
ff_glimpse(mydata)
                         # summary statistics for each variable
## Continuous
##
            label var type
                             n missing_n missing_percent
## year
              year
                      <int> 1704
                                      0
                                                0.0
                                                       1979.5
## lifeExp
             lifeExp
                        <dbl> 1704
                                        0
                                                   0.0
                                                           59.5
## pop
               pop
                     <int> 1704
                                     0
                                                0.0 29601212.3
## gdpPercap gdpPercap
                         <dbl> 1704
                                          0
                                                    0.0
                                                           7215.3
##
                                        median quartile 75
                     min quartile 25
                17.3 1952.0
                                1965.8
                                         1979.5
                                                    1993.2
## year
```

```
9857.5 241.2
                                   1202.1
                                            3531.8
## gdpPercap
                                                       9325.5
##
                       max
## year
                    2007.0
## lifeExp
                      82.6
              1318683096.0
## pop
## gdpPercap
                  113523.1
##
## Categorical
##
             label var_type
                              n missing_n missing_percent levels_n
## country
              country
                        <fct> 1704
                                         0
                                                    0.0
                                                           142
                          <fct> 1704
## continent continent
                                          0
                                                     0.0
                                                              5
##
                                                    levels
## country
## continent "Africa", "Americas", "Asia", "Europe", "Oceania"
##
                  levels_count
                                           levels_percent
## country
## continent 624, 300, 396, 360, 24 36.6, 17.6, 23.2, 21.1, 1.4
```

As can be seen, there are 6 variables, 4 are continuous and 2 are categorical. The categorical variables are already identified as factors. There are no missing data.

#### 6.5 Plot the data

We will start by comparing life expectancy between the 5 continents of the world in two different years. Always plot your data first. Never skip this step! We are particularly interested in the distribution. There's that word again. The shape of the data. Is it normal? Is it skewed? Does it differ between regions and years?

There are three useful plots which can help here:

- Histograms: examine shape of data and compare groups;
- Q-Q plots: are data normally distributed?
- Box-plots: identify outliers, compare shape and groups.

6.5 Plot the data 115

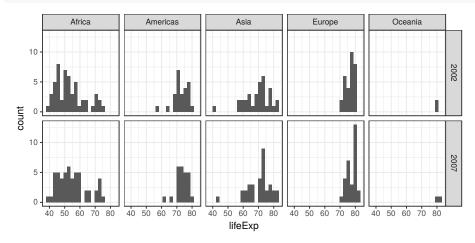
#### 6.5.1 Histogram

```
mydata %>%

filter(year %in% c(2002, 2007)) %>% # only 2002 and 2007

ggplot(aes(x = lifeExp)) + # x-axis is life expectancy

geom_histogram(bins = 20) + # histogram with data chopped into
facet_grid(year ~ continent) # create grid of year by continent
```



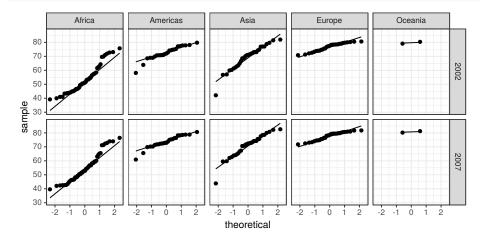
```
# add scale = "free" to facet_grid to
```

What can we see? That life expectancy in Africa is lower than in other regions. That we have little data for Oceania given there are only two countries included, Australia and New Zealand. That Africa and Asia have great variability in life expectancy by country than in the Americas or Europe. That the data follow a reasonably normal shape, with Africa 2002 a little right skewed.

#### 6.5.2 Q-Q plot

A quantile-quantile sounds complicated but is not. It is simply a graphical method for comparing the distribution (think shape) of our own data to a theoretical distribution, such as the normal distribution. In this context, quantiles are just cut points which divide our data into bins each containing the same number of observations. For example, if we have the life expectancy for 100 countries, then quartiles (note the quar-) for life expectancy are the three ages which split the observations into 4 groups each containing 25 countries. A Q-Q plot simply plots the quantiles for our data against the theoretical quantiles for a particular distributions (default below is normal). If our data follow that distribution (e.g. normal), then we get a 45 degree line on the plot.

```
mydata %>%
  filter(year %in% c(2002, 2007)) %>% # only 2002 and 2007
  ggplot(aes(sample = lifeExp)) + # Q-Q plot requires `sample`, rather
  geom_qq() + # geom for qq plot, defaults to norm
  geom_qq_line() + # add 45 degree line
  facet_grid(year ~ continent)
```



What can we see. We are looking to see if the data follow the 45 degree line which is included in the plot. These do reasonably, except for Africa which is curved upwards at each end, suggesting a skew.

We are frequently asked about performing a hypothesis test to check the assumption of normality, such as the Shapiro-Wilk normality test. We do not recommend this, simply because it is often 6.5 Plot the data 117

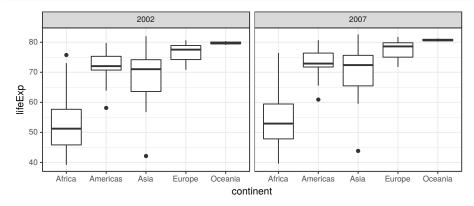
non-significant when the number of observations is small but the data look skewed, and often significant when the number of observations is high but the data look reasonably normal on inspection of plots. It is therefore not useful in practice - common sense should prevail.

#### 6.5.3 Boxplot

Boxplots are our preferred method for comparing a continuous variable such as life expectancy with a categorical explanatory variable. It is much better than a bar plot, or a bar plot with error bars, sometimes called a dynamite plot.

The box represents the median and interquartile range (where 50% of the data sits). The lines (whiskers) by default are 1.5 times the interquartile range. Outliers are represented as points.

Thus it contains information, not only on central tenancy (median), but on the variation in the data and the distribution of the data, for instance a skew should be obvious.

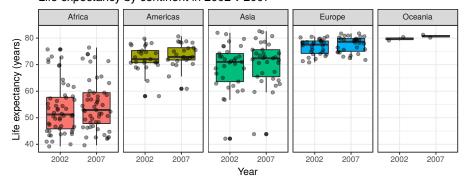


What can we see? The median life expectancy is lower in Africa than in any other continent. The variation in life expectancy is greatest in Africa and smallest in Oceania. The data in Africa looks skewed, particularly in 2002 - the lines/whiskers are unequal lengths.

We can add further arguments

```
mydata %>%
  filter(year %in% c(2002, 2007)) %>%
                                                          # only 2002 and 200
 ggplot(aes(x = factor(year), y = lifeExp)) +
                                                          # specify x and y of
  geom_boxplot(aes(fill = continent)) +
                                                          # add colour to box
  geom_jitter(alpha = 0.4) +
                                                          # add data points,
  facet_grid(. ~ continent) +
                                                          # spread by year, n
 theme(legend.position = "none") +
                                                          # remove legend
 xlab("Year") +
                                                          # adjust x-axis
 ylab("Life expectancy (years)") +
                                                          # adjust y-axis
 ggtitle("Life expectancy by continent in 2002 v 2007") # add title
```

#### Life expectancy by continent in 2002 v 2007



## 6.6 Compare the means of two groups

#### 6.6.1 T-test

A t-test is used to compare the means of two groups of continuous variables. Volumes have been written about this else where, and we won't rehearse it here.

There are various variations on the *t*-test. We will use two here. The most useful in our context is a two-sample test if independent groups (first figure). Repeated-measures data such as comparing the same countries between years can be analysed using a paired *t*-test (second figure)

#### 6.6.2 Two-sample t-tests

Referring to the first figure, let's compare life expectancy between Asia and Europe for 2007. What is imperative, is that you decide what sort of difference exists by looking at the boxplot, rather than relying on the *t*-test output. The median for Europe is clearly higher than in Asia. The distributions overlap, but it looks likely that Europe has a higher life expectancy than Asia.

##

```
## t = -4.6468, df = 41.529, p-value = 3.389e-05
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -9.926525 -3.913705
## sample estimates:
## mean in group Asia mean in group Europe
## 70.72848 77.64860
```

The Welch two-sample t-test is the most flexible and copes with differences in variance (variability) between groups, as in this example. The difference in means is provided at the bottom of the output. The t-value, degrees of freedom (df) and p-value are all provided. The p-value is 0.00003.

The base R output is not that easy to utilise. For reference, the results can be explored and exported. However, more straightforward methods are provided below.

..- attr(\*, "names")= chr "df"

```
names(ttest_result) # Names of elements of result object

## [1] "statistic" "parameter" "p.value" "conf.int" "estimate"

## [6] "null.value" "alternative" "method" "data.name"

str(ttest_result) # Details of result object

## List of 9

## $ statistic : Named num -4.65

## ..- attr(*, "names")= chr "t"

## $ parameter : Named num 41.5
```

```
## $ p.value : num 3.39e-05
## $ conf.int : atomic [1:2] -9.93 -3.91
## ..- attr(*, "conf.level")= num 0.95
## $ estimate : Named num [1:2] 70.7 77.6
## ..- attr(*, "names")= chr [1:2] "mean in group Asia" "mean in group Europe"
## $ null.value : Named num 0
## ..- attr(*, "names")= chr "difference in means"
```

```
## $ alternative: chr "two.sided"
## $ method : chr "Welch Two Sample t-test"
## $ data.name : chr "lifeExp by continent"
## - attr(*, "class")= chr "htest"
```

```
ttest_result$p.value # Extracted element of result object
```

```
## [1] 3.38922e-05
```

The broom package provides useful methods for 'tidying' common model outputs into a tibble.

The whole analysis can be constructed as a single piped function.

```
library(broom)
mydata %>%
  filter(year == 2007) %>%
                                                   # Filter for 2007 only
  filter(continent %in% c("Asia", "Europe")) %>% # Filter for Asia and Europe
  t.test(lifeExp ~ continent, data = .) %>%
  tidy()
## # A tibble: 1 x 10
   estimate estimate1 estimate2 statistic p.value parameter conf.low
##
      <dbl>
              <dbl>
                       <dbl>
                                <dbl> <dbl>
                                                <dbl>
                                                        <dbl>
      -6.92
               70.7
                       77.6
                               -4.65 3.39e-5
                                                41.5
                                                       -9.93
```

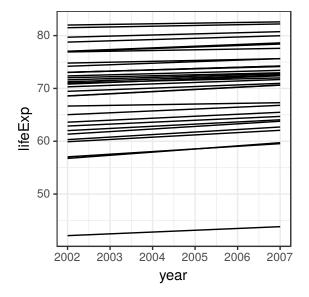
#### 6.6.3 Paired t-tests

alternative <chr>>

Consider that we want to compare the difference in life expectancy in Asian countries between 2002 and 2007. The overall difference is not impressive in the boxplot.

## # ... with 3 more variables: conf.high <dbl>, method <chr>,

We can plot differences at the country level directly.



What is the difference in life expectancy for each individual country. We don't usually have to produce this directly, but here is one method.

```
paired_table = paired_data %>%  # Save object with name paired_data
select(country, year, lifeExp) %>%  # Select variables of interest
spread (year, lifeExp) %>%  # Make wide table
mutate(
   dlifeExp = `2007` - `2002`  # Calculate difference in means
)
```

```
paired_table
## # A tibble: 33 x 4
                         `2002`
                                `2007` dlifeExp
##
      country
      <fct>
##
                          <dbl>
                                 <dbl>
                                           <dbl>
##
    1 Afghanistan
                          42.1
                                  43.8
                                           1.70
    2 Bahrain
                                  75.6
##
                          74.8
                                           0.84
##
    3 Bangladesh
                          62.0
                                  64.1
                                           2.05
##
    4 Cambodia
                                  59.7
                          56.8
                                           2.97
    5 China
                          72.0
                                  73.0
                                           0.933
##
   6 Hong Kong, China
                          81.5
                                  82.2
                                           0.713
##
    7 India
                          62.9
                                  64.7
                                           1.82
##
   8 Indonesia
                          68.6
                                  70.6
                                           2.06
##
    9 Iran
                          69.5
                                  71.0
                                           1.51
                                  59.5
                                           2.50
## 10 Iraq
                          57.0
## # ... with 23 more rows
# Mean of difference in years
paired_table %>% summarise( mean(dlifeExp) )
## # A tibble: 1 x 1
##
     `mean(dlifeExp)`
##
                 <dbl>
## 1
                  1.49
```

On average, therefore, there is an increase in life expectancy of 1.5 years in Asian countries between 2002 and 2007. Let's test whether this number differs from zero with a paired t-test.

```
paired_data %>%
    t.test(lifeExp ~ year, data = .) # Include paired = TRUE

##
## Welch Two Sample t-test
##
```

```
## data: lifeExp by year
## t = -0.74294, df = 63.839, p-value = 0.4602
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -5.513722 2.524510
## sample estimates:
## mean in group 2002 mean in group 2007
## 69.23388 70.72848
```

The results show a highly significant difference. As an exercise you can repeat this analysis simply comparing the means in an unpaired manner. The resulting p-value is R paired\_data %>% t.test(lifeExp ~ year, data = .)\$p.value. Why is there such a difference between the two approaches? This emphasises just how important it is to plot the data first. The average difference of 1.5 years is highly consistent between countries, as show on the line plot, and this differs from zero. It is up to you the investigator to interpret the effect size of 1.5 y in reporting the finding.

## 6.7 Compare the mean of one group

#### 6.7.1 One sample *t*-tests

We can use a t-test to determine whether the mean of a distribution is different to a specific value.

The paired t-test above is is equivalent to a one-sample t-test on the calculated difference in life expectancy being different to zero.

# t.test(paired\_table\$dlifeExp)

```
##
## One Sample t-test
##
## data: paired_table$dlifeExp
```

```
## t = 14.338, df = 32, p-value = 1.758e-15
## alternative hypothesis: true mean is not equal to 0
## 95 percent confidence interval:
## 1.282271 1.706941
## sample estimates:
## mean of x
## 1.494606
```

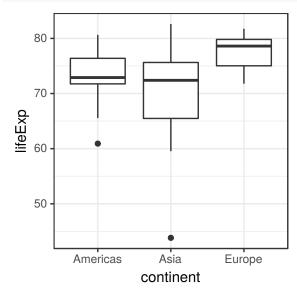
We can compare to values other than zero. For instance, we can test whether the mean life expectancy in each continent was significantly different to 77 years in 2007. We have included some extra code here to demonstrate how to run multiple base R tests in one pipe function.

```
mydata %>%
  filter(year == 2007) %>%
                                      # filter for 2007 only
                                      # split data into chunks by continent
  group_by(continent) %>%
                                      # dplyr function useful in this setting
  do(
                                      # compare mean lifeExp for each contine
    t.test(.$lifeExp, mu = 77) %>%
                                      # tidy into tibble
      tidy()
    )
## # A tibble: 5 x 9
## # Groups:
                continent [5]
    continent estimate statistic p.value parameter conf.low conf.high method
                                         <dbl>
##
    <fct>
               <dbl>
                        <dbl>
                                <dbl>
                                                 <dbl>
                                                          <dbl> <chr>
## 1 Africa
                54.8
                       -16.6 3.15e-22
                                                  52.1
                                                          57.5 One S~
                                            51
## 2 Americas
                 73.6
                         -3.82 8.32e- 4
                                             24
                                                  71.8
                                                           75.4 One S~
## 3 Asia
                70.7
                       -4.52 7.88e- 5
                                           32
                                                 67.9
                                                         73.6 One S~
                77.6
                         1.19 2.43e-1
                                           29
                                                 76.5
## 4 Europe
                                                          78.8 One S~
## 5 Oceania
                 80.7
                         7.22 8.77e- 2
                                             1
                                                 74.2
                                                          87.3 One S~
## # ... with 1 more variable: alternative <chr>
```

The mean life expectancy for Europe and Oceania do not differ from 77, while the others to to varying degrees. In particular, look at the confidence intervals of the tables and whether they include or exclude 77.

#### 6.8 Compare the means of more than two groups

It may be that our question is set around a hypothesis involving more than two groups. For example, we may be interested in comparing life expectancy across 3 continents such as the Americas, Europe and Asia. ### Plot the data



#### 6.8.1 ANOVA

Analysis of variance is a collection of statistical tests which can be used to test the difference in means between two or more groups.

In base R form, it produces an ANOVA table which includes an F-test. This so-called omnibus test tells you whether there are any differences in the comparison of means of the included groups. Again, it is important to plot carefully and be clear what question you are asking.

```
aov_data = mydata %>%
  filter(year == 2007) %>%
  filter(continent %in% c("Americas", "Europe", "Asia"))

fit = aov(lifeExp ~ continent, data = aov_data)
  summary(fit)
```

```
## Df Sum Sq Mean Sq F value Pr(>F)

## continent 2 755.6 377.8 11.63 3.42e-05 ***

## Residuals 85 2760.3 32.5

## ---

## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
```

We can conclude from this, that there is a difference in the means between at least two pairs of the included continents. As above, the output can be neatened up using the tidy function.

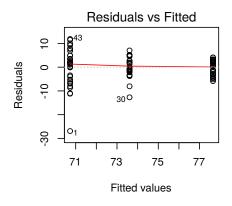
```
library(broom)
mydata %>%
  filter(year == 2007) %>%
  filter(continent %in% c("Americas", "Europe", "Asia")) %>%
  aov(lifeExp~continent, data = .) %>%
  tidy()
```

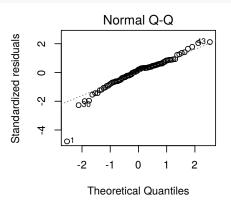
```
## # A tibble: 2 x 6
##
                   df sumsq meansq statistic
     term
                                                  p.value
##
     <chr>
                <dbl> <dbl>
                              <dbl>
                                         <dbl>
                                                     <dbl>
## 1 continent
                    2
                      756.
                              378.
                                         11.6
                                                0.0000342
## 2 Residuals
                   85 2760.
                               32.5
                                         NA
                                               NA
```

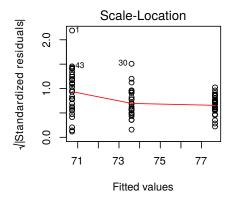
#### 6.8.2 Assumptions

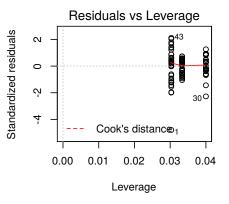
As with the normality assumption of the t-test, there are assumptions of the ANOVA model). These are covered in detail in the linear regression chapter and will not be repeated here. Suffice to say that diagnostic plots can be produced to check that the assumptions are fulfilled.

```
par(mfrow=c(2,2))
plot(fit)
```









```
par(mfrow=c(1,1))
```

#### 6.8.3 Pairwise testing and multiple comparisons

When the F-test is significant, we will often want to proceed to try and determine where the differences lie. This should of course be obvious from the boxplot you have made. However, some are fixated on the p-value!

```
##
##
    Pairwise comparisons using t tests with pooled SD
##
##
          aov data$lifeExp and aov data$continent
   data:
##
##
          Americas Asia
          0.180
## Asia
## Europe 0.031
                    1.9e-05
##
## P value adjustment method: bonferroni
```

A matrix of pairwise p-values is produced. Here we can that there is good evidence of a difference in means between Europe and Asia.

The p-values are corrected for multiple comparisons. When performing a hypothesis test at the 5% level (alpha = 0.05), there is a 5% chance of a type 1 error. That is, a 1 in 20 chance of concluding a difference exists when it in fact does not (formally, this is rejection of a true null hypothesis). As more simultaneous statistical tests are performed, the chance of a type 1 error increases.

There are three approaches to this. The first, is to no perform any correction at all. Some advocate that the best approach is simply to present the results of all the tests that were performed, and let the sceptical reader make adjustments themselves. This is attractive, but presupposes a sophisticated readership who will take the time to consider the results in their entirety.

The second and classical approach, is to control for the so-called

##

data:

family-wise error rate. The "Bonferroni" correction is probably the most famous and most conservative, where the threshold for significance is lowered in proportion to the number of comparisons made. For example, if three comparisons are made, the threshold for significance is lowered to 0.017. Equivalently, any particular p-value can be multiples by 3 and the value compared to a threshold of 0.05, as is done above. The Bonferroni method is particular conservative, meaning that type 2 errors may occur (failure to identify true differences, or false negatives) in favour or minimising type 1 errors (false positives).

The third newer approach controls false-discovery rate. The development of these methods has been driven in part by the needs of areas of science where many different statistical tests are performed at the same time, for instance, examining the influence of 1000 genes simultaneously. In these hypothesis-generating settings, a higher tolerance to type 1 errors may be preferable to missing potential findings through type 2 errors. You can see in our example, that the p-values are lower with the fdr correction when compared to the Bonferroni correction.

```
p.adjust.method = "fdr")

##

## Pairwise comparisons using t tests with pooled SD
##
```

aov data\$lifeExp and aov data\$continent

pairwise.t.test(aov data\$lifeExp, aov data\$continent,

```
##
## Americas Asia
## Asia 0.060 -
## Europe 0.016 1.9e-05
##
## P value adjustment method: fdr
```

Try not to get too hung up on this. Be sensible. Plot the data and look for differences. Focus on effect size, for instance, the actual difference in life expectancy in years, rather than the p-value of

a comparison test. Choose a method which fits with your overall aims. If you are generating hypotheses which you will proceed to test with other methods, the fdr approach may be preferable. If you are trying to capture robust effect and want to minimise type 2 errors, use a family-wise approach.

#### 6.9 Non-parametric data

What if your data is different shape to normal or the ANOVA assumptions are not fulfilled (see linear regression chapter). As always, be sensible! Would your data be expected to be normally distributed given the data-generating process? For instance, if you examining length of hospital stay it is likely that your data are highly right skewed - most patients are discharged from hospital in a few days while a smaller number stay for a long time. Is a comparison of means ever going to be the correct approach here? Perhaps you should consider a time-to-event analysis for instance (see chapter x).

If a comparison of means approach is reasonable, but the normality assumption are not fulfilled there are two approaches,

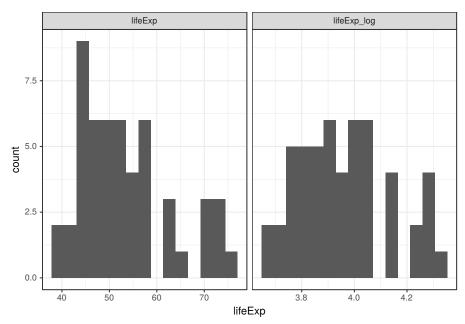
- 1. Transform the data;
- 2. Perform non-parametric tests.

#### 6.9.1 Transforming data

Remember, the Welch *t*-test is reasonably robust to divergence from the normality assumption, so small deviations can be safely ignored.

Otherwise, the data can be transformed to another scale to deal with a skew. A natural log scale is most common.

```
africa_data = mydata %>%
                                          # new object africa_data
  filter(year == 2002) %>%
                                          # only 2002
  filter(continent == "Africa") %>%
                                          # only Africa
                                          # only these variables
  select(country, lifeExp) %>%
  mutate(
                                          # log life expectancy
    lifeExp_log = log(lifeExp)
  )
head(africa_data)
                                          # inspect
## # A tibble: 6 x 3
                  lifeExp lifeExp_log
     country
##
     <fct>
                    <dbl>
                                 <dbl>
## 1 Algeria
                     71.0
                                  4.26
## 2 Angola
                     41.0
                                  3.71
## 3 Benin
                     54.4
                                 4.00
## 4 Botswana
                     46.6
                                  3.84
## 5 Burkina Faso
                                  3.92
                     50.6
## 6 Burundi
                     47.4
                                  3.86
africa_data %>%
                                          # gather original & log vals to saw
  gather(key, lifeExp, -country) %>%
  ggplot(aes(x = lifeExp)) +
  geom_histogram(bins = 15) +
                                          # make histogram
  facet_grid(. ~ key, scales = "free")
                                          # facet by original/log, axes free
```



This has worked well here. The right skew on the Africa data has been dealt with by the transformation. A parametric test such as a *t*-test can now be performed.

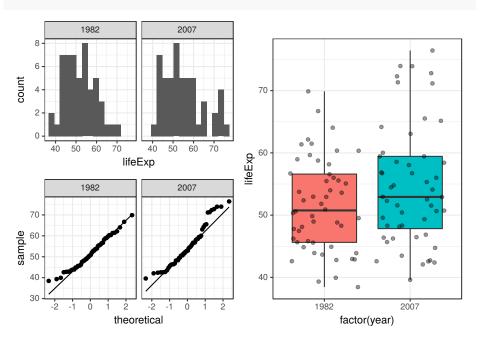
#### 6.9.2 Non-parametric test for comparing two groups

The Mann-Whitney U test is also called the Wilcoxon rank-sum test and uses a rank-based method to compare two groups (note the Wilcoxon signed-rank test is for paired data). We can use if to test for a difference in life expectancies for African countries between 1982 and 2007. Let's do a histogram, Q-Q plot and boxplot first.

```
africa_plot = mydata %>%
  filter(year %in% c(1982, 2007)) %>%  # Filter for 1982 and 2007 of filter(continent %in% c("Africa"))  # Filter for Africa only

p1 = africa_plot %>%  # Plot saved as object p1
  ggplot(aes(x = lifeExp)) +
```

```
geom_histogram(bins = 15) +
                                                 # bins can be adjusted base
  facet_grid(. ~ year)
p2 = africa_plot %>%
                                                 # Plot saved as object p2
  ggplot(aes(sample = lifeExp)) +
                                                 # remember `sample` for Q-Q
  geom_qq() +
  geom_qq_line() +
  facet_grid(. ~ year)
p3 = africa_plot %>%
                                                 # Plot saved as object p3
  ggplot(aes(x = factor(year), y = lifeExp)) + # Year is numeric, specify
  geom_boxplot(aes(fill = factor(year))) +
                                                 # Colour in boxplot
  geom_jitter(alpha = 0.4) +
                                                 # Include individual data po
  theme(legend.position = "none")
                                                 # Remove legend
library(patchwork)
                                                 # Great package for combining
p1 / p2 | p3
```



The data is a little skewed based on the histograms and Q-Q plots.

The difference between 1982 and 2007 is not particularly striking on the boxplot.

```
africa_plot %>%
  wilcox.test(lifeExp ~ year, data = .)

##

## Wilcoxon rank sum test with continuity correction

##

## data: lifeExp by year

## W = 1130, p-value = 0.1499

## alternative hypothesis: true location shift is not equal to 0
```

# 6.9.3 Non-parametric test for comparing more than two groups

The non-parametric equivalent to ANOVA, is the Kruskal-Wallis test. It can be used in base R, or via the finalfit package below.

```
library(broom)
mydata %>%
  filter(year == 2007) %>%
  filter(continent %in% c("Americas", "Europe", "Asia")) %>%
  kruskal.test(lifeExp~continent, data = .) %>%
    tidy()
## # A tibble: 1 x 4
                 p.value parameter method
     statistic
##
##
         <dbl>
                   <dbl>
                              <int> <chr>
## 1
        21.6 0.0000202
                            2 Kruskal-Wallis rank sum test
```

### 6.10 Finalfit approach

The finalfit package provides an easy to use interface for performing non-parametric hypothesis tests. Any number of explanatory variables can be tested against a so-called dependent variable. In this case, this is equivalent to a typical Table 1 in healthcare study.

label	levels	1982	2007	р
lifeExp	Median (IQR)	50.8 (11.0)	52.9 (11.6)	0.150
pop	Median (IQR)	5668228.5 (8218654.0)	10093310.5 (16454428.0)	0.032
gdpPercap	Median (IQR)	1323.7 (1958.9)	1452.3 (3130.6)	0.506

#### 6.11 Conclusions

Continuous data is frequently encountered in a healthcare setting. Liberal use of plotting is required to really understand the underlying data. Comparisons can easily made between two or more groups of data, but always remember what you are actually trying to analyse and don't become fixated on the p-value. In the

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next chapter, we will explore the comparison of two continuous variables together with multivariable models of datasets.

#### 6.12 Exercises

#### 6.12.1 Exercise 1

Make a histogram, Q-Q plot, and a box-plot for the life expectancy for a continent of your choice, but for all years. Do the data appear normally distributed?

#### 6.12.2 Exercise 2

- 1. Select any 2 years in any continent and perform a t-test to determine whether mean life expectancy is significantly different. Remember to plot your data first.
- 2. Extract only the p-value from your t.test() output.

#### 6.12.3 Exercise 3

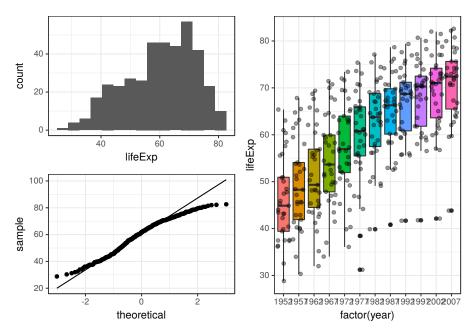
In 2007, in which continents did mean life expectancy differ from 70.

#### 6.12.4 Exercise 4

1. Use ANOVA to determine if the population changed significantly through the 1990s/2000s in individual continents.

#### 6.13 Exercise solutions

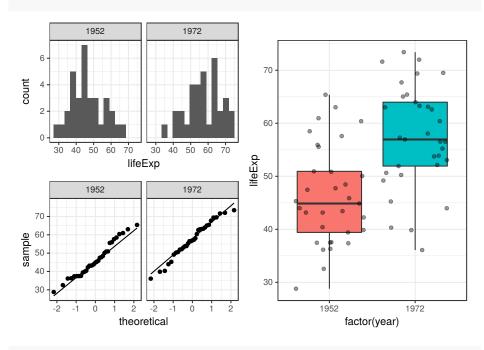
```
# Exerise 1
## Make a histogram, Q-Q plot, and a box-plot for the life expectancy for a
asia_plot = mydata %>%
  filter(continent %in% c("Asia"))
                                                # Filter for Asia only
p1 = asia_plot %>%
                                                # Plot saved as object p1
  ggplot(aes(x = lifeExp)) +
  geom_histogram(bins = 15) #+
                                                 # bins can be adjusted base
  #facet_grid(. ~ year)
                                                 # years, not included, so a
p2 = asia_plot %>%
                                                # Plot saved as object p2
  ggplot(aes(sample = lifeExp)) +
                                                # remember `sample` for Q-Q
  geom_qq() +
  geom_qq_line() #+
  #facet_grid(. ~ year)
                                                 # years not included, so all
p3 = asia_plot %>%
                                                # Plot saved as object p3
  ggplot(aes(x = factor(year), y = lifeExp)) +
                                                # Year is numeric, specify
  geom_boxplot(aes(fill = factor(year))) +
                                                # Colour in boxplot
  geom_jitter(alpha = 0.4) +
                                                # Include individual data po
  theme(legend.position = "none")
                                                # Remove legend
library(patchwork)
                                                # Great package for combining
p1 / p2 | p3
```



```
# Exercise 2
## Select any 2 years in any continent and perform a *t*-test to determine
## Remember to plot your data first.
asia_years = mydata %>%
  filter(continent %in% c("Asia")) %>%
                                                # filter for Asia only
  filter(year %in% c(1952, 1972))
                                                # filter for 1952 and 1972
p1 = asia_years %>%
                                                # Plot saved as object p1
  ggplot(aes(x = lifeExp)) +
  geom_histogram(bins = 15) +
                                                # bins can be adjusted base
  facet_grid(. ~ year)
p2 = asia_years %>%
                                                # Plot saved as object p2
  ggplot(aes(sample = lifeExp)) +
                                                # remember `sample` for Q-Q
  geom_qq() +
  geom_qq_line() +
  facet_grid(. ~ year)
                                               # years not included, so all
```

```
p3 = asia_years %>% # Plot saved as object p3
ggplot(aes(x = factor(year), y = lifeExp)) + # Year is numeric, specify
geom_boxplot(aes(fill = factor(year))) + # Colour in boxplot
geom_jitter(alpha = 0.4) + # Include individual data pa
theme(legend.position = "none") # Remove legend

library(patchwork) # Great package for combining
p1 / p2 | p3
```



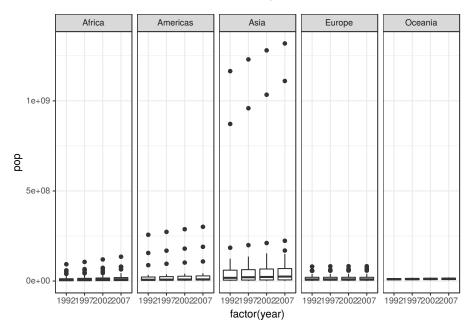
```
asia_years %>%
t.test(lifeExp ~ year, data = .)
```

```
##
## Welch Two Sample t-test
##
## data: lifeExp by year
## t = -4.7007, df = 63.869, p-value = 1.428e-05
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
```

```
6.13 Exercise solutions
```

```
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```

```
## -15.681981 -6.327769
## sample estimates:
## mean in group 1952 mean in group 1972
##
             46.31439
                                57.31927
# Exercise 3
## In 2007, in which continents did mean life expectancy differ from 70.
mydata %>%
  filter(year == 2007) %>%
  group_by(continent) %>%
  do(
    t.test(.$lifeExp, mu = 70) %>% tidy()
                                               # It's somtimes a little awi
  )
## # A tibble: 5 x 9
## # Groups:
              continent [5]
## continent estimate statistic p.value parameter conf.low conf.high method
## <fct>
              <dbl>
                      <dbl>
                             <dbl>
                                       <dbl>
                                              <dbl>
                                                       <dbl> <chr>
## 1 Africa
               54.8 -11.4 1.33e-15
                                               52.1
                                                        57.5 One S~
                                          51
## 2 Americas
                73.6
                       4.06 4.50e-4
                                          24
                                               71.8
                                                       75.4 One S~
               70.7
                      0.525 6.03e-1
                                              67.9
## 3 Asia
                                         32
                                                      73.6 One S~
                                                      78.8 One S~
## 4 Europe
               77.6 14.1 1.76e-14
                                          29
                                               76.5
## 5 Oceania
                80.7 20.8 3.06e-2
                                          1
                                               74.2
                                                       87.3 One S~
## # ... with 1 more variable: alternative <chr>
# Exercise 4
## Use Kruskal-Wallis to determine if the mean population changed significa
mydata %>%
  filter(year >=1990) %>%
  ggplot(aes(x = factor(year), y = pop)) +
  geom_boxplot() +
  facet_grid(. ~ continent)
```



```
mydata %>%
  filter(year >=1990) %>%
  group_by(continent) %>%
  do(
    kruskal.test(pop ~ factor(year), data = .) %>% tidy()
)
```

```
## # A tibble: 5 x 5
               continent [5]
## # Groups:
     continent statistic p.value parameter method
##
     <fct>
                          <dbl>
                   <dbl>
                                      <int> <chr>
## 1 Africa
                2.10 0.553
                                  3 Kruskal-Wallis rank sum test
## 2 Americas
                0.847 0.838
                                   3 Kruskal-Wallis rank sum test
## 3 Asia
               1.57 0.665
                                 3 Kruskal-Wallis rank sum test
## 4 Europe
               0.207 0.977
                                  3 Kruskal-Wallis rank sum test
## 5 Oceania
                1.67
                       0.644
                                  3 Kruskal-Wallis rank sum test
```

# Linear regression

#### 7.1 Data

We will be using the same gapminder dataset as in the last two sessions.

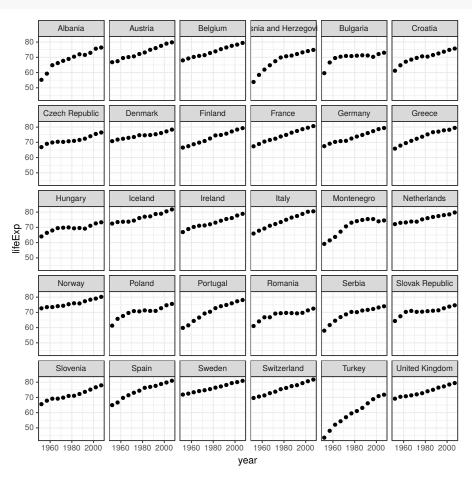
```
library(tidyverse)
library(gapminder) # dataset
library(lubridate) # handles dates
library(broom) # transforms statistical output to data frame
mydata = gapminder
```

#### 7.2 Plotting

Let's plot the life expectancies in European countries over the past 60 years:

```
mydata %>%
  filter(continent == "Europe") %>%
  ggplot(aes(x = year, y = lifeExp)) +
  geom_point() +
  facet_wrap(~country) +
```

```
theme_bw() +
scale_x_continuous(breaks = c(1960, 1980, 2000))
```



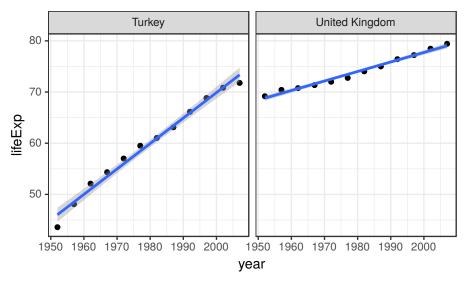
#### 7.2.1 Exercise

Save the above filter into a new variable called eurodata:

```
eurodata = mydata %>%
  filter(continent == "Europe")
```

#### 7.2.2 Exercise

Create the same plot as above (life expectancy over time), but for just Turkey and the United Kingdom, and add linear regression lines. Hint: use + geom\_smooth(method = "lm") for the lines. lm() stands for linear model.



#### 7.3 Simple linear regression

As you can see, ggplot() is very happy to run and plot linear regression for us. To access the results, however, we should save the full results of the linear regression models into variables in our Environment. We can then investigate the intercepts and the slope coefficients (linear increase per year):

```
fit_uk = mydata %>%
  filter(country == "United Kingdom") %>%
  lm(lifeExp~year, data = .) # the data=. argument is necessary
```

```
fit_turkey = mydata %>%
  filter(country == "Turkey") %>%
  lm(lifeExp~year, data = .)

fit_uk$coefficients

fit_turkey$coefficients
```

```
## (Intercept) year

## -294.1965876 0.1859657

## (Intercept) year

## -924.5898865 0.4972399
```

#### 7.3.1 Exercise

To make the intercepts more meaningful, add a new column called year\_from1952 and redo fit\_turkey and fit\_uk using year\_from1952 instead of year.

```
mydata$year_from1952 = mydata$year - 1952

fit_uk = mydata %>%
   filter(country == "United Kingdom") %>%
   lm(lifeExp~year_from1952, data = .)

fit_turkey = mydata %>%
   filter(country == "Turkey") %>%
   lm(lifeExp~year_from1952, data = .)

fit_uk$coefficients
```

fit\_uk %>% tidy()

## fit\_turkey\$coefficients

```
## (Intercept) year_from1952
## 68.8085256 0.1859657
## (Intercept) year_from1952
## 46.0223205 0.4972399
```

#### 7.3.2 Model information: summary(), tidy(), glance()

Accessing all other information about our regression model:

```
fit_uk %>% summary()
```

```
##
## Call:
## lm(formula = lifeExp ~ year_from1952, data = .)
##
## Residuals:
##
        Min
                  1Q
                       Median
                                    3Q
                                            Max
## -0.69767 -0.31962 0.06642 0.36601
                                        0.68165
##
## Coefficients:
##
                  Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                68.808526
                          0.240079 286.61 < 2e-16 ***
## year_from1952 0.185966 0.007394 25.15 2.26e-10 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 0.4421 on 10 degrees of freedom
## Multiple R-squared: 0.9844, Adjusted R-squared: 0.9829
## F-statistic: 632.5 on 1 and 10 DF, p-value: 2.262e-10
```

fit\_uk %>% glance()

```
## # A tibble: 2 x 5
##
     term
                   estimate std.error statistic
##
     <chr>>
                      <dbl>
                                 <dbl>
                                           <dbl>
                                                     <dbl>
## 1 (Intercept)
                      68.8
                               0.240
                                           287. 6.58e-21
## 2 year_from1952
                                            25.1 2.26e-10
                       0.186
                               0.00739
```

```
## # A tibble: 1 x 11
## r.squared adj.r.squared sigma statistic p.value df logLik AIC BIC
## <dbl> >10.984 0.983 0.442 633. 2.26e-10 2 -6.14 18.3 19.7
## # ... with 2 more variables: deviance <dbl>, df.residual <int>
```

#### 7.4 If you are new to linear regression

See these interactive Shiny apps provided by RStudio:

```
https://gallery.shinyapps.io/simple_regression/
https://gallery.shinyapps.io/multi_regression/
```

(library(shiny) is an R package for making your output interactive)

#### 7.4.1 Exercise - Residuals

Open the first Shiny app ("Simple regression"). Move the sliders until the red lines (residuals\*) turn green - this means you've made the line fit the points as well as possible. Look at the intercept and slope - discuss with your neighbour or a tutor what these numbers mean and how they affect the straight line on the plot.

\*Residual is how far away each point (observation) is from the

linear regression line. (In this example it's the linear regression line, but residuals are relevant in many other contexts as well.)

#### 7.5 Multiple linear regression

Multiple linear regression includes more than one predictor variable. There are a few ways to include more variables, depending on whether they should share the intercept and how they interact:

Simple linear regression (exactly one predictor variable):

```
myfit = lm(lifeExp~year, data=eurodata)
```

Multiple linear regression (additive):

```
myfit = lm(lifeExp~year+country, data=eurodata)
```

Multiple linear regression (all interactions):

```
myfit = lm(lifeExp~year*country, data=eurodata)
```

These examples of multiple regression include two variables: year and country, but we could include more by just adding them with +.

#### 7.5.1 Exercise

Open the second Shiny app ("Multiple regression") and see how:

- In simple regression, there is only one intercept and slope for the whole dataset.
- Using the additive model (lm(formula = y ~ x + group) the two lines (one for each group) have different intercepts but the same slope. However, the lm() summary seems to only include one line called "(Intercept)", how to find the intercept for the second group of points?

• Using the interactive model (lm(formula = y ~ x\*group)) the two lines have different intercepts and different slopes.

#### 7.5.2 Exercise

## 2 year from1952

Convince yourself that using an fully interactive multivariable model is similar to running several separate simple linear regression models. Remember that we calculate the life expectancy in 1952 (intercept) and improvement per year (slope) for Turkey and the United Kingdom:

```
fit_uk %>%
  tidy() %>%
  mutate(estimate = round(estimate, 2)) %>%
  select(term, estimate)
## # A tibble: 2 x 2
##
     term
                    estimate
     <chr>
##
                       <dbl>
## 1 (Intercept)
                       68.8
## 2 year_from1952
                        0.19
fit_turkey %>%
  tidy() %>%
 mutate(estimate = round(estimate, 2)) %>%
  select(term, estimate)
## # A tibble: 2 x 2
     term
                    estimate
##
     <chr>>
                       <dbl>
## 1 (Intercept)
                        46.0
```

(The lines tidy(), mutate(), and select() are only included for neater presentation here, you can use summary() instead.)

0.5

We can do this together using year\_from1952\*country in the lm():

```
mydata %>%
  filter(country %in% c("Turkey", "United Kingdom")) %>%
  lm(lifeExp ~ year_from1952*country, data = .) %>%
  tidy() %>%
  mutate(estimate = round(estimate, 2)) %>%
  select(term, estimate)
```

Now. It may seem like R has omitted Turkey but the values for Turkey are actually in the Intercept = 46.02 and in year\_from1952 = 0.50. Can you make out the intercept and slope for the UK? Are they the same as in the simple linear regression model?

#### 7.5.3 Exercise

Add a third country (e.g. "Portugal") to filter(country %in% c("Turkey", "United Kingdom")) in the above example. Do the results change?

#### 7.5.4 Optional (Advanced) Exercise

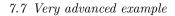
Run separate linear regression models for every country in the dataset at the same time and putting it all in two neat dataframes (one for the coefficients, one for the summary statistics):

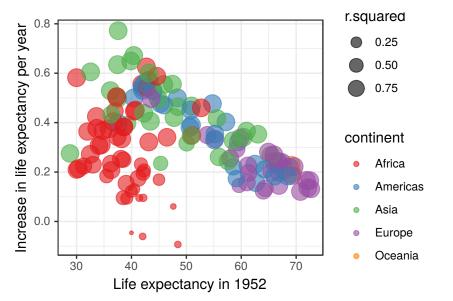
```
linfit_coefficients = mydata %>%
  group_by(country) %>%
  do(
    tidy(
       lm(lifeExp~year, data=.)
    )
)

linfit_overall = mydata %>%
  group_by(country) %>%
  do(
    glance(
       lm(lifeExp~year, data=.)
    )
)
)
```

Plot the linear regression estimate (improvement per year between 1952 - 2007), size the points by their r-squared values, and colour the points by continent (hint: you will have to join mydata, linfit\_coefficients %>% filter(term == "year"), and linfit\_overall):

```
mydata %>%
  filter(year == 1952) %>%
  full_join(linfit_coefficients %>% filter(term == "year"), by = "country")
  full_join(linfit_overall, by = "country") %>%
  ggplot(aes(x = lifeExp, y = estimate, colour = continent, size = r.squared
  geom_point(alpha = 0.6) +
  theme_bw() +
  scale_colour_brewer(palette = "Set1") +
  ylab("Increase in life expectancy per year") +
  xlab("Life expectancy in 1952")
```





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### 7.6 Very advanced example

Or you can do the above in a nested tibble/data frame:

```
nested_linreg = mydata %>%
  group_by(country) %>%
  nest() %>%
  mutate(model = purrr::map(data, ~ lm(lifeExp ~ year, data = .)))
```

#### 7.7 Solutions

#### 6.2.2

```
mydata %>%
  filter(country %in% c("United Kingdom", "Turkey") ) %>%
  ggplot(aes(x = year.formatted, y = lifeExp)) +
  geom_point() +
  facet_wrap(~country) +
  theme_bw() +
  geom_smooth(method = "lm")
```

#### 6.5.3

```
mydata %>%
  filter(country %in% c("Turkey", "United Kingdom", "Portugal")) %>%
  lm(lifeExp ~ year_from1952*country, data = .) %>%
  tidy() %>%
  mutate(estimate = round(estimate, 2)) %>%
  select(term, estimate)
```

Overall, the estimates for Turkey and the UK do not change, but Portugal becomes the reference (alphabetically first) to which you can subtract or add the relevant lines for Turkey and the UK.

# Tests for categorical variables

#### 8.1 Data

We are now changing to a new dataset, melanoma. Click on mydata in your environment and have a look at the values - you'll see that categorical variables are coded as numbers, rather than text. You will need to recode these numbers into proper factors.

```
library(tidyverse)
library(finalfit)
library(broom)
mydata = boot::melanoma
```

#### 8.1.1 Recap on factors

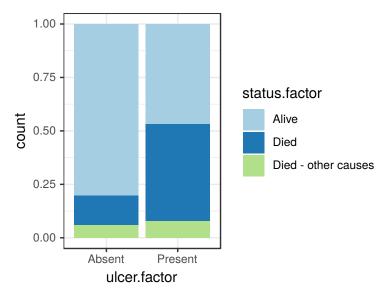
Press F1 on boot::melanoma to see its description. Use the information from help to change the numbers into proper factors (e.g. 0 - female, 1 - male).

#### 8.2 Chi-squared test / Fisher's exact test

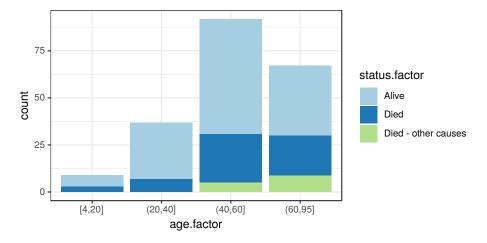
#### 8.2.1 Plotting

Always plot new data first!

```
mydata %>%
    ggplot(aes(x = ulcer.factor, fill=status.factor)) +
        geom_bar(position = "fill") +
        theme_bw() +
        scale_fill_brewer(palette = "Paired")
```

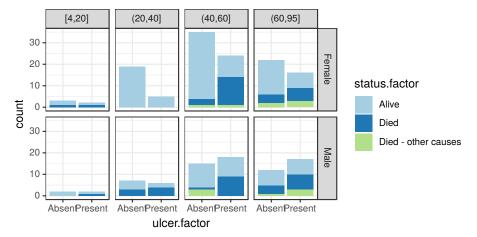


```
mydata %>%
  ggplot(aes(x = age.factor, fill = status.factor)) +
   geom_bar() +
   theme_bw() +
   scale_fill_brewer(palette = "Paired")
```



```
mydata %>%
   ggplot(aes(x = ulcer.factor, fill=status.factor)) +
```





#### 8.3 Analysis

#### 8.3.1 Using base R

First lets group together those that 'died of another cause' with those 'alive', to give a disease-specific mortality variable (fct\_collapse() will help us).

```
mydata$status.factor %>%
   fct_collapse("Alive" = c("Alive", "Died - other causes")) ->
   mydata$status.factor
```

Let's test mortality against sex.

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```
table(mydata$status.factor, mydata$sex.factor)
```

```
## Female Male
## Alive 98 50
## Died 28 29
```

```
chisq.test(mydata$status.factor, mydata$sex.factor)
```

```
##
## Pearson's Chi-squared test with Yates' continuity correction
##
## data: mydata$status.factor and mydata$sex.factor
## X-squared = 4.3803, df = 1, p-value = 0.03636
```

Note that chisq.test() defaults to the Yates' continuity correction.

It is fine to use this, but if you have a particular need not to, turn if off with chisq.test(mydata\$status.factor, mydata\$sex.factor, correct=FALSE).

#### 8.3.2 Using CrossTable

This gives lots of useful information. It is readable in R and has lots of options, including Fisher's exact test. It is not that easy to extract results.

```
library(gmodels)
# F1 CrossTable to see options
CrossTable(mydata$status.factor, mydata$sex.factor, chisq=TRUE)
##
##
##
      Cell Contents
##
## | Chi-square contribution |
                N / Row Total |
## |
## |
                N / Col Total |
             N / Table Total |
##
##
##
## Total Observations in Table:
                                    205
##
##
##
                          | mydata$sex.factor
                                          Male | Row Total |
                             Female |
## mydata$status.factor |
##
                Alive |
                               98 |
                                           50 |
                            0.544 |
                                        0.868 |
##
##
                           0.662 |
                                       0.338 |
##
                            0.778
                                        0.633 |
##
                            0.478 |
                                        0.244
                                28 I
                                           29 I
##
                 Died |
                                                       57
##
                            1.412
                                        2.253 l
##
                           0.491 |
                                       0.509 |
                                                    0.278 |
##
                            0.222 |
                                        0.367 |
##
                            0.137 |
                                        0.141 |
                               126 |
                                           79 |
                                                      205 I
##
          Column Total |
                            0.615 |
##
                                        0.385 |
```

```
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                                            161
## -----|-----|
##
##
## Statistics for All Table Factors
##
##
## Pearson's Chi-squared test
## Chi^2 = 5.076334
                    d.f. = 1 p = 0.0242546
##
## Pearson's Chi-squared test with Yates' continuity correction
## Chi^2 = 4.380312 d.f. = 1 p = 0.03635633
##
##
```

#### 8.3.3 Exercise

Use the 3 methods (table, chisq.test, CrossTable) to test status.factor against ulcer.factor.

```
table(mydata$status.factor, mydata$ulcer.factor)
chisq.test(mydata$status.factor, mydata$ulcer.factor)
```

Using CrossTable

```
CrossTable(mydata$status.factor, mydata$ulcer.factor, chisq=TRUE)
```

#### 8.3.4 Fisher's exact test

An assumption of the chi-squared test is that the 'expected cell count' is greater than 5. If it is less than 5 the test becomes unreliable and the Fisher's exact test is recommended.

Run the following code.

```
library(gmodels)
CrossTable(mydata$status.factor, mydata$age.factor, expected=TRUE, chisq=TRU
## Warning in chisq.test(t, correct = FALSE, ...): Chi-squared approximation
## may be incorrect
##
##
##
      Cell Contents
##
                            N |
##
                  Expected N |
    Chi-square contribution |
##
##
               N / Row Total |
##
               N / Col Total |
##
             N / Table Total |
##
##
## Total Observations in Table:
                                  205
##
##
##
                         | mydata$age.factor
                            [4,20] | (20,40] | (40,60] | (60,95] | Row Total
## mydata$status.factor |
                     ##
             Alive |
                          6 |
                                  30 |
                                            66 |
                                                     46 |
                                                              148 |
##
                      6.498 |
                               26.712 |
                                          66.420 |
                                                     48.371 |
##
                      0.038 |
                                0.405 |
                                          0.003 |
                                                     0.116 |
##
                     0.041 |
                                0.203 |
                                          0.446 |
                                                     0.311 |
                                                               0.722 \, \mathsf{I}
##
                      0.667 |
                                          0.717 |
                                                     0.687 |
                                0.811 |
##
                      0.029
                                0.146 |
                                           0.322 |
                                                     0.224
##
##
             Died |
                          3 |
                                   7 |
                                           26 |
                                                     21 |
                                                              57 |
##
                      2.502
                               10.288 |
                                          25.580 |
                                                     18.629 |
                      0.099 |
                                                     0.302 |
##
                                1.051
                                          0.007 |
##
                      0.053 |
                                0.123 |
                                          0.456 |
                                                     0.368 |
                                                               0.278 |
```

```
##
                    0.333 |
                             0.189 |
                                       0.283 |
                                                0.313 |
##
                    0.015
                             0.034 |
                                       0.127 |
                                                0.102 |
       Column Total |
                          9 |
                                  37 |
                                          92 I
                                                   67 |
                                                           205 |
                                                 0.327 |
##
                    0.044
                             0.180 |
                                       0.449 |
          ##
##
## Statistics for All Table Factors
##
##
## Pearson's Chi-squared test
                        d.f. = 3 p = 0.5682975
## Chi^2 = 2.019848
##
##
##
Why does it give a warning? Run it a second time including
fisher=TRUE.
library(gmodels)
CrossTable(mydata$status.factor, mydata$age.factor, expected=TRUE, chisq=TRU
## Warning in chisq.test(t, correct = FALSE, ...): Chi-squared approximation
## may be incorrect
##
##
##
     Cell Contents
## |
## |
                 Expected N |
## | Chi-square contribution |
## |
              N / Row Total |
## |
              N / Col Total |
            N / Table Total |
```

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d.f. = 3 p = 0.5682975

## Chi<sup>2</sup> = 2.019848

## ## ##

### 8.4 Summarising multiple factors (optional)

CrossTable is useful for summarising single variables. We often want to summarise more than one factor or continuous variable against our dependent variable of interest. Think of Table 1 in a journal article.

### 8.5 Summarising factors with library(finalfit)

This is our own package which we have written and maintain. It contains functions to summarise data for publication tables and figures, and to easily run regression analyses. We specify a dependent or outcome variable, and a set of explanatory or predictor variables.

```
library(finalfit)
mydata %>%
  summary_factorlist(dependent = "status.factor",
                      explanatory = c("sex.factor", "ulcer.factor", "age.fact
                      p = TRUE,
                      column = TRUE)
## Warning in chisq.test(tab, correct = FALSE): Chi-squared approximation may
## be incorrect
##
            label
                   levels
                               Alive
                                          Died
                                                     р
## 5
       sex.factor
                  Female 98 (66.2) 28 (49.1)
                                                 0.024
## 6
                      Male 50 (33.8) 29 (50.9)
## 7 ulcer.factor Absent 99 (66.9) 16 (28.1) <0.001
## 8
                  Present 49 (33.1) 41 (71.9)
                             6 (4.1)
                                       3 (5.3)
## 1
       age.factor
                    [4,20]
                                                0.568
                                     7 (12.3)
## 2
                   (20,40] 30 (20.3)
```

```
## 3 (40,60] 66 (44.6) 26 (45.6)
## 4 (60,95] 46 (31.1) 21 (36.8)
```

#### 8.5.1 Summarising factors with library(tidyverse)

#### **8.5.2** Example

Tidyverse gives the flexibility and power to examine millions of rows of your data any way you wish. The following are intended as an extension to what you have already done. These demonstrate some more advanced approaches to combining tidy functions.

```
# Calculate number of patients in each group
counted_data = mydata %>%
    count(ulcer.factor, status.factor)

# Add the total number of people in each status group
counted_data2 = counted_data %>%
    group_by(status.factor) %>%
    mutate(total = sum(n))
```

```
# Calculate the percentage of n to total
counted_data3 = counted_data2 %>%
  mutate(percentage = round(100*n/total, 1))
```

Create a combined columns of both n and percentage using paste() to add brackets around the percentage.

```
counted_data4 = counted_data3 %>%
  mutate(count_perc = paste0(n, " (", percentage, ")"))
```

Or combine everything together without the intermediate counted\_data breaks.

```
mydata %>%
  count(ulcer.factor, status.factor) %>%
  group_by(status.factor) %>%
  mutate(total = sum(n)) %>%
  mutate(percentage = round(100*n/total, 1)) %>%
  mutate(count_perc = pasteO(n, " (", percentage, ")")) %>%
  select(-total, -n, -percentage) %>%
  spread(status.factor, count_perc)
```

#### 8.5.3 Exercise

By changing one and only one word at a time in the above block (the "Combine everything together" section)

Reproduce this:

```
## age.factor Alive Died

## 1 [4,20] 6 (4.1) 3 (5.3)

## 2 (20,40] 30 (20.3) 7 (12.3)

## 3 (40,60] 66 (44.6) 26 (45.6)

## 4 (60,95] 46 (31.1) 21 (36.8)
```

And then this:

```
## sex.factor Alive Died
## 1 Female 98 (66.2) 28 (49.1)
## 2 Male 50 (33.8) 29 (50.9)
```

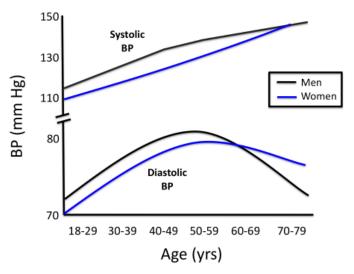
Solution: The only thing you need to change is the first variable in count(), e.g., count(age.factor, ....

# Logistic regression

### 9.1 What is Logistic Regression?

As we have seen in previously, regression analysis is a statistical process for estimating the relationships between variables. For instance, we may try to predict the blood pressure of a group of patients based on their age. As age and blood pressure are on a continuous scale, this is an example of linear regression.

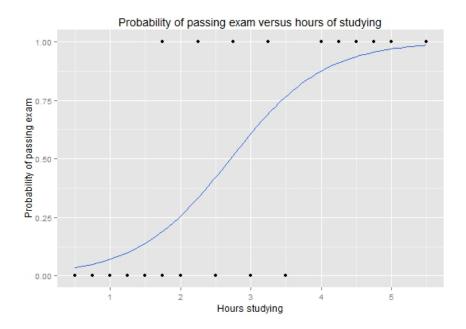
### Changes in Systolic & Diastolic BP with Age



Adapted from: JNC7 & Burt et al (1995) Hypertension 23:305-313

Logistic regression is an extension of this, where the variable being predicted is *categorical*. We will deal with binary logistic regres-

sion, where the variable being predicted has two levels, e.g. yes or no, 0 or 1. In healthcare, this is usually done for an event (like death) occurring or not occurring. Logistic regression can tell us the probability of the outcome occurring.



Logistic regression lets you adjust for the effects of confounding factors on an outcome. When you read a paper that says it has adjusted for confounding factors, this is the usual method which is used.

Adjusting for confounding factors allows us to isolate the true effect of a variable upon an outcome. For example, if we wanted to know the effects of smoking on deaths from heart attacks, we would need to also control for things like sex and diabetes, as we know they contribute towards heart attacks too.

Although in binary logistic regression the outcome must have two levels, the predictor variables (also known as the explanatory variables) can be either continuous or categorical.

Logistic regression can be performed to examine the influence of

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one predictor variable, which is known as a univariable analysis. Or multiple predictor variables, known as a multivariable analysis.

#### 9.2 Definitions

**Dependent** variable (in clinical research usually synonymous to **outcome**) - is what we are trying to explain, i.e. we are trying to identify the factors associated with a particular outcome. In binomial logistic regression, the dependent variable has exactly two levels (e.g. "Died" or "Alive", "Yes - Complications" or "No Complications", "Cured" or "Not Cured", etc.).

Explanatory variables (also known as predictors, confounding variables, or "adjusted for") - patient-level information, usually including demographics (age, gender) as well as clinical information (disease stage, tumour type). Explanatory variables can be categorical as well as continuous, and categorical variables can have more than two levels.

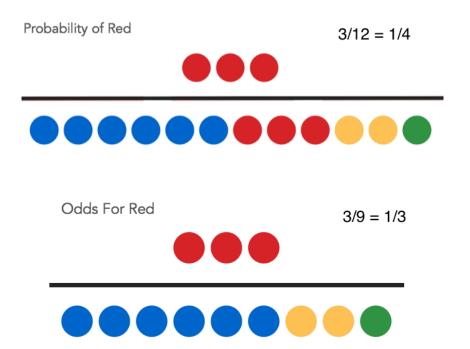
Univariable - analysis with only one Explanatory variable.

Multivariable - analysis with more than one Explanatory variable. Synonymous to "adjusted".

(Multivariate - technically means more than one **Dependent** variable (we will not discuss this type of analysis), but very often used interchangeably with Multivariable.)

### 9.3 Odds and probabilities

Odds and probabilities can get confusing so let's get them straight:



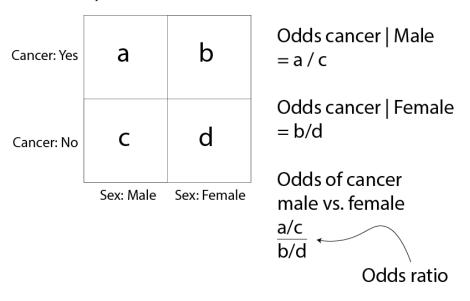
Odds and probabilities can be interconverted. For example, if the odds of a patient dying from a disease are 9 to 1 then the probability of death (also known as risk) is 10%. Odds of 1 to 1 equal 50%.

 $Odds = \frac{p}{1-p}$ , where p is the probability of the outcome occurring (or the circle being red).

Look at the numbers and convince yourself that this works.

#### 9.3.1 Odds ratios

For a given categorical explanatory variable (e.g. gender), the likelihood of an outcome/dependent occurring (e.g cancer) can be expressed in a ratio of odds or odds ratio, e.g. the odds of men developing cancer is 2-times that of females, odds ratio = 2.0.



An alternative is a ratio of probabilites, called a risk ratio or relative risk. Odds ratios have useful mathematical characteristics and are the main expression of results in logistic regression analysis.

#### 9.4 Melanoma dataset

Malignant melanoma is a cancer of the skin. It is agressive and highly invasive, making it difficult to treat.

It's classically divided into 4 stages of severity, based upon the depth of the tumour:

- Stage I: <0.5 mm depth
- Stage II: 0.5 to 1.0 mm depth
- Stage III: 1.0 to 4.0 mm depth
- Stage IV: > 4.0 mm depth

This will be important in our analysis as we will creating a new variable based upon this.

Using logistic regression, we will investigate factors associated with death from malignant melanoma.

#### 9.4.1 Doing logistic regression in R

There are a few different ways of doing logistic regression in R. The glm() function is probably the most common and most flexible one to use. (glm stands for generalised linear model.)

Within the glm() function there are several options in the function we must define to make R run a logistic regression.

data - you must define the dataframe to be used in the regression.

family - this tells R to treat the analysis as a logisitic regression. For our purposes, family will always be "binomial" (as binary data follow this distribution).

 $x \sim a + b + c$  - this is the formula for the logistic regression, with x being the outcome and a, b and c being predictor variables.

Note the outcome is separated from the rest of the formula and

sits on the left hand side of a ~. The confounding variables are on the right side, separated by a + sign.

The final glm() function takes the following form:

```
glm(x ~ a + b + c + d, data = data, family = "binomial")
```

#### 9.5 Setting up your data

The most important step to ensure a good basis to start from is to ensure your variables are well structured and your outcome variable has exactly two outcomes.

We will need to make sure our outcome variables and predictor variables (the ones we want to adjust for) are suitably prepared.

In this example, the outcome variable called status.factor describes whether patients died or not and will be our (dependent) variable of interest.

#### 9.5.1 Worked Example

```
library(tidyverse)

load("melanoma_factored.rda")
#Load in data from the previous session
```

Here status.factor has three levels: Died, Died - other causes and Alive. This is not useful for us, as logistic regression requires outcomes to be binary (exactly two levels).

We want to find out which variables predict death from melanoma. So we should create a new factor variable, died\_melanoma.factor. This will have two outcomes, Yes (did die from melanoma) or No (did not die from melanoma).

### 9.6 Creating categories

Now that we have set up our outcome variable, we should ensure our predictor variables are prepared too.

Remember the stages of melanoma? This is an important predictor of melanoma Mortality based upon the scientific literature.

We should take this into account in our model.

#### 9.6.1 Exercise

Create a new variable called stage.factor to encompass the stages of melanoma based upon the thickness. In this data, the thickness variable is measured in millimetres too.

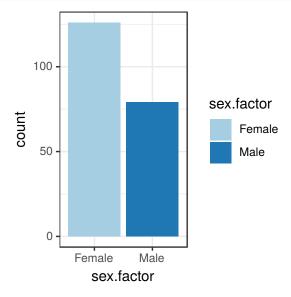
```
#the cut() function makes a continuous variable into a categorical variable
mydata$thickness %>%
    cut(breaks = c(0,0.5,1,4, max(mydata$thickness, na.rm=T)),
        include.lowest = T) ->
mydata$stage.factor
mydata$stage.factor %>% levels()
```

```
## [1] "[0,0.5]" "(0.5,1]" "(1,4]" "(4,17.4]"
```

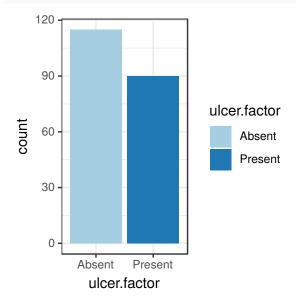
```
## [1] "Stage I" "Stage II" "Stage IV"
```

### 9.6.2 Always plot your data first!

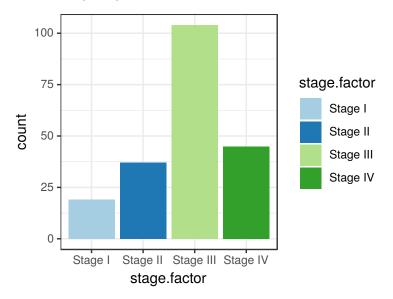
```
mydata %>%
   ggplot(aes(x = sex.factor)) +
   geom_bar(aes(fill = sex.factor))
```



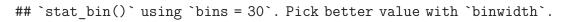
```
mydata %>%
  ggplot(aes(x = ulcer.factor)) +
  geom_bar(aes(fill = ulcer.factor))
```

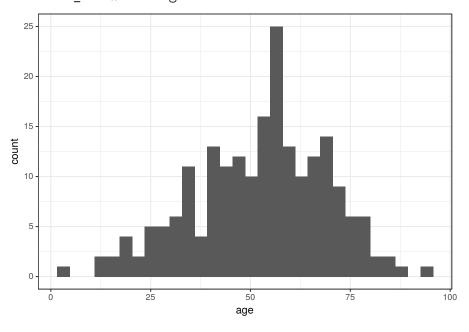


```
mydata %>%
  ggplot(aes(x = stage.factor)) +
  geom_bar(aes(fill = stage.factor))
```



```
mydata %>%
  ggplot(aes(x = age)) +
  geom_histogram(aes(fill = age))
```





Now we are ready for some modelling!

### 9.7 Basic: One explanatory variable (predictor)

Lets find out what the influence of each predictor/confounding variable is on mortality from melanoma, which may help inform a more complicated regression, with multiple predictors/confounders.

We'll start with whether the patient was male or female:

### 9.7.1 Worked example

First we need to create a regression model using glm(). We will then summarise it using summary()

Note, we need to use the family option. Specifying 'binomial' in family tells glm() to switch to logistic regression.

```
#Create a model
glm(died_melanoma.factor ~ sex.factor, data = mydata, family = "binomial")
##
## Call: glm(formula = died_melanoma.factor ~ sex.factor, family = "binomial",
       data = mydata)
##
## Coefficients:
##
      (Intercept)
                   sex.factorMale
           -1.253
                             0.708
##
## Degrees of Freedom: 204 Total (i.e. Null); 203 Residual
## Null Deviance:
                         242.4
## Residual Deviance: 237.4
                                 AIC: 241.4
```

```
model1 = glm(died_melanoma.factor ~ sex.factor, data = mydata, family = "bin
summary(model1)
##
## Call:
## glm(formula = died_melanoma.factor ~ sex.factor, family = "binomial",
       data = mydata)
##
## Deviance Residuals:
       Min
                 1Q
                      Median
                                    3Q
                                             Max
## -0.9565
            -0.7090 -0.7090
                                1.4157
                                          1.7344
##
## Coefficients:
##
                  Estimate Std. Error z value Pr(>|z|)
                  -1.2528
                             0.2143 -5.846 5.03e-09 ***
## (Intercept)
## sex.factorMale
                    0.7080
                                        2.235
                                                0.0254 *
                               0.3169
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
      Null deviance: 242.35 on 204 degrees of freedom
## Residual deviance: 237.35 on 203 degrees of freedom
## AIC: 241.35
##
## Number of Fisher Scoring iterations: 4
Now we have created the model - fantastic!
But this doesn't mean a lot to humans reading a paper - or us in
fact.
```

The estimate output of summary(model\_1) represents the logarithm of the odds ratio. The odds ratio would be a lot easier to understand.

Therefore, to sort that out we should exponentiate the output of the model! The exp() function will do this.

### exp(model1\$coefficients)

```
## (Intercept) sex.factorMale
## 0.2857143 2.0300000
```

This gives us an odds ratio of 2.03 for males. That is to say, males are twice as likely to die from melanoma than females.

Now a confidence interval might be handy. As this will be the logarithm of the confidence interval, we should exponentiate it to make it understandable.

### exp(confint(model1))

```
## Waiting for profiling to be done...
## 2.5 % 97.5 %
## (Intercept) 0.1843592 0.4284939
## sex.factorMale 1.0914854 3.7938450
```

The 2.5% is the lower bound and the 97.5% is the upper bound of the 95% confidence interval.

So we can therefore say that being male doubles your chances of dying from melanoma with an Odds Ratio of 2.03 (95% confidence interval of 1.09 to 3.79)

#### 9.7.2 Exercise

Repeat this for all the variables contained within the data, particulary:

```
stage.factor, age, ulcer.factor, thickness and age.factor.
```

Write their odds ratios and 95% confidence intervals down for the next section!

Congratulations on building your first regression model in R!

### 9.8 Finalfit package

We have developed our finalfit package to help with advanced regression modelling. We will introduce it here, but not go into detail.

See www.finalfit.org for more information and updates.

### 9.9 Summarise a list of variables by another variable

We can use the finalfit package to summarise a list of variables by another variable. This is very useful for "Table 1" in many studies.

```
library(finalfit)
dependent = "died_melanoma.factor"
explanatory = c("age", "sex.factor")

table_result = mydata %>%
   summary_factorlist(dependent, explanatory, p = TRUE)
```

label	levels	No	Yes	р
age	Mean (SD)	51.5 (16.1)	55.1 (17.9)	0.189
sex.factor	Female	98 (77.8)	28 (22.2)	0.024
	Male	50 (63.3)	29 (36.7)	

## 9.10 finalfit function for logistic regression

We can then use the finalfit function to run a logistic regression analysis with similar syntax.

```
dependent = "died_melanoma.factor"
explanatory = c("sex.factor")

model2 = mydata %>%
  finalfit(dependent, explanatory)
```

Dependent: died_melanoma.factor		No	Yes	OR (univariabl
sex.factor	Female	98 (66.2)	28 (49.1)	
	Male	50 (33.8)	29 (50.9)	2.03 (1.09-3.79, p=0.028

### 9.11 Adjusting for multiple variables in R

Your first models only included one variable. It's time to scale them up.

Multivariable models take multiple variables and estimates how each variable predicts an event. It adjusts for the effects of each one, so you end up with a model that calculates the adjusted effect estimate (i.e. the odds ratio), upon an outcome.

When you see the term 'adjusted' in scientific papers, this is what it means.

#### 9.11.1 Worked Example

Lets adjust for age (as a continuous variable), sex.factor and stage.factor. Then output them as odds ratios.

```
dependent = "died_melanoma.factor"
explanatory = c("age", "sex.factor", "stage.factor")

model3 = mydata %>%
  finalfit(dependent, explanatory)
```

Dependent: died_melanoma.factor		No	Yes	OR
age	Mean (SD)	51.5 (16.1)	55.1 (17.9)	1.01 (0.99-1.
sex.factor	Female	98 (66.2)	28 (49.1)	
	Male	50 (33.8)	29 (50.9)	2.03 (1.09-3.
stage.factor	Stage I	18 (12.2)	1 (1.8)	
	Stage II	32 (21.6)	5 (8.8)	2.81 (0.41-56.
	Stage III	75 (50.7)	29 (50.9)	6.96 (1.34-128.
	Stage IV	23 (15.5)	22 (38.6)	17.22 (3.13-322.

```
## Waiting for profiling to be done...
## Waiting for profiling to be done...
## Waiting for profiling to be done...
## Warning: Removed 2 rows containing missing values (geom_errorbarh).

died_melanoma.factor: OR (95% CI, p-value)

age 1.01 (0.99-1.03, p=0.534)

sex.factor Female -

1.5720(0.81-3.21, p=0.167)

stage.factorStage I -

25220(0.140-56.96, p=0.363)

75127(0.187-132.38, p=0.061)

145820(0.2154-270.31, p=0.014)

Odds ratio (95% CI, log scale)
```

When we enter age into regression models, the effect estimate is provided in terms of per unit increase. So in this case it's expressed in terms of an odds ratio per year increase (i.e. for every year in age gained odds of death increases by 1.02).

#### 9.11.2 Exercise

Create a regression that includes ulcer.factor.

### 9.12 Advanced: Fitting the best model

Now we have our preliminary model. We could leave it there.

However, when you publish research, you are often asked to supply a measure of how well the model fitted the data.

There are different approaches to model fitting. Come to our course HealthyR-Advanced: Practical Logistic Regression. At this we describe use of the Akaike Information Criterion (AIC) and the C-statistic.

The C-statistic describes discrimination and anything over 0.60 is considered good. The closer to 1.00 the C-statistic is, the better the fit.

The AIC measure model fit with lower values indicating better fit.

These metrics are available here:

```
mydata %>%
  finalfit(dependent, explanatory, metrics=TRUE)
## Waiting for profiling to be done...
## Setting levels: control = 0, case = 1
## Setting direction: controls < cases
## [[1]]
##
   Dependent: died melanoma.factor
                                                  No
                                                          Yes
## 1
                        age Mean (SD) 51.5 (16.1) 55.1 (17.9)
## 2
                   sex.factor
                                Female 98 (66.2)
                                                   28 (49.1)
## 3
                              Male 50 (33.8) 29 (50.9)
## 4
                 stage.factor Stage I 18 (12.2)
                                                      1 (1.8)
## 5
                           Stage II 32 (21.6)
                                                  5 (8.8)
```

```
TI 75 (50.7) 29 (50.9)

V 23 (15.5) 22 (38.6)

OR (multivariable)

1.01 (0.99-1.03, p=0.534)

-

1.62 (0.81-3.21, p=0.167)
```

9 Logistic regression

## [1] "Number in dataframe = 205, Number in model = 205, Missing = 0, AIC = 232.3

Stage III

Stage IV

### 9.12.1 Extra material: Diagnostics plots

OR (univariable)

1.01 (0.99-1.03, p=0.163)

While outwith the objectives of this course, diagnostic plots for glm models can be produced by:

```
plot(model1)
```

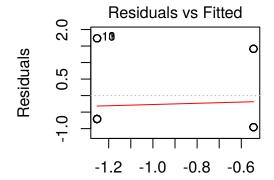
188

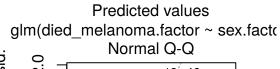
## 6

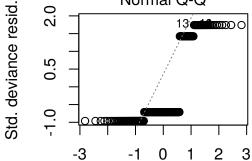
## 7

## 1

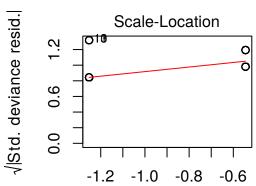
##

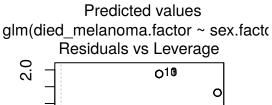


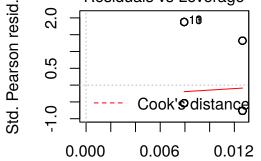




Theoretical Quantiles glm(died\_melanoma.factor ~ sex.factor







Leverage glm(died\_melanoma.factor ~ sex.factor

# Time-to-event data and survival

### 10.1 Data

The boot::melanoma dataset was introduced in chapter 7.

In the previous session, we used logistic regression to investigate death by calculating odds ratios for different factors at a single point in time.

```
library(tidyverse)
library(broom)
library(survival)
library(survminer)
mydata = boot::melanoma
mydata$status %>%
    factor() %>%
    fct_recode("Died" = "1",
                         "Alive" = "2",
                         "Died - other causes" = "3") %>%
    fct_relevel("Alive") -> # move Alive to front (first factor level)
                            # so OR will be relative to that
    mydata$status.factor
mydata$sex %>%
    factor() %>%
    fct_recode("Female" = "0",
                         "Male" = "1") ->
    mydata$sex.factor
```

#### 10.2 Kaplan-Meier survival estimator

The Kaplan-Meier (KM) survival estimator is a non-parametric statistic used to estimate the survival function from time-to-event data.

'Time' is time from event to last known status. This status could be the event, for instance death. Or could be when the patient was last seen, for instance at a clinic. In this circumstance the patient is considered 'censored'.

```
survival_object = Surv(mydata$time, mydata$status.factor == "Died")

# It is often useful to convert days into years
survival_object = Surv(mydata$time/365, mydata$status.factor == "Died")

# Investigate this:
head(survival_object) # + marks censoring in this case "Died of other cause # Or that the follow-up ended and the patient is censored.
```

## [1] 0.02739726+ 0.08219178+ 0.09589041+ 0.27123288+ 0.50684932 0.55890411

### 10.2.1 KM analysis for whole cohort

#### 10.2.2 Model

The survival object is the first step to performing univariable and multivariable survival analyses. A univariable model can then be fitted.

If you want to plot survival stratified by a single grouping variable, you can substitute "survival\_object  $\sim$  1" by "survival\_object  $\sim$  factor"

```
# For all patients
my_survfit = survfit(survival_object ~ 1, data = mydata)
my_survfit # 205 patients, 57 events

## Call: survfit(formula = survival_object ~ 1, data = mydata)
##

##

##

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```

#### 10.2.3 Life table

A life table is the tabular form of a KM plot, which you may be familiar with. It shows survival as a proportion, together with confidence limits. The whole table is shown with, summary(my\_survfit).

```
summary(my_survfit, times = c(0, 1, 2, 3, 4, 5))
## Call: survfit(formula = survival_object ~ 1, data = mydata)
##
```

```
##
   time n.risk n.event survival std.err lower 95% CI upper 95% CI
##
     0
          205
                  0
                       1.000 0.0000
                                           1.000
                                                      1.000
##
     1
          193
                  6
                       0.970 0.0120
                                           0.947
                                                      0.994
     2
##
          183
                  9
                       0.925 0.0187
                                           0.889
                                                      0.962
```

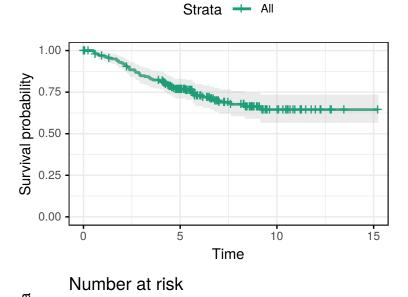
```
##
         167
                 15
                     0.849 0.0255
                                        0.800
                                                   0.900
##
         160
                 6
                     0.818 0.0274
                                        0.766
                                                   0.874
         122
                     0.769 0.0303
                                        0.712
                                                   0.831
##
```

```
# 5 year survival is 77%
# Help is at hand
help(summary.survfit)
```

#### 10.2.4 KM plot

A KM plot can easily be generated using the survminer package.

For more information on how the survminer package draws this plot, or how to modify it: http://www.sthda.com/english/wiki/survminer-r-package-survival-data-analysis-and-visualization and https://github.com/kassambara/survminer



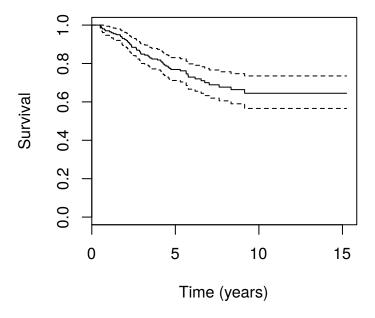
122

```
# Note can also take `ggplot()` options.
my_survplot$plot +
   annotate('text', x = 5, y = 0.25, label='Whole cohort')
```

Time

Here is an alternative plot in base R to compare. Not only does this produce a more basic survival plot, but tailoring the plot can be more difficult to achieve.

Furthermore, appending a life table ('risk.table') alongside the plot can also be difficult, yet this is essential for interpretation.

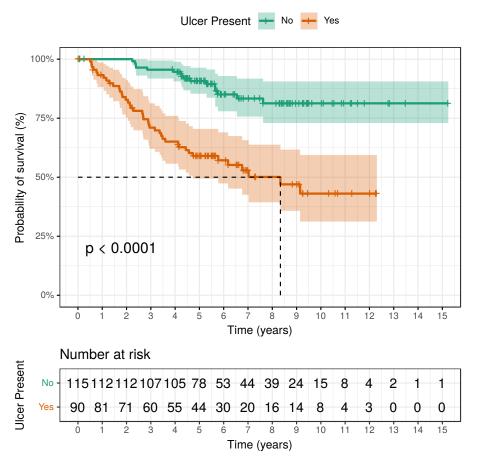


#### 10.2.5 Exercise

Using the above scripts, perform a univariable Kaplan Meier analysis to determine if ulcer.factor influences overall survival. Hint: survival\_object ~ ulcer.factor.

Try modifying the plot produced (see Help for ggsurvplot). For example:

- Add in a medial survival lines: surv.median.line="hv"
- Alter the plot legend: legend.title = "Ulcer Present", legend.labs = c("No", "Yes")
- Change the y-axis to a percentage: ylab = "Probability of survival (%)", surv.scale = "percent"
- Display follow-up up to 10 years, and change the scale to 1 year: xlim = c(0,10), break.time.by = 1)



### 10.2.6 Log-rank test

Two KM survival curves can be compared using the log-rank test. Note survival curves can also be compared using a Wilcoxon test that may be appropriate in some circumstances.

This can easily be performed in library(survival) using the function survdiff().

survdiff(survival\_object ~ ulcer.factor, data = mydata)

```
## Call:
## survdiff(formula = survival_object ~ ulcer.factor, data = mydata)
```

```
##
##
                   N Observed Expected (O-E)^2/E (O-E)^2/V
                                      35.8
## ulcer.factor=Absent 115
                                16
                                              10.9
                                                       29.6
## ulcer.factor=Present 90
                                41
                                      21.2
                                              18.5
                                                       29.6
##
   Chisq= 29.6 on 1 degrees of freedom, p= 5e-08
##
```

Is there a signficiant difference between survival curves?

### 10.3 Cox proportional hazard regression

#### 10.3.1 Model

Multivariable survival analysis can be complex with parametric and semi-parametric methods available. The latter is performed using a Cox proportional hazard regression analysis.

```
# Note several variables are now introduced into the model.
# Variables should be selected carefully based on published methods.
my_hazard = coxph(survival_object~sex.factor+ulcer.factor+age.factor, data=m
summary(my hazard)
## Call:
## coxph(formula = survival_object ~ sex.factor + ulcer.factor +
      age.factor, data = mydata)
##
##
##
    n= 205, number of events= 57
##
##
                 coef exp(coef) se(coef)
                                          z Pr(>|z|)
## sex.factorMale
                    0.48249 1.62011 0.26835 1.798 0.0722 .
## ulcer.factorPresent 1.38972 4.01372 0.29772 4.668 3.04e-06 ***
## age.factor(20,40]
                    ## age.factor(40,60]
                    -0.04513 0.95588 0.61334 -0.074
```

```
## age.factor(60,95]
                       0.17889
                                1.19588 0.62160 0.288 0.7735
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' 1
##
##
                exp(coef) exp(-coef) lower .95 upper .95
## sex.factorMale
                       1.6201
                                 0.6172
                                          0.9575
                                                    2.741
## ulcer.factorPresent
                         4.0137
                                   0.2491
                                            2.2394
                                                      7.194
## age.factor(20,40]
                        0.6661
                                  1.5012
                                            0.1711
                                                     2.593
## age.factor(40,60]
                        0.9559
                                  1.0462
                                            0.2873
                                                     3.180
## age.factor(60,95]
                        1.1959
                                  0.8362
                                            0.3537
                                                     4.044
##
## Concordance= 0.735
                        (se = 0.04)
## Rsquare= 0.153
                     (max possible= 0.937 )
## Likelihood ratio test= 34.08 on 5 df,
                                              p = 2e - 06
## Wald test
                         = 30.19
                                  on 5 df,
                                              p=1e-05
## Score (logrank) test = 35.21
                                  on 5 df,
                                              p=1e-06
library(broom)
tidy(my_hazard)
```

```
## # A tibble: 5 x 7
   term
                estimate std.error statistic p.value conf.low conf.high
                           <dbl>
                                             <dbl>
##
  <chr>
                   <dbl>
                                    <dbl>
                                                     <dbl>
                                                              <dbl>
## 1 sex.factorMale
                     0.482
                               0.268
                                       1.80
                                               7.22e-2 -0.0435
                                                                  1.01
                                               3.04e-6 0.806
## 2 ulcer.factorPr~
                      1.39
                               0.298
                                       4.67
                                                                  1.97
## 3 age.factor(20,~ -0.406
                                       -0.586
                                                5.58e-1 -1.77
                                0.693
                                                                   0.953
## 4 age.factor(40,~ -0.0451
                                       -0.0736
                                                9.41e-1 -1.25
                                                                    1.16
                                0.613
## 5 age.factor(60,~ 0.179
                               0.622
                                       0.288
                                               7.74e-1 -1.04
                                                                  1.40
```

The interpretation of the results of model fitting are beyond the aims of this course. The exponentiated coefficient (exp(coef)) represents the hazard ratio. Therefore, patients with ulcers are 4-times more likely to die at any given time than those without ulcers.

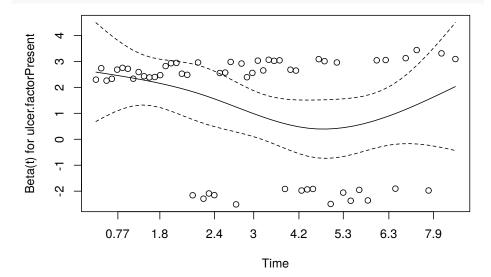
### 10.3.2 Assumptions

The CPH model presumes 'constant hazards'. That means that the risk associated with any given variable (like ulcer status) shouldn't get worse or better over time. This can be checked.

```
ph = cox.zph(my_hazard)
ph
```

```
# GLOBAL shows no overall violation of assumptions.
# Ulcer.status is borderline significant

# Plot Schoenfield residuals to evaluate PH
plot(ph, var=2) # ulcer.status is variable 2
```



10.4 Dates in R 201

```
# help(plot.cox.zph)
```

Hazard decreases a little between 2 and 5 years, but is acceptable.

#### 10.3.3 Exercise

Create a new CPH model, but now include the variable **thickness** as a variable. How would you interpret the output? Is it an independent predictor of overall survival in this model? Are CPH assumptions maintained?

#### 10.4 Dates in R

#### 10.4.1 Converting dates to survival time

In the melanoma example dataset, we already had the time in a convenient format for survial analysis - survival time in days since the operation. This section shows how to convert dates into "days from event". First we will generate a dummy operation date and censoring date based on the melanoma data.

```
library(lubridate)
first_date = ymd("1966-01-01")  # let's create made-up dates for to
last_date = first_date + days(nrow(mydata)-1) # assume tone every day from soperation_date = seq(from = first_date, to = last_date, by = "1 day") # create
mydata$operation_date = operation_date # add the created sequence to melanow
```

Now we will to create a 'censoring' date by adding time from the melanoma dataset to our made up operation date.

Remember the censoring date is either when an event occurred (e.g. death) or the last known alive status of the patient.

```
mydata = mydata %>%
  mutate(censoring_date = operation_date + days(time))

# (Same as doing:):
mydata$censoring_date = mydata$operation_date + days(mydata$time)
```

Now consider if we only had the operation date and censoring date. We want to create the time variable.

```
mydata = mydata %>%
mutate(time_days = censoring_date - operation_date)
```

The Surv() function expects a number (numeric variable), rather than a date object, so we'll convert it:

```
# Surv(mydata$time_days, mydata$status==1) # this doesn't work

mydata %>%
   mutate(time_days_numeric = as.numeric(time_days)) ->
   mydata

survival_object = Surv(mydata$time_days_numeric, mydata$status.factor == "Di
```

### 10.5 Solutions

#### 9.2.2

```
# Fit survival model
my_survfit.solution = survfit(survival_object ~ ulcer.factor, data = mydata)
# Show results
```

10.5 Solutions 203

```
my survfit.solution
summary(my_survfit.solution, times=c(0,1,2,3,4,5))
# Plot results
my_survplot.solution = ggsurvplot(my_survfit.solution,
                         data = mydata,
                         palette = 'Dark2',
                         risk.table = TRUE,
                         ggtheme = theme_bw(),
                         conf.int = TRUE,
                         pval=TRUE,
                         # Add in a medial survival line.
                         surv.median.line="hv",
                         # Alter the plot legend (change the names)
                         legend.title = "Ulcer Present",
                         legend.labs = c("No", "Yes"),
                         # Change the y-axis to a percentage
                         ylab = "Probability of survival (%)",
                         surv.scale = "percent",
                         # Display follow-up up to 10 years, and change the
                         xlab = "Time (years)",
                         # present narrower X axis, but not affect survival
                         xlim = c(0,10),
                         # break X axis in time intervals by 1 year
                         break.time.by = 1)
my_survplot.solution
```

```
# Fit model
my_hazard = coxph(survival_object~sex.factor+ulcer.factor+age.factor+thickne
summary(my_hazard)

# Melanoma thickness has a HR 1.12 (1.04 to 1.21).
# This is interpretted as a 12% increase in the
# risk of death at any time for each 1 mm increase in thickness.

# Check assumptions
ph = cox.zph(my_hazard)
ph
# GLOBAL shows no overall violation of assumptions.
# Plot Schoenfield residuals to evaluate PH
plot(ph, var=6)
```

Part III

Workflow

Notebooks and markdown

Missing data

Encryption

Exporting tables and plots

### RStudio settings, good practise

### 15.1 Script vs Console

Throughout this course, don't copy or type code directly into the Console. We will only be using the Console for viewing output, warnings, and errors. All code should be in a script and executed (=run) using Ctrl+Enter (line or section) or Ctrl+Shift+Enter (whole script). Make sure you are always working in a project (the right-top corner of your RStudio interface should say "HealthyR").

### 15.2 Starting with a blank canvas

In the first session we loaded some data that we then plotted. When we import data, R stores it and displays it in the Environment tab.

It's good practice to restart R before commencing new work. This is to avoid accidentally using the wrong data or functions stored in the environment.

Restarting R only takes a second!

• Restart R (Ctrl+Shift+F10 or select it from Session -> Restart R).

RStudio has a default setting that is no longer considered best practice. You should do this once:

• Go to Tools -> Global Options -> General and set "Save .RData on exit" to Never. This does not mean you can't or shouldn't

save your work in .RData files. But it is best to do it consciously and load exactly what you need to load, rather than letting R always save and load everything for you, as this could also include broken data or objects.

### Bibliography

Xie, Y. (2015). Dynamic Documents with R and knitr. Chapman and Hall/CRC, Boca Raton, Florida, 2nd edition. ISBN 978-1498716963.

Xie, Y. (2018). bookdown: Authoring Books and Technical Documents with R Markdown. R package version 0.7.15.

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