Data Science for Biological, Medical and Health Research: Notes for 432

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Introduction

These Notes provide a series of examples using R to work through issues that are likely to come up in PQHS/CRSP/MPHP 432.

While these Notes share some of the features of a textbook, they are neither comprehensive nor completely original. The main purpose is to give students in 432 a set of common materials on which to draw during the course. In class, we will sometimes:

- reiterate points made in this document,
- amplify what is here,
- simplify the presentation of things done here,
- use new examples to show some of the same techniques,
- refer to issues not mentioned in this document,

but what we don't (always) do is follow these notes very precisely. We assume instead that you will read the materials and try to learn from them, just as you will attend classes and try to learn from them. We welcome feedback of all kinds on this document or anything else. Just email us at 431-help at case dot edu, or submit a pull request. Note that we still use 431-help even though we're now in 432.

What you will mostly find are brief explanations of a key idea or summary, accompanied (most of the time) by R code and a demonstration of the results of applying that code.

Everything you see here is available to you as HTML or PDF. You will also have access to the R Markdown files, which contain the code which generates everything in the document, including all of the R results. We will demonstrate the use of R Markdown (this document is generated with the additional help of an R package called bookdown) and R Studio (the "program" which we use to interface with the R language) in class.

To download the data and R code related to these notes, visit the Data and Code section of the 432 course website.

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R Packages used in these notes

Here, we'll load in the packages used in these notes.

library(tableone)
library(skimr)
library(broom)
library(tidyverse)

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Data used in these notes

Here, we'll load in the data sets used in these notes.

```
fakestroke <- read.csv("data/fakestroke.csv") %>% tbl_df
bloodbrain <- read.csv("data/bloodbrain.csv") %>% tbl_df
smartcle1 <- read.csv("data/smartcle1.csv") %>% tbl_df
```

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Chapter 1

Building Table 1

Many scientific articles involve direct comparison of results from various exposures, perhaps treatments. In 431, we studied numerous methods, including various sorts of hypothesis tests, confidence intervals, and descriptive summaries, which can help us to understand and compare outcomes in such a setting. One common approach is to present what's often called Table 1. Table 1 provides a summary of the characteristics of a sample, or of groups of samples, which is most commonly used to help understand the nature of the data being compared.

1.1 Two examples from the New England Journal of Medicine

1.1.1 A simple Table 1

Table 1 is especially common in the context of clinical research. Consider the excerpt below, from a January 2015 article in the New England Journal of Medicine (Tolaney et al., 2015).

Table 1. Baseline Characteristics of the Patients.*					
Characteristic	Patients (N=406)				
	no. (%)				
Age group					
<50 yr	132 (32.5)				
50–59 yr	137 (33.7)				
60–69 yr	96 (23.6)				
≥70 yr	41 (10.1)				
Sex					
Female	405 (99.8)				
Male	1 (0.2)				
Race†					
White	351 (86.5)				
Black	28 (6.9)				
Asian	11 (2.7)				
Other	16 (3.9)				

This (partial) table reports baseline characteristics on age group, sex and race, describing 406 patients with

HER2-positive¹ invasive breast cancer that began the protocol therapy. Age, sex and race (along with severity of illness) are the most commonly identified characteristics in a Table 1.

In addition to the measures shown in this excerpt, the full Table also includes detailed information on the primary tumor for each patient, including its size, nodal status and histologic grade. Footnotes tell us that the percentages shown are subject to rounding, and may not total 100, and that the race information was self-reported.

1.1.2 A group comparison

A more typical Table 1 involves a group comparison, for example in this excerpt from Roy et al. (2008). This Table 1 describes a multi-center randomized clinical trial comparing two different approaches to caring for patients with heart failure and atrial fibrillation².

Table 1. Baseline Characteristics of the Patients.*							
Variable	Rhythm-Control Group (N = 682)	Rate-Control Group (N = 694)					
Male sex (%)	78	85					
Age (yr)	66±11	67±11					
Body-mass index†	27.8±5.4	28.0±5.1					
Nonwhite race (%)‡	16	13					
NYHA class III or IV (%)							
At baseline	32	31					
During previous 6 mo	76	76					
Predominant cardiac diagnosis (%)∫							
Coronary artery disease	48	48					
Valvular heart disease	5	5					
Nonischemic cardiomyopathy	36	39					
Congenital heart disease	1	1					
Hypertensive heart disease	10	7					

The article provides percentages, means and standard deviations across groups, but note that it does not provide p values for the comparison of baseline characteristics. This is a common feature of NEJM reports on randomized clinical trials, where we anticipate that the two groups will be well matched at baseline. Note that the patients in this study were *randomly* assigned to either the rhythm-control group or to the rate-control group, using blocked randomizations stratified by study center.

1.2 The MR CLEAN trial

Berkhemer et al. (2015) reported on the MR CLEAN trial, involving 500 patients with acute ischemic stroke caused by a proximal intracranial arterial occlusion. The trial was conducted at 16 medical centers in the Netherlands, where 233 were randomly assigned to the intervention (intraarterial treatment plus usual care) and 267 to control (usual care alone.) The primary outcome was the modified Rankin scale score at 90 days; this categorical scale measures functional outcome, with scores ranging from 0 (no symptoms) to 6 (death). The fundamental conclusion of Berkhemer et al. (2015) was that in patients with acute ischemic stroke

¹HER2 = human epidermal growth factor receptor type 2. Over-expression of this occurs in 15-20% of invasive breast cancers, and has been associated with poor outcomes.

²The complete Table 1 appears on pages 2668-2669 of Roy et al. (2008), but I have only reproduced the first page and the footnote in this excerpt.

caused by a proximal intracranial occlusion of the anterior circulation, intraarterial treatment administered within 6 hours after stroke onset was effective and safe.

Here's the Table 1 from Berkhemer et al. (2015).

Characteristic	Intervention (N = 233)	Control (N = 267)
Age — yr		
Median	65.8	65.7
Interquartile range	54.5-76.0	55.5-76.4
Male sex — no. (%)	135 (57.9)	157 (58.8)
NIHSS score†		
Median (interquartile range)	17 (14–21)	18 (14-22)
Range	3-30	4-38
Location of stroke in left hemisphere — no. (%)	116 (49.8)	153 (57.3)
History of ischemic stroke — no. (%)	29 (12.4)	25 (9.4)
Atrial fibrillation — no. (%)	66 (28.3)	69 (25.8)
Diabetes mellitus — no. (%)	34 (14.6)	34 (12.7)
Prestroke modified Rankin scale score — no. (%)‡		
0	190 (81.5)	214 (80.1)
1	21 (9.0)	29 (10.9)
2	12 (5.2)	13 (4.9)
>2	10 (4.3)	11 (4.1)
Systolic blood pressure — mm Hg∫	146±26.0	145±24.4
Treatment with IV alteplase — no. (%)	203 (87.1)	242 (90.6)
Time from stroke onset to start of IV alteplase — min		
Median	85	87
Interquartile range	67-110	65-116
ASPECTS — median (interquartile range)¶	9 (7-10)	9 (8-10)
Intracranial arterial occlusion — no./total no. (%)		
Intracranial ICA	1/233 (0.4)	3/266 (1.1)
ICA with involvement of the M1 middle cerebral artery segment	59/233 (25.3)	75/266 (28.2)
M1 middle cerebral artery segment	154/233 (66.1)	165/266 (62.0)
M2 middle cerebral artery segment	18/233 (7.7)	21/266 (7.9)
A1 or A2 anterior cerebral artery segment	1/233 (0.4)	2/266 (0.8)
Extracranial ICA occlusion — no./total no. (%) **	75/233 (32.2)	70/266 (26.3)
Time from stroke onset to randomization — min††		
Median	204	196
Interquartile range	152-251	149–266
Time from stroke onset to groin puncture — min		
Median	260	NA
Interquartile range	210-313	

The Table was accompanied by the following notes.

- * The intervention group was assigned to intraarterial treatment plus usual care, and the control group was assigned to usual care alone. Plus-minus values are means ±SD. ICA denotes internal carotid artery, IV intravenous, and NA not applicable.
- † Scores on the National Institutes of Health Stroke Scale (NIHSS) range from 0 to 42, with higher scores indicating more severe neurologic deficits. The NIHSS is a 15-item scale, and values for 30 of the 7500 items were missing (0.4%). The highest number of missing items for a single patient was 6.
- Scores on the modified Rankin scale of functional disability range from 0 (no symptoms) to 6 (death). A score of 2 or less indicates functional independence.
- Data on systolic blood pressure at baseline were missing for one patient assigned to the control group.
- The Alberta Stroke Program Early Computed Tomography Score (ASPECTS) is a measure of the extent of stroke. Scores ranges from 0 to 10, with higher scores indicating fewer early ischemic changes. Scores were not available for four patients assigned to the control group: noncontrast computed tomography was not performed in one patient, and three patients had strokes in the territory of the anterior cerebral artery.
- Vessel imaging was not performed in one patient in the control group, so the level of occlusion was not known.
- ** Extracranial ICA occlusions were reported by local investigators.
- †† Data were missing for two patients in the intervention group.

1.3 Simulated fakestroke data

Consider the simulated data, available on the Data and Code page of our course website in the fakestroke.csv file, which I built to let us mirror the Table 1 for MR CLEAN (Berkhemer et al., 2015). The fakestroke.csv file contains the following 18 variables for 500 patients.

studyid	Study ID # (z001 through z500)
trt	Treatment group (Intervention or Control)
age	Age in years
sex	Male or Female
nihss	NIH Stroke Scale Score (can range from 0-42; higher scores
	indicate more severe neurological deficits)
location	Stroke Location - Left or Right Hemisphere
hx.isch	History of Ischemic Stroke (Yes/No)
afib	Atrial Fibrillation $(1 = Yes, 0 = No)$
dm	Diabetes Mellitus $(1 = Yes, 0 = No)$
mrankin	Pre-stroke modified Rankin scale score $(0, 1, 2 \text{ or } > 2)$
	indicating functional disability - complete range is 0 (no
	symptoms) to 6 (death)
sbp	Systolic blood pressure, in mm Hg
iv.altep	Treatment with IV alterplase (Yes/No)
time.iv	Time from stroke onset to start of IV alteplase (minutes) if
	iv.altep=Yes
aspects	Alberta Stroke Program Early Computed Tomography
	score, which measures extent of stroke from 0 - 10; higher
	scores indicate fewer early ischemic changes
ia.occlus	Intracranial arterial occlusion, based on vessel imaging -
	five categories ³
extra.ica	Extracranial ICA occlusion $(1 = Yes, 0 = No)$
time.rand	Time from stroke onset to study randomization, in minutes
time.punc	Time from stroke onset to groin puncture, in minutes (only
	if Intervention)

Here's a quick look at the simulated data in fakestroke.

³The five categories are Intracranial ICA, ICA with involvement of the M1 middle cerebral artery segment, M1 middle cerebral artery segment, M2 middle cerebral artery segment, A1 or A2 anterior cerebral artery segment

fakestroke

```
# A tibble: 500 x 18
   studyid trt
                      age sex
                                nihss location hx.isch afib
                                                                 dm mrankin
           <fct>
   <fct>
                    <dbl> <fct> <int> <fct>
                                                <fct>
                                                        <int> <int> <fct>
 1 z001
           Control 53.0 Male
                                                            0
                                                                  0 2
                                   21 Right
                                                No
 2 z002
           Interve~ 51.0 Male
                                                                  0 0
                                   23 Left
                                                No
                                                            1
                     68.0 Fema~
 3 z003
                                                            0
                                                                  0 0
           Control
                                   11 Right
                                                No
 4 z004
           Control
                     28.0 Male
                                   22 Left
                                                No
                                                            0
                                                                  0 0
                                                            0
 5 z005
           Control
                     91.0 Male
                                   24 Right
                                                No
                                                                  0 0
 6 z006
           Control
                     34.0 Fema~
                                   18 Left
                                                No
                                                                  0 2
 7 z007
                                   25 Right
                                                            0
                                                                  0 0
           Interve~ 75.0 Male
                                                No
 8 z008
           Control
                     89.0 Fema~
                                   18 Right
                                                No
                                                            0
                                                                  0 0
9 z009
           Control
                     75.0 Male
                                   25 Left
                                                No
                                                            1
                                                                  0 2
10 z010
           Interve~ 26.0 Fema~
                                   27 Right
                                                            0
                                                                  0 0
                                                No
# ... with 490 more rows, and 8 more variables: sbp <int>, iv.altep <fct>,
   time.iv <int>, aspects <int>, ia.occlus <fct>, extra.ica <int>,
   time.rand <int>, time.punc <int>
```

1.4 Building Table 1 for fakestroke: Attempt 1

Our goal, then, is to take the data in fakestroke.csv and use it to generate a Table 1 for the study that compares the 233 patients in the Intervention group to the 267 patients in the Control group, on all of the other variables (except study ID #) available. I'll use the tableone package of functions available in R to help me complete this task. We'll make a first attempt, using the CreateTableOne function in the tableone package. To use the function, we'll need to specify:

- the vars or variables we want to place in the rows of our Table 1 (which will include just about everything in the fakestroke data except the studyid code and the trt variable for which we have other plans, and the time.punc which applies only to subjects in the Intervention group.)
 - A useful trick here is to use the dput function, specifically something like dput (names (fakestroke)) can be used to generate a list of all of the variables included in the fakestroke tibble, and then this can be copied and pasted into the vars specification, saving some typing.
- the strata which indicates the levels want to use in the columns of our Table 1 (for us, that's trt)

Stratified by trt Control Intervention test 267 233 age (mean (sd)) 65.38 (16.10) 63.93 (18.09) 0.343 sex = Male (%) 157 (58.8) 135 (57.9) 0.917 nihss (mean (sd)) 18.08 (4.32) 17.97 (5.04) 0.787 117 (50.2) location = Right (%) 114 (42.7) 0.111

hx.isch = Yes (%)	25	(9.4)	29	(12.4)	0.335
afib (mean (sd))	0.26	(0.44)	0.28	(0.45)	0.534
dm (mean (sd))	0.13	(0.33)	0.12	(0.33)	0.923
mrankin (%)					0.922
> 2	11	(4.1)	10	(4.3)	
0	214	(80.1)	190	(81.5)	
1	29	(10.9)	21	(9.0)	
2	13	(4.9)	12	(5.2)	
sbp (mean (sd))	145.00	(24.40)	146.03	(26.00)	0.647
iv.altep = Yes (%)	242	(90.6)	203	(87.1)	0.267
time.iv (mean (sd))	87.96	(26.01)	98.22	(45.48)	0.003
aspects (mean (sd))	8.65	(1.47)	8.35	(1.64)	0.033
ia.occlus (%)					0.795
A1 or A2	2	(0.8)	1	(0.4)	
ICA with M1	75	(28.2)	59	(25.3)	
Intracranial ICA	3	(1.1)	1	(0.4)	
M1	165	(62.0)	154	(66.1)	
M2	21	(7.9)	18	(7.7)	
extra.ica (mean (sd))	0.26	(0.44)	0.32	(0.47)	0.150
time.rand (mean (sd))	213.88	(70.29)	202.51	(57.33)	0.051

1.4.1 Some of this is very useful, and other parts need to be fixed.

- 1. The 1/0 variables (afib, dm, extra.ica) might be better if they were treated as the factors they are, and reported as the Yes/No variables are reported, with counts and percentages rather than with means and standard deviations.
- 2. In some cases, we may prefer to re-order the levels of the categorical (factor) variables, particularly the mrankin variable, but also the ia.occlus variable. It would also be more typical to put the Intervention group to the left and the Control group to the right, so we may need to adjust our trt variable's levels accordingly.
- 3. For each of the quantitative variables (age, nihss, sbp, time.iv, aspects, extra.ica, time.rand and time.punc) we should make a decision whether a summary with mean and standard deviation is appropriate, or whether we should instead summarize with, say, the median and quartiles. A mean and standard deviation really only yields an appropriate summary when the data are least approximately Normally distributed. This will make the p values a bit more reasonable, too. The test column in the first attempt will soon have something useful to tell us.
- 4. If we'd left in the time.punc variable, we'd get some warnings, having to do with the fact that time.punc is only relevant to patients in the Intervention group.

1.4.2 fakestroke Cleaning Up Categorical Variables

Let's specify each of the categorical variables as categorical explicitly. This helps the CreateTableOne function treat them appropriately, and display them with counts and percentages. This includes all of the 1/0, Yes/No and multi-categorical variables.

Then we simply add a factorVars = fs.factorvars call to the CreateTableOne function.

We also want to re-order some of those categorical variables, so that the levels are more useful to us. Specifically, we want to:

- place Intervention before Control in the trt variable,
- reorder the mrankin scale as 0, 1, 2, > 2, and

• rearrange the ia.occlus variable to the order⁴ presented in Berkhemer et al. (2015).

To accomplish this, we'll use the fct_relevel function from the forcats package (loaded with the rest of the core tidyverse packages) to reorder our levels manually.

1.5 fakestroke Table 1: Attempt 2

;	Stratified by trt							
	Interve	ention	Control	L	p	test		
n	233		267					
age (mean (sd))	63.93	(18.09)	65.38	(16.10)	0.343			
sex = Male (%)		(57.9)		(58.8)				
nihss (mean (sd))	17.97	(5.04)	18.08	(4.32)	0.787			
<pre>location = Right (%)</pre>	117	(50.2)	114	(42.7)	0.111			
hx.isch = Yes (%)	29	(12.4)	25	(9.4)	0.335			
afib = 1 (%)	66	(28.3)	69	(25.8)	0.601			
dm = 1 (%)	29	(12.4)	34	(12.7)	1.000			
mrankin (%)					0.922			
0	190	(81.5)	214	(80.1)				
1		(9.0)						
2	12	(5.2)	13	(4.9)				
> 2	10	(4.3)	11	(4.1)				
sbp (mean (sd))	146.03	(26.00)	145.00	(24.40)	0.647			
iv.altep = Yes (%)	203	(87.1)	242	(90.6)	0.267			
time.iv (mean (sd))	98.22	(45.48)	87.96	(26.01)	0.003			
aspects (mean (sd))	8.35	(1.64)	8.65	(1.47)	0.033			
ia.occlus (%)					0.795			
Intracranial ICA	1	(0.4)	3	(1.1)				
ICA with M1	59	(25.3)	75	(28.2)				
M1	154	(66.1)	165	(62.0)				
M2	18	(7.7)	21	(7.9)				
A1 or A2	1	(0.4)	2	(0.8)				
extra.ica = 1 (%)	75	(32.2)	70	(26.3)	0.179			
time.rand (mean (sd))	202.51	(57.33)	213.88	(70.29)	0.051			

The categorical data presentation looks much improved.

⁴We might also have considered reordering the ia.occlus factor by its frequency, using the fct_infreq function

1.5.1 What summaries should we show?

Now, we'll move on to the issue of making a decision about what type of summary to show for the quantitative variables. Since the fakestroke data are just simulated and only match the summary statistics of the original results, not the details, we'll adopt the decisions made by Berkhemer et al. (2015), which were to use medians and interquartile ranges to summarize the distributions of all of the continuous variables except systolic blood pressure.

- Specifying certain quantitative variables as *non-normal* causes R to show them with medians and the 25th and 75th percentiles, rather than means and standard deviations, and also causes those variables to be tested using non-parametric tests, like the Wilcoxon signed rank test, rather than the t test. The test column indicates this with the word nonnorm.
- Specifying *exact* tests for certain categorical variables (we'll try this for the location and mrankin variables) can be done, and these changes will be noted in the test column, as well.

To accomplish this, we need to specify which variables should be treated as non-Normal in the print statement - notice that we don't need to redo the CreateTableOne for this change.

```
Stratified by trt
                          Intervention
                                                   Control
                             233
                                                      267
n
                           65.80 [54.50, 76.00]
                                                    65.70 [55.75, 76.20]
age (median [IQR])
sex = Male (%)
                             135 (57.9)
                                                      157 (58.8)
nihss (median [IQR])
                           17.00 [14.00, 21.00]
                                                    18.00 [14.00, 22.00]
location = Right (%)
                             117 (50.2)
                                                      114 (42.7)
hx.isch = Yes (%)
                              29 (12.4)
                                                       25 (9.4)
afib = 1 (%)
                              66 (28.3)
                                                       69 (25.8)
dm = 1 (\%)
                              29 (12.4)
                                                       34 (12.7)
mrankin (%)
   0
                             190 (81.5)
                                                      214 (80.1)
   1
                              21 (9.0)
                                                       29 (10.9)
   2
                              12 (5.2)
                                                       13 (4.9)
   > 2
                              10 (4.3)
                                                       11 (4.1)
sbp (mean (sd))
                          146.03 (26.00)
                                                   145.00 (24.40)
                                                      242 (90.6)
iv.altep = Yes (%)
                             203 (87.1)
time.iv (median [IQR])
                           85.00 [67.00, 110.00]
                                                    87.00 [65.00, 116.00]
                            9.00 [7.00, 10.00]
aspects (median [IQR])
                                                     9.00 [8.00, 10.00]
ia.occlus (%)
   Intracranial ICA
                               1 (0.4)
                                                        3 (1.1)
   ICA with M1
                              59 (25.3)
                                                       75 (28.2)
   M1
                             154 (66.1)
                                                      165 (62.0)
   M2
                              18 (7.7)
                                                       21 (7.9)
   A1 or A2
                               1 (0.4)
                                                        2 (0.8)
                              75 (32.2)
                                                       70 (26.3)
extra.ica = 1 (\%)
time.rand (median [IQR]) 204.00 [152.00, 249.50] 196.00 [149.00, 266.00]
                         Stratified by trt
                                 test
age (median [IQR])
                           0.579 nonnorm
sex = Male (%)
                           0.917
nihss (median [IQR])
                           0.453 nonnorm
location = Right (%)
                           0.106 exact
```

```
hx.isch = Yes (%)
                           0.335
afib = 1 (%)
                           0.601
dm = 1 (\%)
                           1.000
                           0.917 exact
mrankin (%)
   0
   1
   2
   > 2
sbp (mean (sd))
                           0.647
iv.altep = Yes (%)
                           0.267
time.iv (median [IQR])
                           0.596 nonnorm
aspects (median [IQR])
                           0.075 nonnorm
ia.occlus (%)
                           0.795
   Intracranial ICA
   ICA with M1
   M1
   M2
   A1 or A2
extra.ica = 1 (%)
                           0.179
time.rand (median [IQR])
                           0.251 nonnorm
```

1.6 Obtaining a more detailed Summary

If this was a real data set, we'd want to get a more detailed description of the data to make decisions about things like potentially collapsing categories of a variable, or whether or not a normal distribution was useful for a particular continuous variable, etc. You can do this with the summary command applied to a created Table 1, which shows, among other things, the effect of changing from normal to non-normal p values for continuous variables, and from approximate to "exact" p values for categorical factors.

Note in the summary below that we have some missing values here. Often, we'll present this information within the Table 1, as well.

```
summary(att2)
```

196 149 266 120 360 0.508 -0.93

Summary of continuous variables

```
trt: Intervention
            n miss p.miss mean sd median p25 p75 min max
          233
                 0
                      0.0
                             64 18
                                       66 54
                                               76
                                                    23
                                                        96 -0.34 -0.52
age
nihss
          233
                 0
                      0.0
                             18 5
                                       17
                                          14
                                               21
                                                    10
                                                        28
                                                           0.48 - 0.74
sbp
          233
                 0
                      0.0 146 26
                                      146 129 164
                                                    78 214 -0.07 -0.22
          233
                     12.9
                             98 45
                                                    42 218
time.iv
                30
                                       85
                                           67 110
                                                           1.03 0.08
          233
                 0
                      0.0
                              8
                               2
                                        9
                                            7 10
                                                     5
                                                       10 -0.56 -0.98
aspects
time.rand 233
                 2
                      0.9
                           203 57
                                      204 152 250 100 300
                                                            0.01 - 1.16
trt: Control
            n miss p.miss mean sd median p25 p75 min max
                                                             skew kurt
                      0.0
                             65 16
                                               76
                                                    24
                                                        94 -0.296 -0.28
age
          267
                                           56
                      0.0
                                               22
                                                        25
nihss
          267
                 0
                             18
                               4
                                       18
                                          14
                                                    11
                                                            0.017 - 1.24
          267
                 1
                      0.4
                           145 24
                                      145 128 161
                                                    82 231
                                                            0.156
sbp
                             88 26
                                                    44 130
time.iv
          267
                25
                      9.4
                                       87
                                           65 116
                                                           0.001 -1.32
          267
                      1.5
                              9
                                        9
                                            8
                                              10
                                                     5
                                                       10 -1.071 0.36
aspects
```

0.0 214 70

0

time.rand 267

p-values

pNormal pNonNormal age 0.342813660 0.57856976 nihss 0.787487252 0.45311695 sbp 0.647157646 0.51346132 time.iv 0.003073372 0.59641104 aspects 0.032662901 0.07464683 time.rand 0.050803672 0.25134327

Standardize mean differences

1 vs 2
age 0.08478764
nihss 0.02405390
sbp 0.04100833
time.iv 0.27691223
aspects 0.19210662
time.rand 0.17720957

Summary of categorical variables

trt: Interv	ent:	ion					
var	n	miss	p.miss	level	freq	percent	cum.percent
sex	233	0	0.0	Female	98	42.1	42.1
				Male	135	57.9	100.0
location	233	0	0.0	Left	116	49.8	49.8
				Right	117	50.2	100.0
hx.isch	233	0	0.0	No	204	87.6	87.6
				Yes	29	12.4	100.0
afib	233	0	0.0	0	167	71.7	71.7
				1	66	28.3	100.0
dm	233	0	0.0	0	204	87.6	87.6
				1	29	12.4	100.0
mrankin	233	0	0.0	0	190	81.5	81.5
				1	21	9.0	90.6
				2	12	5.2	95.7
				> 2	10	4.3	100.0
iv.altep	233	0	0.0	No	30	12.9	12.9
•				Yes	203	87.1	100.0
ia.occlus	233	0	0.0	Intracranial ICA	1	0.4	0.4
				ICA with M1	59	25.3	25.8
				M1	154		91.8
				M2	18		99.6
				A1 or A2	1	0.4	100.0

extra.ica	233	0	0.0	0	158 75	67.8 32.2	67.8 100.0
trt: Contro	 วไ						
var		miss	p.miss	level	freq	percent	cum.percent
sex	267	0	-	Female	-	41.2	41.2
				Male	157	58.8	100.0
location	267	0	0.0	Left	153	57.3	57.3
				Right	114	42.7	100.0
hx.isch	267	0	0.0	No	242	90.6	90.6
				Yes	25	9.4	100.0
afib	267	0	0.0	0	198	74.2	74.2
				1	69	25.8	100.0
dm	267	0	0.0	0	233	87.3	87.3
				1	34	12.7	100.0
mrankin	267	0	0.0	0	214	80.1	80.1
				1	29	10.9	91.0
				2	13	4.9	95.9
				> 2	11	4.1	100.0
iv.altep	267	0	0.0	No	25	9.4	9.4
•				Yes	242	90.6	100.0
ia.occlus	267	1	0.4	Intracranial ICA	3	1.1	1.1
				ICA with M1	75	28.2	29.3
				M1	165	62.0	91.4
				M2	21	7.9	99.2
				A1 or A2	2	0.8	100.0
extra.ica	267	1	0.4	0	196	73.7	73.7
				1	70	26.3	100.0
p-values	μAq	prox	рЕха	act			

pApprox pExact sex 0.9171387 0.8561188 location 0.1113553 0.1056020 hx.isch 0.3352617 0.3124683 afib 0.6009691 0.5460206 dm 1.0000000 mrankin 0.9224798 0.9173657 iv.altep 0.2674968 0.2518374 ia.occlus 0.7945580 0.8189090 extra.ica 0.1793385 0.1667574

 ${\tt Standardize\ mean\ differences}$

1 vs 2

sex 0.017479025

```
location 0.151168444
hx.isch 0.099032275
afib 0.055906317
dm 0.008673478
mrankin 0.062543164
iv.altep 0.111897009
ia.occlus 0.117394890
extra.ica 0.129370206
```

Again, I have simulated the data to mirror the results in the published Table 1 for this study. In no way have I captured the full range of the real data, or any of the relationships in that data, so it's more important here to see what's available in the analysis, rather than to interpret it closely in the clinical context.

1.7 Exporting the Completed Table 1 from R to Excel or Word

Once you've built the table and are generally satisfied with it, you'll probably want to be able to drop it into Excel or Word for final cleanup.

1.7.1 Approach A: Save and open in Excel

One option is to save the Table 1 to a .csv file, which you can then open directly in Excel. This is the approach I generally use. Note the addition of some quote, noSpaces and printToggle selections here.

When I then open the fs-table1.csv file in Excel, it looks like this:

4	Α	В	С	D	E
1		Intervention	Control	p	test
2	n	233	267		
3	age (median [IQR])	65.80 [54.50, 76.00]	65.70 [55.75, 76.20]	0.579	nonnorm
4	sex = Male (%)	135 (57.9)	157 (58.8)	0.917	
5	nihss (median [IQR])	17.00 [14.00, 21.00]	18.00 [14.00, 22.00]	0.453	nonnorm
6	location = Right (%)	117 (50.2)	114 (42.7)	0.111	
7	hx.isch = Yes (%)	29 (12.4)	25 (9.4)	0.335	
8	afib = 1 (%)	66 (28.3)	69 (25.8)	0.601	
9	dm = 1 (%)	29 (12.4)	34 (12.7)	1	
10	mrankin (%)			0.922	
11	0	190 (81.5)	214 (80.1)		
12	1	21 (9.0)	29 (10.9)		
13	2	12 (5.2)	13 (4.9)		
14	>2	10 (4.3)	11 (4.1)		
15	sbp (mean (sd))	146.03 (26.00)	145.00 (24.40)	0.647	
16	iv.altep = Yes (%)	203 (87.1)	242 (90.6)	0.267	
17	time.iv (median [IQR])	85.00 [67.00, 110.00]	87.00 [65.00, 116.00]	0.596	nonnorm
18	aspects (median [IQR])	9.00 [7.00, 10.00]	9.00 [8.00, 10.00]	0.075	nonnorm
19	ia.occlus (%)			0.795	
20	Intracranial ICA	1 (0.4)	3 (1.1)		
21	ICA with M1	59 (25.3)	75 (28.2)		
22	M1	154 (66.1)	165 (62.0)		
23	M2	18 (7.7)	21 (7.9)		
24	A1 or A2	1 (0.4)	2 (0.8)		
25	extra.ica = 1 (%)	75 (32.2)	70 (26.3)	0.179	
26	time.rand (median [IQR])	204.00 [152.00, 249.50]	196.00 [149.00, 266.00]	0.251	nonnorm
27	time.punc (median [IQR])	260.00 [212.00, 313.00]	NA [NA, NA]	NA	nonnorm
28					

And from here, I can either drop it directly into Word, or present it as is, or start tweaking it to meet formatting needs.

1.7.2 Approach B: Produce the Table so you can cut and paste it

This will look like a mess by itself, but if you:

- 1. copy and paste that mess into Excel
- 2. select Text to Columns from the Data menu
- 3. select Delimited, then Space and select Treat consecutive delimiters as one

you should get something usable again.

Or, in Word,

1. insert the text

- 2. select the text with your mouse
- 3. select Insert ... Table ... Convert Text to Table
- 4. place a quotation mark in the "Other" area under Separate text at ...

After dropping blank columns, the result looks pretty good.

1.8 A Controlled Biological Experiment - The Blood-Brain Barrier

My source for the data and the following explanatory paragraph is page 307 from Ramsey and Schafer (2002). The original data come from Barnett et al. (1995).

The human brain (and that of rats, coincidentally) is protected from the bacteria and toxins that course through the bloodstream by something called the blood-brain barrier. After a method of disrupting the barrier was developed, researchers tested this new mechanism, as follows. A series of 34 rats were inoculated with human lung cancer cells to induce brain tumors. After 9-11 days they were infused with either the barrier disruption (BD) solution or, as a control, a normal saline (NS) solution. Fifteen minutes later, the rats received a standard dose of a particular therapeutic antibody (L6-F(ab')2. The key measure of the effectiveness of transmission across the brain-blood barrier is the ratio of the antibody concentration in the brain tumor to the antibody concentration in normal tissue outside the brain. The rats were then sacrificed, and the amounts of antibody in the brain tumor and in normal tissue from the liver were measured. The study's primary objective is to determine whether the antibody concentration in the tumor increased when the blood-barrier disruption infusion was given, and if so, by how much?

1.9 The bloodbrain.csv file

Consider the data, available on the Data and Code page of our course website in the bloodbrain.csv file, which includes the following variables:

Variable	Description
case	identification number for the rat (1 - 34)
brain	an outcome: Brain tumor antibody count (per gram)
liver	an outcome: Liver antibody count (per gram)
tlratio	an outcome: tumor / liver concentration ratio
solution	the treatment: BD (barrier disruption) or NS (normal saline)
sactime	a design variable: Sacrifice time (hours; either 0.5, 3, 24 or 72)
postin	covariate: Days post-inoculation of lung cancer cells (9, 10 or
	11)
sex	covariate: M or F
wt.init	covariate: Initial weight (grams)
wt.loss	covariate: Weight loss (grams)
wt.tumor	covariate: Tumor weight (10 ⁻⁴ grams)

And here's what the data look like in R.

bloodbrain

```
# A tibble: 34 x 11

case brain liver tlratio solution sactime postin sex wt.init
<int> <int> <int> <int> <fct> <fct> <dbl> <fct> <dbl> <int> <fct> <int> 239
```

```
2 44286 1602171 0.0276 BD
                                           0.500
                                                     10 F
                                                                  225
 3
      3 102926 1601936 0.0642 BD
                                           0.500
                                                     10 F
                                                                  224
                                                     10 F
 4
      4 25927 1776411 0.0146 BD
                                           0.500
                                                                  184
 5
      5 42643 1351184 0.0316 BD
                                           0.500
                                                     10 F
                                                                  250
 6
      6
         31342 1790863 0.0175 NS
                                           0.500
                                                     10 F
                                                                  196
7
                                           0.500
      7 22815 1633386 0.0140 NS
                                                     10 F
                                                                  200
                                           0.500
8
        16629 1618757 0.0103 NS
                                                     10 F
                                                                  273
9
      9
         22315 1567602 0.0142 NS
                                           0.500
                                                     10 F
                                                                  216
     10
         77961 1060057 0.0735 BD
                                           3.00
                                                     10 F
                                                                  267
# ... with 24 more rows, and 2 more variables: wt.loss <dbl>, wt.tumor
    <int>
```

1.10 A Table 1 for bloodbrain

Barnett et al. (1995) did not provide a Table 1 for these data, so let's build one to compare the two solutions (BD vs. NS) on the covariates and outcomes, plus the natural logarithm of the tumor/liver concentration ratio (tlratio). We'll opt to treat the sacrifice time (sactime) and the days post-inoculation of lung cancer cells (postin) as categorical rather than quantitative variables.

Summary of continuous variables

```
solution: BD
                                              p25
         n miss p.miss
                         mean
                                  sd median
                                                    p75
                                                            min
                                                                  max
wt.init
        17
              0
                          243 3e+01
                                     2e+02
                                            2e+02 3e+02
                                                         2e+02 3e+02
wt.loss 17
              0
                     0
                             3 5e+00
                                     4e+00
                                            1e+00 6e+00 -5e+00 1e+01
wt.tumor 17
              0
                     0
                          157 8e+01
                                     2e+02
                                            1e+02 2e+02
                                                         2e+01 4e+02
              0
                     0 56043 3e+04 5e+04 4e+04 8e+04
                                                         6e+03 1e+05
brain
         17
                     0 672577 7e+05 6e+05 2e+04 1e+06
                                                         2e+03 2e+06
liver
         17
              0
                            2 3e+00 1e-01 6e-02 3e+00 1e-02 9e+00
tlratio
        17
              0
                     0
logTL
               0
                           -1 2e+00 -2e+00 -3e+00 1e+00 -4e+00 2e+00
         17
         skew kurt
wt.init -0.39 0.7
wt.loss -0.10 0.2
```

```
wt.tumor 0.53 1.0
brain 0.29 -0.6
                0.35 - 1.7
liver
tlratio 1.58 1.7
logTL 0.08 -1.7
 -----
solution: NS
                   n miss p.miss mean sd median p25 p75 min max
wt.init 17 0 0 240 3e+01 2e+02 2e+02 3e+02 2e+02 3e+02

      wt.lnit
      17
      0
      0
      240 Se+01
      2e+02
      2e+02
      3e+02
      2e+02
      3e+02
      2e+02
      3e+02
      2e+02
      3e+02
      3e+04
      1e+03
      3e+0
                   skew kurt
wt.init 0.33 -0.48
wt.loss -0.09 0.08
wt.tumor 0.63 0.77
brain 0.30 -0.35
liver
                 0.40 - 1.56
tlratio 2.27 4.84
logTL
                0.27 - 1.61
p-values
                          pNormal pNonNormal
wt.init 0.807308940 0.641940278
wt.loss 0.683756156 0.876749808
wt.tumor 0.151510151 0.190482094
brain 0.001027678 0.002579901
liver
                  0.974853609 0.904045603
tlratio 0.320501715 0.221425879
logTL
                 0.351633525 0.221425879
Standardize mean differences
                          1 vs 2
wt.init 0.08435244
wt.loss 0.14099823
wt.tumor 0.50397184
brain 1.23884159
liver 0.01089667
tlratio 0.34611465
logTL 0.32420504
          ### Summary of categorical variables ###
solution: BD
```

```
var n miss p.miss level freq percent cum.percent sactime 17 0 0.0 0.5 5 29.4 29.4 3 4 23.5 52.9 24 4 23.5 76.5 72 4 23.5 100.0
```

postin	17	0	0.0	9	1	5.9	5.9	
				10	14	82.4	88.2	
				11	2	11.8	100.0	
sex	17	0	0.0	F	13	76.5	76.5	
				M	4	23.5	100.0	
solution: NS								
var	n	miss	p.miss	level	freq	percent	cum.percent	
sactime	17	0	0.0	0.5	4	23.5	23.5	
				3	5	29.4	52.9	
				24	4	23.5	76.5	
				72	4	23.5	100.0	
postin	17	0	0.0	9	2	11.8	11.8	
				10	13	76.5	88.2	
				11	2	11.8	100.0	
sex	17	0	0.0	F	13	76.5	76.5	
				М	4	23.5	100.0	

p-values

pApprox pExact sactime 0.9739246 1 1 postin 0.8309504 1 sex 1.0000000 1

Standardize mean differences

1 vs 2 sactime 0.1622214 postin 0.2098877 sex 0.0000000

1.10.1 Generate final Table 1 for bloodbrain

I'll choose to treat tlratio as non-Normal, but otherwise, use t tests.

print(bb.att1, nonnormal = c("tlratio", "logTL"))

	Stratified	by solution			
	BD		NS		
n	17			17	
<pre>sactime (%)</pre>					
0.5	5	(29.4)		4	(23.5)
3	4	(23.5)		5	(29.4)
24	4	(23.5)		4	(23.5)
72	4	(23.5)		4	(23.5)
postin (%)					
9	1	(5.9)		2	(11.8)
10	14	(82.4)		13	(76.5)
11	2	(11.8)		2	(11.8)

```
4 (23.5)
sex = M (\%)
                                4 (23.5)
wt.init (mean (sd))
                          242.82 (27.23)
                                                   240.47 (28.54)
wt.loss (mean (sd))
                            3.34 (4.68)
                                                     3.94 (3.88)
wt.tumor (mean (sd))
                           157.29 (84.00)
                                                   208.53 (116.68)
brain (mean (sd))
                        56043.41 (33675.40)
                                                 23887.18 (14610.53)
liver (mean (sd))
                       672577.35 (694479.58)
                                                664975.47 (700773.13)
tlratio (median [IQR])
                            0.12 [0.06, 2.84]
                                                     0.05 [0.03, 0.94]
                            -2.10 [-2.74, 1.04]
                                                    -2.95 [-3.41, -0.07]
logTL (median [IQR])
                       Stratified by solution
                               test
                       р
n
                        0.974
sactime (%)
   0.5
   3
   24
   72
postin (%)
                         0.831
   10
   11
sex = M (\%)
                         1.000
wt.init (mean (sd))
                         0.807
wt.loss (mean (sd))
                         0.684
wt.tumor (mean (sd))
                         0.152
brain (mean (sd))
                         0.001
liver (mean (sd))
                         0.975
tlratio (median [IQR]) 0.221 nonnorm
logTL (median [IQR])
                        0.221 nonnorm
```

Or, we can get an Excel-readable version, using

A	A	В	С	D	E
1		BD	NS	р	test
2	n	17	17		
3	sex = M (%)	4 (23.5)	4 (23.5)	1	
4	sactime (%)			0.974	
5	0.5	5 (29.4)	4 (23.5)		
6	3	4 (23.5)	5 (29.4)		
7	24	4 (23.5)	4 (23.5)		
8	72	4 (23.5)	4 (23.5)		
9	postin (%)			0.831	
10	9	1 (5.9)	2 (11.8)		
11	10	14 (82.4)	13 (76.5)		
12	11	2 (11.8)	2 (11.8)		
13	wt.init (mean (sd))	242.82 (27.23)	240.47 (28.54)	0.807	
14	wt.loss (mean (sd))	3.34 (4.68)	3.94 (3.88)	0.684	
15	wt.tumor (mean (sd))	157.29 (84.00)	208.53 (116.68)	0.152	
16	brain (mean (sd))	56043.41 (33675.40)	23887.18 (14610.53)	0.001	
17	liver (mean (sd))	672577.35 (694479.58)	664975.47 (700773.13)	0.975	
18	tlratio (median [IQR])	0.12 [0.06, 2.84]	0.05 [0.03, 0.94]	0.221	nonnorm
19	logTL (median [IQR])	-2.10 [-2.74, 1.04]	-2.95 [-3.41, -0.07]	0.221	nonnorm
20					

One thing I would definitely clean up here, in practice, is to change the presentation of the p value for sex from 1 to > 0.99, or just omit it altogether. I'd also drop the computer-ese where possible, add units for the measures, round a lot, identify the outcomes carefully, and use notes to indicate deviations from the main approach.

1.10.2 A More Finished Version (after Cleanup in Word)

Table 1. Comparing Rats Receiving BD to those Receiving NS on Available Covariates and Design Variables, and Key Outcomes

	Barrier Disruption	Normal Saline	
	(BD: treatment)	(NS: control)	р
# of Rats	17	17	
Sex = Male	4 (23.5)	4 (23.5)	-
Sacrifice Time (hours)			0.97
0.5	5 (29.4)	4 (23.5)	
3	4 (23.5)	5 (29.4)	
24	4 (23.5)	4 (23.5)	
72	4 (23.5)	4 (23.5)	
Days post-inoculation of			0.83
lung cancer cells			0.03
9	1 (5.9)	2 (11.8)	
10	14 (82.4)	13 (76.5)	
11	2 (11.8)	2 (11.8)	
Initial Weight (g)	243 (27)	240 (29)	0.81
Weight Loss (g)	3.3 (4.7)	3.9 (3.9)	0.68
Tumor Weight (10 ⁻⁴ g)	157.3 (84.0)	208.5 (116.7)	0.15
Key Outcomes: mean (sd) unless otherw	ise indicated		
Brain Tumor Antibody Count (per g)	56,043 (33,675)	23,887 (14,611)	0.001
Liver Antibody Count (per g)	672,577 (694,480)	664,975 (700,773)	0.98
Tumor/Liver Ratio	0.12	0.05	0.22
(median [Q25, Q75])	[0.06, 2.84]	[0.03, 0.94]	0.22
Natural Log of Tumor/Liver Ratio	-2.10	-2.95	0.22
(median [Q25, Q75])	[-2.74, 1.04]	[-3.41, -0.07]	0.22

Table 1 Notes:

- Categorical variables are summarized with counts, percentages and p values based on approximate chi-square tests.
- Continuous variables, unless otherwise indicated, are summarized with means, standard deviations and p values based on t tests.
- The Tumor / Liver ratio and its natural logarithm are summarized with the median and quartiles and a p value from a non-parametric (Wilcoxon signed rank) test.

Chapter 2

Linear Regression on a small SMART data set

2.1 BRFSS and SMART

The Centers for Disease Control analyzes Behavioral Risk Factor Surveillance System (BRFSS) survey data for specific metropolitan and micropolitan statistical areas (MMSAs) in a program called the Selected Metropolitan/Micropolitan Area Risk Trends of BRFSS (SMART BRFSS.)

In this work, we will focus on data from the 2016 SMART, and in particular on data from the Cleveland-Elyria, OH, Metropolitan Statistical Area. The purpose of this survey is to provide localized health information that can help public health practitioners identify local emerging health problems, plan and evaluate local responses, and efficiently allocate resources to specific needs.

2.1.1 Key resources

- the full data are available in the form of the 2016 SMART BRFSS MMSA Data, found in a zipped SAS Transport Format file. The data were released in August 2017.
- the MMSA Variable Layout PDF which simply lists the variables included in the data file
- the Calculated Variables PDF which describes the risk factors by data variable names there is also an online summary matrix of these calculated variables, as well.
- the lengthy 2016 Survey Questions PDF which lists all questions asked as part of the BRFSS in 2016
- the enormous Codebook for the 2016 BRFSS Survey PDF which identifies the variables by name for

Later this term, we'll use all of those resources to help construct a more complete data set than we'll study today. I'll also demonstrate how I built the smartcle1 data set that we'll use in this Chapter.

2.2 The smartcle1 data: Cookbook

The smartcle1.csv data file available on the Data and Code page of our website describes information on 11 variables for 1036 respondents to the BRFSS 2016, who live in the Cleveland-Elyria, OH, Metropolitan Statistical Area. The variables in the smartcle1.csv file are listed below, along with (in some cases) the BRFSS items that generate these responses.

Variable	Description
SEQNO	respondent identification number (all begin with 2016)

Variable	Description
physhealth	Now thinking about your physical health, which includes physical illness and injury, for how many days during the past 30 days was your physical health not good?
menthealth	Now thinking about your mental health, which includes stress, depression, and problems with emotions, for how many days during the past 30 days was your mental health not good?
poorhealth	During the past 30 days, for about how many days did poor physical or mental health keep you from doing your usual activities, such as self-care, work, or recreation?
genhealth	Would you say that in general, your health is (five categories: Excellent, Very Good, Good, Fair or Poor)
bmi	Body mass index, in kg/m ²
female	Sex, $1 = \text{female}$, $0 = \text{male}$
internet30	Have you used the internet in the past 30 days? $(1 = yes, 0 = no)$
exerany	During the past month, other than your regular job, did you participate in any physical activities or exercises such as running, calisthenics, golf, gardening, or walking for exercise? $(1 = yes, 0 = no)$
sleephrs	On average, how many hours of sleep do you get in a 24-hour period?
alcdays	How many days during the past 30 days did you have at least one drink of any alcoholic beverage such as beer, wine, a malt beverage or liquor?

str(smartcle1)

```
Classes 'tbl_df', 'tbl' and 'data.frame': 1036 obs. of 11 variables:
           : num 2.02e+09 2.02e+09 2.02e+09 2.02e+09 2.02e+09 ...
$ physhealth: int  0 0 1 0 5 4 2 2 0 0 ...
$ menthealth: int  0 0 5 0 0 18 0 3 0 0 ...
 $ poorhealth: int NA NA O NA O 6 O O NA NA ...
 $ genhealth : Factor w/ 5 levels "1_Excellent",..: 2 1 2 3 1 2 3 3 2 3 ...
 $ bmi
            : num 26.7 23.7 26.9 21.7 24.1 ...
$ female
            : int 1001001100...
$ internet30: int 1 1 1 1 1 1 1 1 1 1 ...
          : int 1 1 0 1 1 1 1 1 1 0 ...
 $ exerany
 $ sleephrs : int 6 6 8 9 7 5 9 7 7 7 ...
            : int 1 4 4 3 2 28 4 2 4 25 ...
 $ alcdays
```

2.2.1 Omitting Missing Observations: Complete-Case Analyses

For the purpose of fitting our first few models, we will eliminate the missingness problem, and look only at the *complete cases* in our smartcle1 data.

To inspect the missingness in our data, we might consider using the skim function from the skimr package. We'll exclude the respondent identifier code (SEQNO) from this summary as uninteresting. In this case, we'll use the skim_with command to set some options that will make this book work a bit more effectively, by eliminating the spark graphs that we'll demonstrate in class.

```
skim_with(numeric = list(hist = NULL), ts = list(line_graph = NULL))
smartcle1 %>% skim(-SEQNO)
```

Skim summary statistics

n obs: 1036
n variables: 11

```
Variable type: factor
     variable missing complete
                                                                                 n n unique
  genhealth
                                                            1033 1036
                                            3
                                                                            top_counts ordered
  2 V: 350, 3 G: 344, 1 E: 173, 4 F: 122
                                                                                                              FALSE
Variable type: integer
       variable missing complete
                                                                                                            sd p0 p25 median p75 p100
                                                                                    n mean
          alcdays
                                            46
                                                                 990 1036 4.65 8.05 0
                                                                                                                                 0
          exerany
                                               3
                                                               1033 1036 0.76 0.43
                                                                                                                                 1
                                                                                                                                                    1
                                                                                                                                                                           1
                                                                                                                                                               1
                                               0
                                                               1036 1036 0.6 0.49
             female
                                                                                                                    0
                                                                                                                                 0
                                                                                                                                                    1
                                                                                                                                                               1
                                                                                                                                                                            1
   internet30
                                               6
                                                              1030 1036 0.81 0.39 0
                                                                                                                                                                           1
                                                                                                                                 1
                                                                                                                                                    1
                                                                                                                                                               1
                                                                                                                                                               2
  menthealth
                                           11
                                                              1025 1036 2.72 6.82 0
                                                                                                                                 0
                                                                                                                                                                         30
                                          17
                                                               1019 1036 3.97 8.67 0
                                                                                                                                                    0
                                                                                                                                                               2
                                                                                                                                                                         30
  physhealth
                                                                                                                                 0
  poorhealth
                                          543
                                                                493 1036 4.07 8.09 0
                                                                                                                                 0
                                                                                                                                                    0
                                                                                                                                                               3
                                                                                                                                                                         30
                                                               1028 1036 7.02 1.53 1
                                                                                                                                                                         20
                                               8
       sleephrs
            hist
   <U+2587><U+2582><U+2581><U+2581><U+2581><U+2581><U+2581>
   <U+2582><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581>
   <U+2585><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2587>
   <U+2582><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+258
   <U+2587><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581>
   <U+2587><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581><U+2581>
   <U+2587><U+2581><U+2581><U+2581><U+2581><U+2581>
   <U+2581><U+2581><U+2587><U+2581><U+2581><U+2581><U+2581>
Variable type: numeric
                                                                               n mean
  variable missing complete
                                                                                                          sd
                                                                                                                         p0 p25 median
                                                                                                                                                                      p75 p100
               bmi
                                                             952 1036 27.89 6.47 12.71 23.7 26.68 30.53 66.06
```

Now, we'll create a new tibble called smartcle2 which contains every variable except poorhealth, and which includes all respondents with complete data on the variables (other than poorhealth). We'll store those observations with complete data in the smartcle2 tibble.

```
smartcle2 <- smartcle1 %>%
    select(-poorhealth) %>%
    filter(complete.cases(.))
smartcle2
```

A tibble: 896 x 10 SEQNO physhealth menthealth genhealth bmi female internet30 exerany <dbl> <int> <int> <fct> <dbl> <int> <int> <int> 1 2.02e9 0 0 2_VeryGo~ 26.7 1 1 1 2 2.02e9 0 0 1_Excell~ 23.7 0 1 1 3 2.02e9 5 2_VeryGo~ 0 1 26.9 0 1 4 2.02e9 0 0 3_Good 21.7 1 1 1 5 2.02e9 5 0 1_Excell~ 24.1 0 1 1 6 2.02e9 4 18 2_VeryGo~ 27.6 0 1 1 7 2.02e9 2 0 3_Good 25.7 1 1 1 8 2.02e9 2 3 3_Good 28.5 1 1 1 9 2.02e9 0 0 2_VeryGo~ 28.6 0 1 1 10 2.02e9 0 23.1 0 0 0 3 Good # ... with 886 more rows, and 2 more variables: sleephrs <int>, alcdays

<int>

Note that there are only 896 respondents with **complete** data on the 10 variables (excluding poorhealth) in the smartcle2 tibble, as compared to our original smartcle1 data which described 1036 respondents and 11 variables, but with lots of missing data.

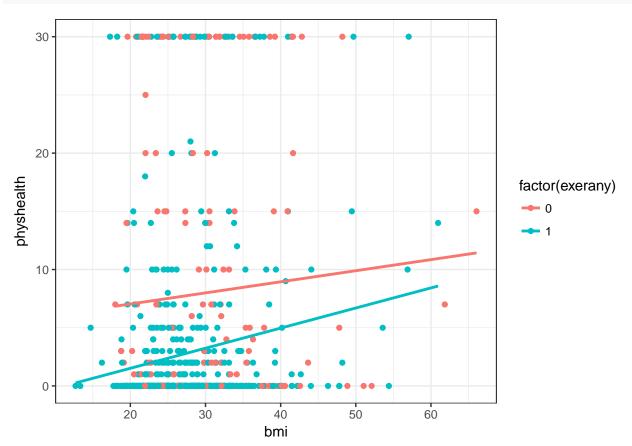
2.3 Model A: Predicting physhealth

We'll begin by investigating the problem of predicting physhealth, at first with just two predictor variables: exerany and bmi, in our new smartcle2 data set.

2.3.1 Graphing The Data

We'll start by building an exploratory figure to show the relationship between bmi and physhealth within each of the two exerany groups.

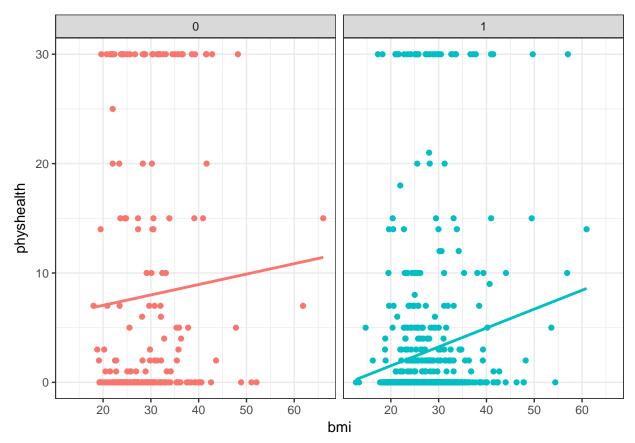
```
ggplot(smartcle2, aes(x = bmi, y = physhealth, group = exerany, color = factor(exerany))) +
    geom_point() +
    geom_smooth(method = "lm", se = FALSE) +
    theme_bw()
```



The figure *could* be improved by separating the two groups into facets.

```
ggplot(smartcle2, aes(x = bmi, y = physhealth, group = exerany, color = factor(exerany))) +
    geom_point() +
    geom_smooth(method = "lm", se = FALSE) +
```

```
theme_bw() +
guides(color = FALSE) +
facet_wrap(~ exerany)
```



Now, what can we learn from these plots?

- 1. Does physhealth look like a good candidate for a linear model?
- 2. Does there seem to be a meaningful difference in the slopes of the two fitted lines?
- 3. In what BMI range can we make a reasonable prediction of physhealth?

2.3.2 Building Model A

First, we'll fit a simple model describing the main effects but not the interaction of exerany and bmi, without doing any of the exploratory ground work we should do in advance.

```
modA <- lm(physhealth ~ exerany * bmi, data = smartcle2)</pre>
glance(modA)
   r.squared adj.r.squared
                              sigma statistic
                                                   p.value df
                                                                 logLik
1 0.07607716
                 0.0729698 8.318521
                                     24.48286 3.14356e-15 4 -3167.527
       AIC
                BIC deviance df.residual
1 6345.053 6369.043 61724.43
tidy(modA)
                 estimate std.error statistic
                                                   p.value
         term
1 (Intercept) 5.13460484 2.3570652 2.1783890 0.02963831
      exerany -7.06992980 2.8044237 -2.5209921 0.01187578
```

```
bmi 0.09527214 0.0772901 1.2326565 0.21802872
4 exerany:bmi 0.07732883 0.0944822 0.8184487 0.41331959
```

summary(modA)

Call:

lm(formula = physhealth ~ exerany * bmi, data = smartcle2)

Residuals:

Min 1Q Median 3Q -10.0992 -3.4231 -2.3780 -0.9199 28.9493

Coefficients:

Estimate Std. Error t value Pr(>|t|) (Intercept) 5.13460 2.35707 2.178 0.0296 * exerany -7.06993 2.80442 -2.521 0.0119 * 0.09527 0.07729 1.233 0.2180 exerany:bmi 0.07733 0.09448 0.818 0.4133

Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1

Residual standard error: 8.319 on 892 degrees of freedom Multiple R-squared: 0.07608, Adjusted R-squared: 0.07297 F-statistic: 24.48 on 3 and 892 DF, p-value: 3.144e-15

Bibliography

- Barnett, P. A., Roman-Golstein, S., Ramsey, F., et al. (1995). Differential permeability and quantitative mr imaging of a human lung carcinoma brain xenograft in the nude rat. *American Journal of Pathology*, 146(2):436–449.
- Berkhemer, O. A., Fransen, P. S. S., Buemer, D., et al. (2015). A randomized trial of intraarterial treatment for acute ischemic stroke. *New England Journal of Medicine*, 372:11–20.
- Ramsey, F. L. and Schafer, D. W. (2002). The Statistical Sleuth: A Course in Methods of Data Analysis. Duxbury, Pacific Grove, CA, second edition.
- Roy, D., Talajic, M., Nattel, S., et al. (2008). Rhythm control versus rate control for atrial fibrillation and heart failure. *New England Journal of Medicine*, 358:2667–2677.
- Tolaney, S. M., Barry, W. T., Chau, T. D., et al. (2015). Adjuvant paclitaxel and trastuzumab for nodengative, her2-positive breast cancer. *New England Journal of Medicine*, 372:134–141.