

Book by M. A. Hernán and J. M. Robins — R code by Joy Shi and Sean McGrath Stata code by Eleanor Murray and Roger Logan — R Markdown code by Tom Palmer

 $14~\mathrm{May}~2020$ 

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# **Preface**

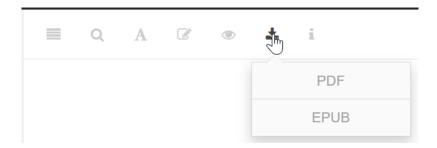
This book presents code examples from Hernán and Robins (2020), which is available in draft form from the following webpage.

https://www.hsph.harvard.edu/miguel-hernan/causal-inference-book/

The R code is based on the code by Joy Shi and Sean McGrath given here.

The Stata code is based on the code by Eleanor Murray and Roger Logan given here.

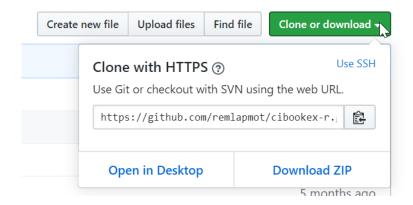
This repo is rendered at https://remlapmot.github.io/cibookex-r/. Click the download button above for the pdf and eBook versions.



# Downloading the code

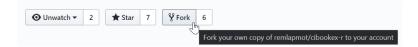
The repo is available on GitHub here. There are a number of ways to download the code. Either,

• click the green Clone or download button then choose to Open in Desktop or Download ZIP.



The *Desktop* option means open in the GitHub Desktop app (if you have that installed on your machine). The *ZIP* option will give you a zip archive of the repo, which you then unzip.

 or fork the repo into your own GitHub account and then clone or download your forked repo to your machine.



## Installing dependency packages

It is easiest to open the repo in RStudio, as an RStudio project, by doubling click the .Rproj file. This makes sure that R's working directory is at the top level of the repo. If you don't want to open the repo as a project set the working directory to the top level of the repo directories using setwd(). Then run:

```
# install.packages('devtools') # uncomment if devtools not
# installed
devtools::install_dev_deps()
```

# Downloading the datasets

We assume that you have downloaded the data from the Causal Inference Book website and saved it to a data subdirectory. You can do this manually or with the following code (nb. we use the here package to reference the data subdirectory).

```
library(here)
```

```
dataurls <- list()
dataurls[[1]] <- "https://cdn1.sph.harvard.edu/wp-content/uploads/sites/1268/2012/10/nhefs_sas.zip"
dataurls[[2]] <- "https://cdn1.sph.harvard.edu/wp-content/uploads/sites/1268/2012/10/nhefs_stata.zip"
dataurls[[3]] <- "https://cdn1.sph.harvard.edu/wp-content/uploads/sites/1268/2017/01/nhefs_excel.zip"
dataurls[[4]] <- "https://cdn1.sph.harvard.edu/wp-content/uploads/sites/1268/1268/20/nhefs.csv"

temp <- tempfile()
for (i in 1:3) {
    download.file(dataurls[[i]], temp)
    unzip(temp, exdir = "data")
}
download.file(dataurls[[4]], here("data", "nhefs.csv"))</pre>
```

# R code

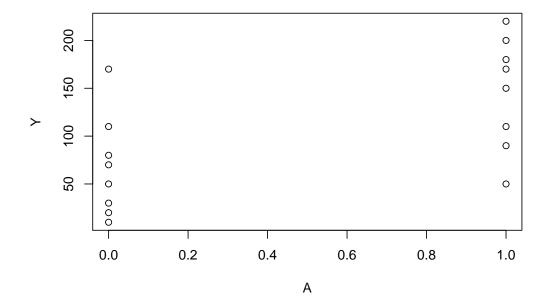
# 11. Why model?

# Program 11.1

- Sample averages by treatment level
- $\bullet$  Data from Figures 11.1 and 11.2

```
A <- c(1, 1, 1, 1, 1, 1, 1, 1, 0, 0, 0, 0, 0, 0, 0, 0)
Y <- c(200, 150, 220, 110, 50, 180, 90, 170, 170, 30, 70, 110, 80, 50, 10, 20)

plot(A, Y)
```



```
summary(Y[A == 0])
```

#### summary(Y[A == 1]) ## Min. 1st Qu. Median Mean 3rd Qu. Max. 50.0 105.0 ## 160.0 146.2 185.0 220.0 $A2 \leftarrow c(1, 1, 1, 1, 2, 2, 2, 2, 3, 3, 3, 3, 4, 4, 4, 4)$ Y2 <- c(110, 80, 50, 40, 170, 30, 70, 50, 110, 50, 180, 130, 200, 150, 220, 210) plot(A2, Y2) 000 200 0 0 150 0 0 0 0 100 0 0 00 0 0 0 1.0 1.5 2.0 2.5 3.0 3.5 4.0 Α2 summary(Y2[A2 == 1]) ## Min. 1st Qu. Median Mean 3rd Qu. Max. ## 40.0 47.5 65.0 70.0 87.5 110.0 summary(Y2[A2 == 2]) ## Min. 1st Qu. Median Mean 3rd Qu. Max. ## 30 45 60 80 95 170 summary(Y2[A2 == 3]) ## Min. 1st Qu. Median Mean 3rd Qu. Max.

180.0

117.5 142.5

##

50.0 95.0

120.0

```
summary(Y2[A2 == 4])
```

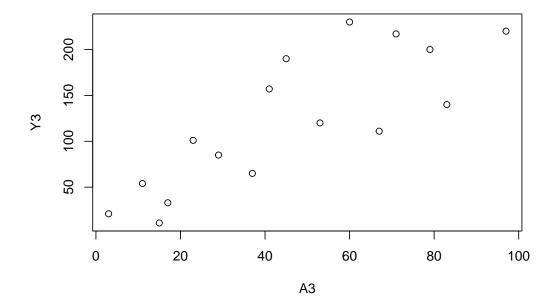
```
## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 150.0 187.5 205.0 195.0 212.5 220.0
```

- 2-parameter linear model
- Data from Figures 11.3 and 11.1

```
A3 <-
    c(3, 11, 17, 23, 29, 37, 41, 53, 67, 79, 83, 97, 60, 71, 15, 45)

Y3 <-
    c(21, 54, 33, 101, 85, 65, 157, 120, 111, 200, 140, 220, 230, 217,
        11, 190)

plot(Y3 ~ A3)
```



```
summary(glm(Y3 ~ A3))
```

```
##
## Call:
## glm(formula = Y3 ~ A3)
##
```

```
## Deviance Residuals:
##
           1Q Median 3Q
                                         Max
## -61.930 -30.564 -5.741 30.653
                                      77.225
##
## Coefficients:
              Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) 24.5464
                                   1.151 0.269094
                          21.3300
                2.1372
                           0.3997
                                   5.347 0.000103 ***
## A3
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 1944.109)
##
      Null deviance: 82800 on 15 degrees of freedom
##
## Residual deviance: 27218 on 14 degrees of freedom
## AIC: 170.43
## Number of Fisher Scoring iterations: 2
predict(glm(Y3 ~ A3), data.frame(A3 = 90))
##
## 216.89
summary(glm(Y ~ A))
##
## Call:
## glm(formula = Y \sim A)
## Deviance Residuals:
      Min
                1Q Median
                                 3Q
## -96.250 -40.000
                   3.125 35.938 102.500
##
## Coefficients:
              Estimate Std. Error t value Pr(>|t|)
                                   3.424 0.00412 **
## (Intercept)
                 67.50
                            19.72
                 78.75
                            27.88
                                   2.824 0.01352 *
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 3109.821)
##
##
      Null deviance: 68344 on 15 degrees of freedom
## Residual deviance: 43538 on 14 degrees of freedom
## AIC: 177.95
##
## Number of Fisher Scoring iterations: 2
```

## 197.1269

- 3-parameter linear model
- Data from Figure 11.3

```
Asq <- A3 * A3
mod3 <- glm(Y3 ~ A3 + Asq)
summary(mod3)
##
## Call:
## glm(formula = Y3 ~ A3 + Asq)
##
## Deviance Residuals:
     Min
              1Q Median
                               3Q
                                     Max
## -65.27 -34.41 13.21 26.11
                                    64.36
##
## Coefficients:
              Estimate Std. Error t value Pr(>|t|)
## (Intercept) -7.40688
                         31.74777 -0.233
                                            0.8192
## A3
               4.10723
                          1.53088
                                     2.683
                                            0.0188 *
              -0.02038
                          0.01532 -1.331
                                            0.2062
## Asq
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 1842.697)
##
##
       Null deviance: 82800 on 15 degrees of freedom
## Residual deviance: 23955 on 13 degrees of freedom
## AIC: 170.39
##
## Number of Fisher Scoring iterations: 2
predict(mod3, data.frame(cbind(A3 = 90, Asq = 8100)))
##
          1
```

# 12. IP Weighting and Marginal Structural Models

# Program 12.1

library(here)

• Descriptive statistics from NHEFS data (Table 12.1)

```
# install.packages("readxl") # install package if required
library("readxl")

nhefs <- read_excel(here("data", "NHEFS.xls"))
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)

# provisionally ignore subjects with missing values for weight in 1982
nhefs.nmv <-
nhefs[which(!is.na(nhefs$wt82)),]

lm(wt82_71 ~ qsmk, data = nhefs.nmv)</pre>
```

```
##
## Call:
## lm(formula = wt82_71 ~ qsmk, data = nhefs.nmv)
##
## Coefficients:
## (Intercept) qsmk
## 1.984 2.541

# Smoking cessation
predict(lm(wt82_71 ~ qsmk, data = nhefs.nmv), data.frame(qsmk = 1))
```

```
## 1
## 4.525079
```

```
# No smoking cessation
predict(lm(wt82_71 ~ qsmk, data = nhefs.nmv), data.frame(qsmk = 0))
         1
## 1.984498
# Table
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 0),]$age)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
##
    25.00 33.00 42.00
                           42.79
                                   51.00 72.00
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 0),]$wt71)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
##
    40.82
          59.19
                  68.49
                          70.30 79.38 151.73
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 0),]$smokeintensity)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
##
     1.00 15.00
                   20.00
                           21.19 30.00
                                           60.00
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 0),]$smokeyrs)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
                   23.00
                           24.09
                                   32.00
##
     1.00
           15.00
                                           64.00
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 1),]$age)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
    25.00 35.00 46.00 46.17 56.00 74.00
##
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 1),]$wt71)
##
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
          60.67
                   71.21
                           72.35
                                   81.08 136.98
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 1),]$smokeintensity)
     Min. 1st Qu. Median
##
                            Mean 3rd Qu.
                                            Max.
##
      1.0 10.0
                    20.0
                            18.6
                                    25.0
                                            80.0
```

```
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 1),]$smokeyrs)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                              Max.
##
      1.00
           15.00
                     26.00
                             26.03
                                     35.00
                                             60.00
table(nhefs.nmv$qsmk, nhefs.nmv$sex)
##
##
         0
             1
     0 542 621
##
     1 220 183
##
prop.table(table(nhefs.nmv$qsmk, nhefs.nmv$sex), 1)
##
##
               0
     0 0.4660361 0.5339639
##
     1 0.5459057 0.4540943
##
table(nhefs.nmv$qsmk, nhefs.nmv$race)
##
##
         0
##
     0 993 170
     1 367 36
prop.table(table(nhefs.nmv$qsmk, nhefs.nmv$race), 1)
##
                0
##
     0 0.85382631 0.14617369
##
##
     1 0.91066998 0.08933002
table(nhefs.nmv$qsmk, nhefs.nmv$education)
##
##
         1
             2
                 3
                     4
     0 210 266 480 92 115
     1 81 74 157 29 62
prop.table(table(nhefs.nmv$qsmk, nhefs.nmv$education), 1)
##
##
     0 0.18056750 0.22871883 0.41272571 0.07910576 0.09888220
##
     1 0.20099256 0.18362283 0.38957816 0.07196030 0.15384615
##
```

```
table(nhefs.nmv$qsmk, nhefs.nmv$exercise)
##
##
         0
             1
     0 237 485 441
##
     1 63 176 164
##
prop.table(table(nhefs.nmv$qsmk, nhefs.nmv$exercise), 1)
##
##
               0
##
     0 0.2037833 0.4170249 0.3791917
     1 0.1563275 0.4367246 0.4069479
##
table(nhefs.nmv$qsmk, nhefs.nmv$active)
##
##
         0 1
##
     0 532 527 104
     1 170 188 45
##
prop.table(table(nhefs.nmv$qsmk, nhefs.nmv$active), 1)
##
               0
##
     0 0.4574377 0.4531384 0.0894239
##
     1 0.4218362 0.4665012 0.1116625
##
```

- Estimating IP weights
- Data from NHEFS

```
# Estimation of ip weights via a logistic model
fit <- glm(
  qsmk ~ sex + race + age + I(age ^ 2) +
    as.factor(education) + smokeintensity +
    I(smokeintensity ^ 2) + smokeyrs + I(smokeyrs ^ 2) +
    as.factor(exercise) + as.factor(active) + wt71 + I(wt71 ^ 2),
  family = binomial(),
  data = nhefs.nmv
)
summary(fit)</pre>
```

```
##
## Call:
## glm(formula = qsmk ~ sex + race + age + I(age^2) + as.factor(education) +
      smokeintensity + I(smokeintensity^2) + smokeyrs + I(smokeyrs^2) +
      as.factor(exercise) + as.factor(active) + wt71 + I(wt71^2),
##
      family = binomial(), data = nhefs.nmv)
##
##
## Deviance Residuals:
               1Q
##
      Min
                    Median
                                3Q
                                       Max
## -1.5127 -0.7907 -0.6387
                                     2.3729
                            0.9832
##
## Coefficients:
                         Estimate Std. Error z value Pr(>|z|)
##
                       -2.2425191 1.3808360 -1.624 0.104369
## (Intercept)
## sex
                       -0.5274782   0.1540496   -3.424   0.000617 ***
## race
                       2.364 0.018068 *
## age
                        0.1212052 0.0512663
                       ## I(age^2)
## as.factor(education)2 -0.0287755 0.1983506 -0.145 0.884653
## as.factor(education)3 0.0864318 0.1780850
                                           0.485 0.627435
## as.factor(education)4 0.0636010 0.2732108 0.233 0.815924
## as.factor(education)5 0.4759606 0.2262237
                                             2.104 0.035384 *
                       -0.0772704 0.0152499 -5.067 4.04e-07 ***
## smokeintensity
                                            3.647 0.000265 ***
## I(smokeintensity^2)
                        0.0010451 0.0002866
## smokeyrs
                       -0.0735966  0.0277775  -2.650  0.008061 **
                        0.0008441 0.0004632 1.822 0.068398 .
## I(smokeyrs^2)
## as.factor(exercise)1 0.3548405 0.1801351 1.970 0.048855 *
## as.factor(exercise)2  0.3957040  0.1872400  2.113  0.034571 *
## as.factor(active)1
                        0.0319445 0.1329372 0.240 0.810100
                        0.1767840 0.2149720 0.822 0.410873
## as.factor(active)2
## wt71
                       ## I(wt71^2)
                        0.0001352 0.0001632 0.829 0.407370
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
  (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 1786.1 on 1565 degrees of freedom
## Residual deviance: 1676.9 on 1547
                                   degrees of freedom
## AIC: 1714.9
## Number of Fisher Scoring iterations: 4
p.qsmk.obs <-
 ifelse(nhefs.nmv$qsmk == 0,
        1 - predict(fit, type = "response"),
        predict(fit, type = "response"))
nhefs.nmv$w <- 1 / p.qsmk.obs
```

```
summary(nhefs.nmv$w)
##
      Min. 1st Qu. Median Mean 3rd Qu.
                                              Max.
##
     1.054 1.230 1.373 1.996 1.990 16.700
sd(nhefs.nmv$w)
## [1] 1.474787
# install.packages("geepack") # install package if required
library("geepack")
msm.w <- geeglm(
  wt82_{71} \sim qsmk
 data = nhefs.nmv,
  weights = w,
 id = seqn,
  corstr = "independence"
summary(msm.w)
##
## Call:
## geeglm(formula = wt82_71 ~ qsmk, data = nhefs.nmv, weights = w,
       id = seqn, corstr = "independence")
##
##
## Coefficients:
               Estimate Std.err Wald Pr(>|W|)
##
## (Intercept) 1.7800 0.2247 62.73 2.33e-15 ***
                 3.4405 0.5255 42.87 5.86e-11 ***
## qsmk
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Correlation structure = independence
## Estimated Scale Parameters:
##
               Estimate Std.err
##
## (Intercept)
                 65.06 4.221
## Number of clusters: 1566 Maximum cluster size: 1
beta <- coef(msm.w)</pre>
SE <- coef(summary(msm.w))[, 2]</pre>
lcl <- beta - qnorm(0.975) * SE</pre>
ucl <- beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
##
                beta lcl ucl
## (Intercept) 1.780 1.340 2.22
## qsmk
              3.441 2.411 4.47
```

```
{\it \# no \ association \ between \ sex \ and \ qsmk \ in \ pseudo-population}
xtabs(nhefs.nmv$w ~ nhefs.nmv$sex + nhefs.nmv$qsmk)
                nhefs.nmv$qsmk
## nhefs.nmv$sex
                     0
##
               0 763.6 763.6
##
               1 801.7 797.2
# "check" for positivity (White women)
table(nhefs.nmv$age[nhefs.nmv$race == 0 & nhefs.nmv$sex == 1],
     nhefs.nmv$qsmk[nhefs.nmv$race == 0 & nhefs.nmv$sex == 1])
##
##
         0 1
##
     25 24 3
##
     26 14 5
##
     27 18 2
##
     28 20 5
##
     29 15 4
##
     30 14 5
##
     31 11 5
##
     32 14 7
##
     33 12 3
##
     34 22 5
     35 16 5
##
##
     36 13 3
##
     37 14 1
##
     38 6 2
##
     39 19 4
##
     40 10 4
##
     41 13 3
##
     42 16 3
     43 14 3
##
##
     44 9 4
##
     45 12 5
##
     46 19 4
##
     47 19 4
##
     48 19 4
     49 11 3
##
##
     50 18 4
     51 9 3
##
##
     52 11 3
##
     53 11 4
     54 17 9
##
     55 9 4
##
##
     56 8 7
##
     57 9 2
```

##

58 8 4

```
##
    59 5 4
##
    60 5 4
##
    61
       5
##
    62 6 5
##
    63 3 3
##
    64 7 1
##
    65 3 2
    66 4 0
##
    67 2 0
##
##
    69 6 2
    70 2 1
##
##
    71 0 1
    72 2 2
##
    74 0 1
##
```

- Estimating stabilized IP weights
- Data from NHEFS

```
# estimation of denominator of ip weights
denom.fit <-
glm(
    qsmk ~ as.factor(sex) + as.factor(race) + age + I(age ^ 2) +
    as.factor(education) + smokeintensity +
    I(smokeintensity ^ 2) + smokeyrs + I(smokeyrs ^ 2) +
    as.factor(exercise) + as.factor(active) + wt71 + I(wt71 ^ 2),
    family = binomial(),
    data = nhefs.nmv
)
summary(denom.fit)</pre>
```

```
##
## Call:
  glm(formula = qsmk ~ as.factor(sex) + as.factor(race) + age +
       I(age^2) + as.factor(education) + smokeintensity + I(smokeintensity^2) +
       smokeyrs + I(smokeyrs^2) + as.factor(exercise) + as.factor(active) +
##
       wt71 + I(wt71^2), family = binomial(), data = nhefs.nmv)
##
##
## Deviance Residuals:
##
     Min
              1Q Median
                               3Q
                                     Max
## -1.513 -0.791 -0.639
                          0.983
                                    2.373
##
## Coefficients:
##
                         Estimate Std. Error z value Pr(>|z|)
                                              -1.62 0.10437
## (Intercept)
                        -2.242519
                                     1.380836
## as.factor(sex)1
                        -0.527478
                                     0.154050 -3.42 0.00062 ***
## as.factor(race)1
                                    0.210067 -4.00 6.5e-05 ***
                        -0.839264
```

```
## age
                         0.121205
                                    0.051266
                                                2.36 0.01807 *
## I(age^2)
                        -0.000825
                                    0.000536
                                               -1.54 0.12404
## as.factor(education)2 -0.028776
                                    0.198351
                                               -0.15 0.88465
## as.factor(education)3 0.086432
                                    0.178085
                                                0.49 0.62744
## as.factor(education)4 0.063601
                                    0.273211
                                                0.23 0.81592
## as.factor(education)5 0.475961
                                    0.226224
                                                2.10 0.03538 *
## smokeintensity
                        -0.077270
                                    0.015250
                                               -5.07 4.0e-07 ***
## I(smokeintensity^2)
                                                3.65 0.00027 ***
                         0.001045
                                    0.000287
## smokeyrs
                                               -2.65 0.00806 **
                        -0.073597
                                    0.027777
## I(smokeyrs^2)
                         0.000844
                                    0.000463
                                                1.82 0.06840 .
## as.factor(exercise)1
                                    0.180135
                                                1.97 0.04885 *
                         0.354841
## as.factor(exercise)2 0.395704
                                    0.187240
                                                2.11 0.03457 *
## as.factor(active)1
                                                0.24 0.81010
                         0.031944
                                    0.132937
## as.factor(active)2
                         0.176784
                                    0.214972
                                                0.82 0.41087
## wt71
                        -0.015236
                                    0.026316
                                               -0.58 0.56262
## I(wt71^2)
                         0.000135
                                    0.000163
                                                0.83 0.40737
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
  (Dispersion parameter for binomial family taken to be 1)
##
##
      Null deviance: 1786.1 on 1565 degrees of freedom
## Residual deviance: 1676.9 on 1547 degrees of freedom
## AIC: 1715
##
## Number of Fisher Scoring iterations: 4
pd.qsmk <- predict(denom.fit, type = "response")</pre>
# estimation of numerator of ip weights
numer.fit <- glm(qsmk ~ 1, family = binomial(), data = nhefs.nmv)</pre>
summary(numer.fit)
##
## Call:
## glm(formula = qsmk ~ 1, family = binomial(), data = nhefs.nmv)
## Deviance Residuals:
##
     Min
              1Q Median
                              3Q
                                     Max
## -0.771 -0.771 -0.771
                                   1.648
                           1.648
##
## Coefficients:
              Estimate Std. Error z value Pr(>|z|)
##
                           0.0578 -18.3 <2e-16 ***
## (Intercept) -1.0598
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
```

```
Null deviance: 1786.1 on 1565 degrees of freedom
## Residual deviance: 1786.1 on 1565 degrees of freedom
## AIC: 1788
##
## Number of Fisher Scoring iterations: 4
pn.qsmk <- predict(numer.fit, type = "response")</pre>
nhefs.nmv$sw <-
  ifelse(nhefs.nmv\$qsmk == 0, ((1 - pn.qsmk) / (1 - pd.qsmk)),
         (pn.qsmk / pd.qsmk))
summary(nhefs.nmv$sw)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
##
     0.331
           0.867
                    0.950
                             0.999
                                    1.079
                                             4.298
msm.sw <- geeglm(</pre>
 wt82_{71} \sim qsmk
 data = nhefs.nmv,
 weights = sw,
 id = seqn,
  corstr = "independence"
)
summary(msm.sw)
##
## Call:
## geeglm(formula = wt82_71 ~ qsmk, data = nhefs.nmv, weights = sw,
       id = seqn, corstr = "independence")
##
## Coefficients:
               Estimate Std.err Wald Pr(>|W|)
##
                 1.780 0.225 62.7 2.3e-15 ***
## (Intercept)
                  3.441 0.525 42.9 5.9e-11 ***
## qsmk
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Correlation structure = independence
## Estimated Scale Parameters:
##
               Estimate Std.err
##
## (Intercept)
                   60.7
                           3.71
## Number of clusters: 1566 Maximum cluster size: 1
beta <- coef(msm.sw)</pre>
SE <- coef(summary(msm.sw))[, 2]</pre>
lcl <- beta - qnorm(0.975) * SE</pre>
```

```
ucl <- beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
##
              beta lcl ucl
## (Intercept) 1.78 1.34 2.22
## qsmk
              3.44 2.41 4.47
# no association between sex and qsmk in pseudo-population
xtabs(nhefs.nmv$sw ~ nhefs.nmv$sex + nhefs.nmv$qsmk)
##
               nhefs.nmv$qsmk
## nhefs.nmv$sex
                 0 1
              0 567 197
##
              1 595 205
##
```

• Estimating the parameters of a marginal structural mean model with a continuous treatment Data from NHEFS

```
# Analysis restricted to subjects reporting <=25 cig/day at baseline
nhefs.nmv.s <- subset(nhefs.nmv, smokeintensity <= 25)</pre>
# estimation of denominator of ip weights
den.fit.obj <- lm(</pre>
  smkintensity82_71 ~ as.factor(sex) +
    as.factor(race) + age + I(age ^ 2) +
    as.factor(education) + smokeintensity + I(smokeintensity ^ 2) +
    smokeyrs + I(smokeyrs ^ 2) + as.factor(exercise) + as.factor(active) + wt71 +
    I(wt71 ^ 2),
  data = nhefs.nmv.s
p.den <- predict(den.fit.obj, type = "response")</pre>
dens.den <-
  dnorm(nhefs.nmv.s$smkintensity82_71,
        p.den,
        summary(den.fit.obj)$sigma)
# estimation of numerator of ip weights
num.fit.obj <- lm(smkintensity82_71 ~ 1, data = nhefs.nmv.s)</pre>
p.num <- predict(num.fit.obj, type = "response")</pre>
dens.num <-
  dnorm(nhefs.nmv.s$smkintensity82_71,
        p.num,
        summary(num.fit.obj)$sigma)
```

```
nhefs.nmv.s$sw.a <- dens.num / dens.den
summary(nhefs.nmv.s$sw.a)
##
     Min. 1st Qu. Median
                              Mean 3rd Qu.
                                              Max.
##
      0.19
           0.89
                      0.97
                              1.00
                                      1.05
                                              5.10
msm.sw.cont <-
 geeglm(
   wt82_71 ~ smkintensity82_71 + I(smkintensity82_71 * smkintensity82_71),
   data = nhefs.nmv.s,
   weights = sw.a,
   id = seqn,
   corstr = "independence"
 )
summary(msm.sw.cont)
##
## Call:
## geeglm(formula = wt82_71 ~ smkintensity82_71 + I(smkintensity82_71 *
       smkintensity82_71), data = nhefs.nmv.s, weights = sw.a, id = seqn,
##
       corstr = "independence")
##
  Coefficients:
##
##
                                            Estimate Std.err Wald Pr(>|W|)
                                             2.00452  0.29512  46.13  1.1e-11 ***
## (Intercept)
## smkintensity82_71
                                            -0.10899 0.03154 11.94 0.00055 ***
## I(smkintensity82_71 * smkintensity82_71) 0.00269 0.00242 1.24 0.26489
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Correlation structure = independence
## Estimated Scale Parameters:
##
##
               Estimate Std.err
                            4.5
                   60.5
## (Intercept)
## Number of clusters:
                        1162 Maximum cluster size: 1
beta <- coef(msm.sw.cont)</pre>
SE <- coef(summary(msm.sw.cont))[, 2]
lcl <- beta - qnorm(0.975) * SE</pre>
ucl <- beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
##
                                                beta
                                                          lcl
                                                                   ucl
## (Intercept)
                                             2.00452 1.42610 2.58295
## smkintensity82_71
                                            -0.10899 -0.17080 -0.04718
## I(smkintensity82_71 * smkintensity82_71) 0.00269 -0.00204 0.00743
```

- Estimating the parameters of a marginal structural logistic model
- Data from NHEFS

```
table(nhefs.nmv$qsmk, nhefs.nmv$death)
```

```
##
## 0 1
## 0 963 200
## 1 312 91

# First, estimation of stabilized weights sw (same as in Program 12.3)
# Second, fit logistic model below
msm.logistic <- geeglm(
   death ~ qsmk,
   data = nhefs.nmv,
   weights = sw,
   id = seqn,
   family = binomial(),
   corstr = "independence"
)</pre>
```

## Warning in eval(family\$initialize): non-integer #successes in a binomial glm!

```
summary(msm.logistic)
```

```
##
## Call:
## geeglm(formula = death ~ qsmk, family = binomial(), data = nhefs.nmv,
      weights = sw, id = seqn, corstr = "independence")
##
##
  Coefficients:
##
##
              Estimate Std.err
                                 Wald Pr(>|W|)
## (Intercept) -1.4905 0.0789 356.50
                                        <2e-16 ***
                0.0301 0.1573
                                          0.85
## qsmk
                               0.04
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Correlation structure = independence
## Estimated Scale Parameters:
##
##
              Estimate Std.err
## (Intercept)
                     1 0.0678
## Number of clusters:
                       1566 Maximum cluster size: 1
```

- Assessing effect modification by sex using a marginal structural mean model
- Data from NHEFS

```
table(nhefs.nmv$sex)
##
##
    0
## 762 804
# estimation of denominator of ip weights
denom.fit <-
  glm(
   qsmk ~ as.factor(sex) + as.factor(race) + age + I(age ^ 2) +
      as.factor(education) + smokeintensity +
      I(smokeintensity ^ 2) + smokeyrs + I(smokeyrs ^ 2) +
      as.factor(exercise) + as.factor(active) + wt71 + I(wt71 ^ 2),
   family = binomial(),
   data = nhefs.nmv
  )
summary(denom.fit)
```

```
##
## Call:
## glm(formula = qsmk ~ as.factor(sex) + as.factor(race) + age +
       I(age^2) + as.factor(education) + smokeintensity + I(smokeintensity^2) +
##
       smokeyrs + I(smokeyrs^2) + as.factor(exercise) + as.factor(active) +
##
##
       wt71 + I(wt71^2), family = binomial(), data = nhefs.nmv)
##
## Deviance Residuals:
               1Q Median
##
     Min
                               3Q
                                      Max
## -1.513 -0.791 -0.639 0.983
                                    2.373
##
## Coefficients:
                          Estimate Std. Error z value Pr(>|z|)
##
```

```
## (Intercept)
                        -2.242519
                                    1.380836
                                               -1.62 0.10437
## as.factor(sex)1
                        -0.527478
                                    0.154050
                                               -3.42 0.00062 ***
## as.factor(race)1
                        -0.839264
                                    0.210067
                                               -4.00 6.5e-05 ***
## age
                         0.121205
                                    0.051266
                                                2.36 0.01807 *
## I(age^2)
                        -0.000825
                                    0.000536
                                               -1.54 0.12404
## as.factor(education)2 -0.028776
                                    0.198351
                                               -0.15 0.88465
## as.factor(education)3 0.086432
                                    0.178085
                                                0.49 0.62744
## as.factor(education)4 0.063601
                                    0.273211
                                                0.23 0.81592
## as.factor(education)5 0.475961
                                    0.226224
                                                2.10 0.03538 *
## smokeintensity
                                               -5.07 4.0e-07 ***
                        -0.077270
                                    0.015250
## I(smokeintensity^2)
                                                3.65 0.00027 ***
                         0.001045
                                    0.000287
## smokeyrs
                        -0.073597
                                    0.027777
                                               -2.65 0.00806 **
## I(smokeyrs^2)
                         0.000844
                                    0.000463
                                                1.82 0.06840 .
## as.factor(exercise)1 0.354841
                                    0.180135
                                                1.97 0.04885 *
## as.factor(exercise)2
                                    0.187240
                                                2.11 0.03457 *
                         0.395704
## as.factor(active)1
                         0.031944
                                   0.132937
                                                0.24 0.81010
## as.factor(active)2
                         0.176784
                                    0.214972
                                                0.82 0.41087
## wt71
                        -0.015236
                                    0.026316
                                              -0.58 0.56262
## I(wt71^2)
                         0.000135
                                    0.000163
                                                0.83 0.40737
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
##
      Null deviance: 1786.1 on 1565 degrees of freedom
## Residual deviance: 1676.9 on 1547 degrees of freedom
## AIC: 1715
##
## Number of Fisher Scoring iterations: 4
pd.qsmk <- predict(denom.fit, type = "response")</pre>
# estimation of numerator of ip weights
numer.fit <-
 glm(qsmk ~ as.factor(sex), family = binomial(), data = nhefs.nmv)
summary(numer.fit)
##
## Call:
## glm(formula = qsmk ~ as.factor(sex), family = binomial(), data = nhefs.nmv)
##
## Deviance Residuals:
##
     Min
              1Q Median
                              3Q
                                     Max
## -0.825 -0.825 -0.719 1.576
                                   1.720
##
## Coefficients:
##
                  Estimate Std. Error z value Pr(>|z|)
                   -0.9016
## (Intercept)
                               0.0799 -11.28
                                                <2e-16 ***
## as.factor(sex)1 -0.3202
                               0.1160
                                      -2.76
                                                0.0058 **
```

```
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 1786.1 on 1565 degrees of freedom
##
## Residual deviance: 1778.4 on 1564 degrees of freedom
## AIC: 1782
## Number of Fisher Scoring iterations: 4
pn.qsmk <- predict(numer.fit, type = "response")</pre>
nhefs.nmv$sw.a <-
  ifelse(nhefs.nmv$qsmk == 0, ((1 - pn.qsmk) / (1 - pd.qsmk)),
         (pn.qsmk / pd.qsmk))
summary(nhefs.nmv$sw.a)
##
     Min. 1st Qu. Median
                              Mean 3rd Qu.
                                              Max.
##
      0.29
             0.88
                      0.96
                              1.00
                                      1.08
                                              3.80
sd(nhefs.nmv$sw.a)
## [1] 0.271
# Estimating parameters of a marginal structural mean model
msm.emm <- geeglm(</pre>
 wt82_71 ~ as.factor(qsmk) + as.factor(sex)
 + as.factor(qsmk):as.factor(sex),
 data = nhefs.nmv,
 weights = sw.a,
 id = seqn,
  corstr = "independence"
summary(msm.emm)
##
## Call:
## geeglm(formula = wt82_71 ~ as.factor(qsmk) + as.factor(sex) +
       as.factor(qsmk):as.factor(sex), data = nhefs.nmv, weights = sw.a,
       id = seqn, corstr = "independence")
##
##
  Coefficients:
##
##
                                    Estimate Std.err Wald Pr(>|W|)
                                     1.78445 0.30984 33.17 8.5e-09 ***
## (Intercept)
## as.factor(qsmk)1
                                     3.52198  0.65707  28.73  8.3e-08 ***
## as.factor(sex)1
                                    -0.00872 0.44882 0.00
                                                                0.98
```

```
## as.factor(qsmk)1:as.factor(sex)1 -0.15948 1.04608 0.02
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Correlation structure = independence
## Estimated Scale Parameters:
##
               Estimate Std.err
##
                   60.8
                           3.71
## (Intercept)
## Number of clusters:
                         1566 Maximum cluster size: 1
beta <- coef(msm.emm)</pre>
SE <- coef(summary(msm.emm))[, 2]
lcl <- beta - qnorm(0.975) * SE</pre>
ucl <- beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
##
                                         beta
                                                 lcl ucl
## (Intercept)
                                      1.78445 1.177 2.392
## as.factor(qsmk)1
                                     3.52198 2.234 4.810
## as.factor(sex)1
                                    -0.00872 -0.888 0.871
## as.factor(qsmk)1:as.factor(sex)1 -0.15948 -2.210 1.891
Program 12.7
  • Estimating IP weights to adjust for selection bias due to censoring
  • Data from NHEFS
table(nhefs$qsmk, nhefs$cens)
##
##
          0
               1
##
     0 1163
              38
     1 403
summary(nhefs[which(nhefs$cens == 0),]$wt71)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
```

```
## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 36.2 63.1 72.1 76.6 87.9 169.2
```

69.2

summary(nhefs[which(nhefs\$cens == 1),]\$wt71)

70.8

79.8

##

39.6

59.5

151.7

```
# estimation of denominator of ip weights for A
denom.fit <-
  glm(
    qsmk ~ as.factor(sex) + as.factor(race) + age + I(age ^ 2) +
      as.factor(education) + smokeintensity +
      I(smokeintensity ^ 2) + smokeyrs + I(smokeyrs ^ 2) +
      as.factor(exercise) + as.factor(active) + wt71 + I(wt71 ^ 2),
   family = binomial(),
    data = nhefs
  )
summary(denom.fit)
##
## Call:
## glm(formula = qsmk ~ as.factor(sex) + as.factor(race) + age +
       I(age^2) + as.factor(education) + smokeintensity + I(smokeintensity^2) +
##
       smokeyrs + I(smokeyrs^2) + as.factor(exercise) + as.factor(active) +
##
       wt71 + I(wt71^2), family = binomial(), data = nhefs)
##
## Deviance Residuals:
     Min
              10 Median
                               30
                                      Max
## -1.465 -0.804 -0.646
                           1.058
                                    2.355
## Coefficients:
##
                          Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                         -1.988902
                                     1.241279
                                                -1.60 0.10909
## as.factor(sex)1
                         -0.507522
                                     0.148232
                                                -3.42 0.00062 ***
## as.factor(race)1
                         -0.850231
                                     0.205872
                                                -4.13 3.6e-05 ***
## age
                          0.103013
                                     0.048900
                                                 2.11 0.03515 *
## I(age^2)
                         -0.000605
                                     0.000507
                                                -1.19 0.23297
## as.factor(education)2 -0.098320
                                     0.190655
                                                -0.52 0.60607
## as.factor(education)3 0.015699
                                     0.170714
                                                 0.09 0.92673
## as.factor(education)4 -0.042526
                                     0.264276
                                                -0.16 0.87216
## as.factor(education)5 0.379663
                                     0.220395
                                                 1.72 0.08495 .
## smokeintensity
                         -0.065156
                                     0.014759
                                                -4.41 1.0e-05 ***
## I(smokeintensity^2)
                          0.000846
                                     0.000276
                                                 3.07 0.00216 **
## smokeyrs
                         -0.073371
                                     0.026996
                                                -2.72 0.00657 **
## I(smokeyrs^2)
                          0.000838
                                     0.000443
                                                 1.89 0.05867 .
## as.factor(exercise)1
                          0.291412
                                     0.173554
                                                 1.68 0.09314 .
## as.factor(exercise)2
                          0.355052
                                     0.179929
                                                 1.97 0.04846 *
## as.factor(active)1
                                     0.129832
                                                 0.08 0.93324
                          0.010875
## as.factor(active)2
                          0.068312
                                     0.208727
                                                 0.33 0.74346
## wt71
                         -0.012848
                                     0.022283
                                                -0.58 0.56423
## I(wt71^2)
                          0.000121
                                     0.000135
                                                 0.89 0.37096
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
```

```
Null deviance: 1876.3 on 1628 degrees of freedom
## Residual deviance: 1766.7 on 1610 degrees of freedom
## AIC: 1805
##
## Number of Fisher Scoring iterations: 4
pd.qsmk <- predict(denom.fit, type = "response")</pre>
# estimation of numerator of ip weights for A
numer.fit <- glm(qsmk ~ 1, family = binomial(), data = nhefs)</pre>
summary(numer.fit)
##
## glm(formula = qsmk ~ 1, family = binomial(), data = nhefs)
##
## Deviance Residuals:
     Min
              1Q Median
                                      Max
## -0.781 -0.781 -0.781
                          1.635
                                    1.635
##
## Coefficients:
##
               Estimate Std. Error z value Pr(>|z|)
## (Intercept) -1.0318
                            0.0563 -18.3 <2e-16 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
       Null deviance: 1876.3 on 1628 degrees of freedom
##
## Residual deviance: 1876.3 on 1628 degrees of freedom
## AIC: 1878
##
## Number of Fisher Scoring iterations: 4
pn.qsmk <- predict(numer.fit, type = "response")</pre>
# estimation of denominator of ip weights for C
denom.cens <- glm(</pre>
  cens ~ as.factor(qsmk) + as.factor(sex) +
   as.factor(race) + age + I(age ^ 2) +
   as.factor(education) + smokeintensity +
   I(smokeintensity ^ 2) + smokeyrs + I(smokeyrs ^ 2) +
    as.factor(exercise) + as.factor(active) + wt71 + I(wt71 ^ 2),
  family = binomial(),
  data = nhefs
summary(denom.cens)
```

##

```
## Call:
## glm(formula = cens ~ as.factor(qsmk) + as.factor(sex) + as.factor(race) +
       age + I(age^2) + as.factor(education) + smokeintensity +
       I(smokeintensity^2) + smokeyrs + I(smokeyrs^2) + as.factor(exercise) +
##
       as.factor(active) + wt71 + I(wt71^2), family = binomial(),
##
       data = nhefs)
##
##
## Deviance Residuals:
##
     Min
               10 Median
                                     Max
## -1.097 -0.287 -0.207 -0.157
                                    2.996
##
## Coefficients:
##
                         Estimate Std. Error z value Pr(>|z|)
                                                 1.56
## (Intercept)
                         4.014466
                                    2.576106
                                                       0.1192
## as.factor(qsmk)1
                         0.516867
                                    0.287716
                                                 1.80
                                                       0.0724 .
## as.factor(sex)1
                         0.057313
                                    0.330278
                                                0.17
                                                       0.8622
## as.factor(race)1
                        -0.012271
                                    0.452489
                                               -0.03
                                                      0.9784
                                               -2.30
## age
                        -0.269729
                                    0.117465
                                                      0.0217 *
                                    0.001114
                                                2.59
                                                      0.0096 **
## I(age^2)
                         0.002884
## as.factor(education)2 -0.440788
                                              -1.05
                                                       0.2933
                                    0.419399
## as.factor(education)3 -0.164688
                                               -0.44
                                                      0.6567
                                    0.370547
## as.factor(education)4 0.138447
                                    0.569797
                                                0.24
                                                       0.8080
## as.factor(education)5 -0.382382
                                    0.560181
                                               -0.68
                                                       0.4949
                                                0.45
## smokeintensity
                         0.015712
                                    0.034732
                                                       0.6510
## I(smokeintensity^2)
                        -0.000113
                                    0.000606
                                              -0.19
                                                      0.8517
## smokeyrs
                                    0.074958
                                                1.05
                                                       0.2944
                         0.078597
## I(smokeyrs^2)
                        -0.000557
                                    0.001032
                                               -0.54
                                                      0.5894
## as.factor(exercise)1 -0.971471
                                    0.387810
                                               -2.51
                                                       0.0122 *
## as.factor(exercise)2 -0.583989
                                    0.372313
                                               -1.57
                                                      0.1168
## as.factor(active)1
                        -0.247479
                                    0.325455
                                               -0.76
                                                       0.4470
## as.factor(active)2
                         0.706583
                                    0.396458
                                                1.78
                                                       0.0747 .
## wt71
                        -0.087887
                                    0.040012
                                               -2.20
                                                       0.0281 *
## I(wt71^2)
                         0.000635
                                    0.000226
                                                2.81
                                                       0.0049 **
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
       Null deviance: 533.36 on 1628 degrees of freedom
## Residual deviance: 465.36 on 1609 degrees of freedom
## AIC: 505.4
##
## Number of Fisher Scoring iterations: 7
pd.cens <- 1 - predict(denom.cens, type = "response")</pre>
# estimation of numerator of ip weights for C
numer.cens <-
  glm(cens ~ as.factor(qsmk), family = binomial(), data = nhefs)
```

```
summary(numer.cens)
##
## Call:
## glm(formula = cens ~ as.factor(qsmk), family = binomial(), data = nhefs)
## Deviance Residuals:
     Min
               1Q Median
## -0.347 -0.254 -0.254 -0.254
                                    2.628
##
## Coefficients:
                    Estimate Std. Error z value Pr(>|z|)
##
## (Intercept)
                      -3.421
                                  0.165 -20.75
                                                  <2e-16 ***
                                  0.264
                                           2.43
                                                    0.015 *
## as.factor(qsmk)1
                       0.641
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 533.36 on 1628 degrees of freedom
## Residual deviance: 527.76 on 1627 degrees of freedom
## AIC: 531.8
## Number of Fisher Scoring iterations: 6
pn.cens <- 1 - predict(numer.cens, type = "response")</pre>
nhefs$sw.a <-
  ifelse(nhefs\$qsmk == 0, ((1 - pn.qsmk) / (1 - pd.qsmk)),
         (pn.qsmk / pd.qsmk))
nhefs$sw.c <- pn.cens / pd.cens</pre>
nhefs$sw <- nhefs$sw.c * nhefs$sw.a
summary(nhefs$sw.a)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
      0.33
              0.86
##
                      0.95
                              1.00
                                      1.08
                                              4.21
sd(nhefs$sw.a)
## [1] 0.284
summary(nhefs$sw.c)
      Min. 1st Qu. Median
                              Mean 3rd Qu.
##
                                              Max.
##
      0.94
              0.98
                      0.99
                              1.01
                                      1.01
                                              7.58
```

```
sd(nhefs$sw.c)
## [1] 0.178
summary(nhefs$sw)
##
      Min. 1st Qu. Median
                             Mean 3rd Qu.
                                              Max.
##
      0.35
              0.86
                      0.94
                              1.01
                                      1.08
                                             12.86
sd(nhefs$sw)
## [1] 0.411
msm.sw <- geeglm(
  wt82_{71} \sim qsmk,
 data = nhefs,
 weights = sw,
 id = seqn,
  corstr = "independence"
summary(msm.sw)
##
## Call:
## geeglm(formula = wt82_71 ~ qsmk, data = nhefs, weights = sw,
##
       id = seqn, corstr = "independence")
##
## Coefficients:
               Estimate Std.err Wald Pr(>|W|)
##
                1.662 0.233 51.0 9.3e-13 ***
## (Intercept)
## qsmk
                  3.496 0.526 44.2 2.9e-11 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Correlation structure = independence
## Estimated Scale Parameters:
##
               Estimate Std.err
##
                   61.8
## (Intercept)
                           3.83
## Number of clusters: 1566 Maximum cluster size: 1
beta <- coef(msm.sw)</pre>
SE <- coef(summary(msm.sw))[, 2]
lcl <- beta - qnorm(0.975) * SE</pre>
ucl <- beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
```

# 13. Standardization and the parametric G-formula

- Estimating the mean outcome within levels of treatment and confounders
- Data from NHEFS

```
library(here)
```

```
#install.packages("readxl") # install package if required
library("readxl")
nhefs <- read_excel(here("data", "NHEFS.xls"))

# some preprocessing of the data
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)

fit <-
glm(
    wt82_71 ~ qsmk + sex + race + age + I(age * age) + as.factor(education)
    + smokeintensity + I(smokeintensity * smokeintensity) + smokeyrs
    + I(smokeyrs * smokeyrs) + as.factor(exercise) + as.factor(active)
    + wt71 + I(wt71 * wt71) + qsmk * smokeintensity,
    data = nhefs
)
summary(fit)</pre>
```

```
##
## Call:
## glm(formula = wt82_71 ~ qsmk + sex + race + age + I(age * age) +
## as.factor(education) + smokeintensity + I(smokeintensity *
## smokeintensity) + smokeyrs + I(smokeyrs * smokeyrs) + as.factor(exercise) +
## as.factor(active) + wt71 + I(wt71 * wt71) + qsmk * smokeintensity,
## data = nhefs)
##
## Deviance Residuals:
## Min 1Q Median 3Q Max
```

```
## -42.056 -4.171 -0.343
                              3.891
                                     44.606
##
## Coefficients:
##
                                      Estimate Std. Error t value Pr(>|t|)
                                    -1.5881657 4.3130359 -0.368 0.712756
## (Intercept)
## qsmk
                                     2.5595941 0.8091486 3.163 0.001590 **
                                    -1.4302717   0.4689576   -3.050   0.002328 **
## sex
## race
                                     0.3596353  0.1633188  2.202  0.027809 *
## age
## I(age * age)
                                    -0.0061010 0.0017261 -3.534 0.000421 ***
                                     0.7904440 0.6070005 1.302 0.193038
## as.factor(education)2
## as.factor(education)3
                                     0.5563124 0.5561016 1.000 0.317284
## as.factor(education)4
                                     1.4915695 0.8322704 1.792 0.073301 .
## as.factor(education)5
                                    -0.1949770 0.7413692 -0.263 0.792589
## smokeintensity
                                     0.0491365 0.0517254
                                                         0.950 0.342287
## I(smokeintensity * smokeintensity) -0.0009907 0.0009380 -1.056 0.291097
## smokeyrs
                                     0.1343686 0.0917122
                                                         1.465 0.143094
## I(smokeyrs * smokeyrs)
                                   -0.0018664 0.0015437 -1.209 0.226830
## as.factor(exercise)1
                                    0.2959754 0.5351533 0.553 0.580298
## as.factor(exercise)2
                                    ## as.factor(active)1
                                   -0.9475695  0.4099344  -2.312  0.020935 *
## as.factor(active)2
                                   -0.2613779   0.6845577   -0.382   0.702647
## wt71
                                    0.0455018 0.0833709
                                                         0.546 0.585299
## I(wt71 * wt71)
                                   -0.0009653 0.0005247 -1.840 0.066001 .
## qsmk:smokeintensity
                                     0.0466628 0.0351448
                                                         1.328 0.184463
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 53.5683)
##
##
      Null deviance: 97176 on 1565 degrees of freedom
## Residual deviance: 82763 on 1545 degrees of freedom
    (63 observations deleted due to missingness)
## AIC: 10701
## Number of Fisher Scoring iterations: 2
nhefs$predicted.meanY <- predict(fit, nhefs)</pre>
nhefs[which(nhefs$seqn == 24770), c(
 "predicted.meanY",
 "qsmk",
 "sex",
 "race",
 "age",
 "education",
 "smokeintensity",
 "smokeyrs",
 "exercise",
```

```
"active",
 "wt71"
)]
## # A tibble: 1 x 11
##
     predicted.meanY qsmk sex race
                                        age education smokeintensity smokeyrs
              <dbl> <dbl> <dbl> <dbl> <dbl> <
                                                <dbl>
                                                               <dbl>
                                                                        <dbl>
## 1
                        0
                                                                  15
                                                                           12
## # ... with 3 more variables: exercise <dbl>, active <dbl>, wt71 <dbl>
summary(nhefs$predicted.meanY[nhefs$cens == 0])
      Min. 1st Qu. Median
                             Mean 3rd Qu.
                                             Max.
## -10.876
           1.116 3.042
                            2.638
                                            9.876
summary(nhefs$wt82_71[nhefs$cens == 0])
      Min. 1st Qu. Median
                             Mean 3rd Qu.
                                             Max.
## -41.280 -1.478 2.604
                            2.638
                                    6.690 48.538
```

- Standardizing the mean outcome to the baseline confounders
- Data from Table 2.2

```
id <- c(
  "Rheia",
  "Kronos",
  "Demeter",
  "Hades",
  "Hestia",
  "Poseidon",
  "Hera",
  "Zeus",
  "Artemis",
  "Apollo",
  "Leto",
  "Ares",
  "Athena",
  "Hephaestus",
  "Aphrodite",
  "Cyclope",
  "Persephone",
  "Hermes",
  "Hebe",
  "Dionysus"
```

```
N <- length(id)
L \leftarrow c(0, 0, 0, 0, 0, 0, 0, 1, 1, 1, 1, 1, 1, 1, 1, 1, 1, 1, 1)
A \leftarrow c(0, 0, 0, 0, 1, 1, 1, 1, 0, 0, 0, 1, 1, 1, 1, 1, 1, 1, 1, 1)
Y \leftarrow c(0, 1, 0, 0, 0, 0, 1, 1, 1, 0, 1, 1, 1, 1, 1, 1, 0, 0, 0)
interv \leftarrow rep(-1, N)
observed <- cbind(L, A, Y, interv)</pre>
untreated <- cbind(L, rep(0, N), rep(NA, N), rep(0, N))
treated <- cbind(L, rep(1, N), rep(NA, N), rep(1, N))
table22 <- as.data.frame(rbind(observed, untreated, treated))</pre>
table22$id <- rep(id, 3)
glm.obj <- glm(Y ~ A * L, data = table22)</pre>
summary(glm.obj)
##
## Call:
## glm(formula = Y \sim A * L, data = table22)
##
## Deviance Residuals:
##
        Min
                    1Q
                          Median
                                         3Q
                                                  Max
## -0.66667 -0.25000
                        0.04167
                                  0.33333
                                              0.75000
##
## Coefficients:
##
                 Estimate Std. Error t value Pr(>|t|)
## (Intercept) 2.500e-01 2.552e-01
                                        0.980
                                                  0.342
               -4.164e-16 3.608e-01
                                        0.000
                                                  1.000
## L
                4.167e-01 3.898e-01
                                        1.069
                                                  0.301
## A:L
                3.237e-16 4.959e-01 0.000
                                                  1.000
##
## (Dispersion parameter for gaussian family taken to be 0.2604167)
##
##
       Null deviance: 5.0000 on 19 degrees of freedom
## Residual deviance: 4.1667 on 16 degrees of freedom
     (40 observations deleted due to missingness)
## AIC: 35.385
## Number of Fisher Scoring iterations: 2
table22$predicted.meanY <- predict(glm.obj, table22)</pre>
mean(table22$predicted.meanY[table22$interv == -1])
## [1] 0.5
mean(table22$predicted.meanY[table22$interv == 0])
## [1] 0.5
```

```
mean(table22$predicted.meanY[table22$interv == 1])
```

```
## [1] 0.5
```

- Standardizing the mean outcome to the baseline confounders:
- Data from NHEFS

```
# create a dataset with 3 copies of each subject
nhefs$interv <- -1 # 1st copy: equal to original one
interv0 <- nhefs # 2nd copy: treatment set to 0, outcome to missing
interv0$interv <- 0
interv0$qsmk <- 0
interv0$wt82_71 <- NA
interv1 <- nhefs # 3rd copy: treatment set to 1, outcome to missing
interv1$interv <- 1</pre>
interv1$qsmk <- 1
interv1$wt82_71 <- NA
onesample <- rbind(nhefs, interv0, interv1) # combining datasets</pre>
# linear model to estimate mean outcome conditional on treatment and confounders
# parameters are estimated using original observations only (nhefs)
# parameter estimates are used to predict mean outcome for observations with
# treatment set to 0 (interv=0) and to 1 (interv=1)
std <- glm(</pre>
 wt82_71 ~ qsmk + sex + race + age + I(age * age)
 + as.factor(education) + smokeintensity
 + I(smokeintensity * smokeintensity) + smokeyrs
 + I(smokeyrs * smokeyrs) + as.factor(exercise)
 + as.factor(active) + wt71 + I(wt71 * wt71) + I(qsmk * smokeintensity),
  data = onesample
summary(std)
```

```
##
## Call:
## glm(formula = wt82_71 ~ qsmk + sex + race + age + I(age * age) +
## as.factor(education) + smokeintensity + I(smokeintensity *
## smokeintensity) + smokeyrs + I(smokeyrs * smokeyrs) + as.factor(exercise) +
## as.factor(active) + wt71 + I(wt71 * wt71) + I(qsmk * smokeintensity),
## data = onesample)
##
```

```
## Deviance Residuals:
##
      Min
                1Q Median
                                         Max
                    -0.343
##
  -42.056
           -4.171
                               3.891
                                       44.606
##
## Coefficients:
                                      Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                                     -1.5881657 4.3130359 -0.368 0.712756
                                      2.5595941 0.8091486
                                                            3.163 0.001590 **
## qsmk
## sex
                                     -1.4302717   0.4689576   -3.050   0.002328 **
## race
                                      ## age
                                      0.3596353 0.1633188
                                                            2.202 0.027809 *
                                     -0.0061010 0.0017261 -3.534 0.000421 ***
## I(age * age)
## as.factor(education)2
                                      0.7904440 0.6070005 1.302 0.193038
## as.factor(education)3
                                      0.5563124 0.5561016 1.000 0.317284
                                      1.4915695 0.8322704 1.792 0.073301 .
## as.factor(education)4
## as.factor(education)5
                                     -0.1949770 0.7413692 -0.263 0.792589
                                      0.0491365 0.0517254
                                                           0.950 0.342287
## smokeintensity
## I(smokeintensity * smokeintensity) -0.0009907 0.0009380 -1.056 0.291097
                                      0.1343686 0.0917122
                                                           1.465 0.143094
## smokeyrs
## I(smokeyrs * smokeyrs)
                                     -0.0018664 0.0015437 -1.209 0.226830
## as.factor(exercise)1
                                     0.2959754 0.5351533 0.553 0.580298
## as.factor(exercise)2
                                     0.3539128 0.5588587
                                                            0.633 0.526646
## as.factor(active)1
                                    -0.9475695 0.4099344 -2.312 0.020935 *
## as.factor(active)2
                                    -0.2613779   0.6845577   -0.382   0.702647
                                                           0.546 0.585299
## wt.71
                                     0.0455018 0.0833709
## I(wt71 * wt71)
                                     -0.0009653 0.0005247 -1.840 0.066001 .
## I(qsmk * smokeintensity)
                                     0.0466628 0.0351448
                                                          1.328 0.184463
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for gaussian family taken to be 53.5683)
##
##
      Null deviance: 97176 on 1565 degrees of freedom
## Residual deviance: 82763 on 1545 degrees of freedom
     (3321 observations deleted due to missingness)
## AIC: 10701
##
## Number of Fisher Scoring iterations: 2
onesample$predicted_meanY <- predict(std, onesample)</pre>
# estimate mean outcome in each of the groups interv=0, and interv=1
# this mean outcome is a weighted average of the mean outcomes in each combination
# of values of treatment and confounders, that is, the standardized outcome
mean(onesample[which(onesample$interv == -1), ]$predicted_meanY)
```

## [1] 2.56319

```
mean(onesample[which(onesample$interv == 0), ]$predicted_meanY)

## [1] 1.660267

mean(onesample[which(onesample$interv == 1), ]$predicted_meanY)

## [1] 5.178841
```

- Computing the 95% confidence interval of the standardized means and their difference
- Data from NHEFS

```
#install.packages("boot") # install package if required
library(boot)
# function to calculate difference in means
standardization <- function(data, indices) {</pre>
  # create a dataset with 3 copies of each subject
 d <- data[indices, ] # 1st copy: equal to original one`</pre>
  d$interv <- -1
  d0 <- d # 2nd copy: treatment set to 0, outcome to missing
  dO$interv <- 0
 d0$qsmk <- 0
 d0$wt82_71 <- NA
  d1 <- d # 3rd copy: treatment set to 1, outcome to missing
  d1$interv <- 1
  d1$qsmk <- 1
  d1$wt82_71 <- NA
  d.onesample <- rbind(d, d0, d1) # combining datasets
  # linear model to estimate mean outcome conditional on treatment and confounders
  # parameters are estimated using original observations only (interv= -1)
  # parameter estimates are used to predict mean outcome for observations with set
  # treatment (interv=0 and interv=1)
  fit <- glm(</pre>
    wt82_71 ~ qsmk + sex + race + age + I(age * age) +
      as.factor(education) + smokeintensity +
      I(smokeintensity * smokeintensity) + smokeyrs + I(smokeyrs *
      as.factor(exercise) + as.factor(active) + wt71 + I(wt71 *
                                                             wt71),
    data = d.onesample
  d.onesample$predicted_meanY <- predict(fit, d.onesample)</pre>
```

```
# estimate mean outcome in each of the groups interv=-1, interv=0, and interv=1
  return(c(
    mean(d.onesample$predicted_meanY[d.onesample$interv == -1]),
    mean(d.onesample$predicted_meanY[d.onesample$interv == 0]),
    mean(d.onesample$predicted_meanY[d.onesample$interv == 1]),
    mean(d.onesample$predicted_meanY[d.onesample$interv == 1]) -
      mean(d.onesample$predicted_meanY[d.onesample$interv == 0])
 ))
}
# bootstrap
results <- boot(data = nhefs,
                statistic = standardization,
                R = 5)
# generating confidence intervals
se <- c(sd(results$t[, 1]),</pre>
        sd(results$t[, 2]),
        sd(results$t[, 3]),
        sd(results$t[, 4]))
mean <- results$t0</pre>
11 < -mean - qnorm(0.975) * se
ul <- mean + qnorm(0.975) * se
bootstrap <-
  data.frame(cbind(
    c(
      "Observed",
      "No Treatment",
      "Treatment",
      "Treatment - No Treatment"
    ),
    mean,
    se,
    11.
    ul
  ))
bootstrap
```

```
##
                           V1
                                                                                 11
                                          mean
## 1
                     Observed 2.56188497106103
                                                 0.11833271542945 2.32995711062648
## 2
                 No Treatment 1.65212306626746 0.149274183026408 1.35955104371406
                    Treatment 5.11474489549347 0.152007614290188 4.81681544610884
## 4 Treatment - No Treatment 3.46262182922601 0.0778256612844246 3.31008633603552
##
                   ul
## 1 2.79381283149558
## 2 1.94469508882086
## 3 5.41267434487809
```

#### ## 4 3.61515732241649

# 14. G-estimation of Structural Nested Models

- Preprocessing, ranks of extreme observations, IP weights for censoring
- Data from NHEFS

```
library(here)
```

```
#install.packages("readxl") # install package if required
library("readxl")
nhefs <- read_excel(here("data", "NHEFS.xls"))

# some processing of the data
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)

# ranking of extreme observations
#install.packages("Hmisc")
library(Hmisc)</pre>
```

```
## Loading required package: lattice

## Loading required package: survival

## Loading required package: Formula

## Loading required package: ggplot2

## ## Attaching package: 'Hmisc'

## The following objects are masked from 'package:base': ##

## format.pval, units
```

```
describe(nhefs$wt82_71)
## nhefs$wt82_71
##
         n missing distinct
                                Info
                                         Mean
                                                   Gmd
                                                            .05
                                                                    .10
                 63
                                        2.638
                                                 8.337
                                                         -9.752
                                                                  -6.292
##
      1566
                        1510
                                   1
##
       .25
                         .75
                                          .95
                .50
                                  .90
##
    -1.478
              2.604
                       6.690
                              11.117
                                       14.739
##
## lowest : -41.28047 -30.50192 -30.05007 -29.02579 -25.97056
## highest: 34.01780 36.96925 37.65051 47.51130 48.53839
# estimation of denominator of ip weights for C
cw.denom <- glm(cens==0 ~ qsmk + sex + race + age + I(age^2)</pre>
                    + as.factor(education) + smokeintensity + I(smokeintensity^2)
                    + smokeyrs + I(smokeyrs^2) + as.factor(exercise)
                    + as.factor(active) + wt71 + I(wt71^2),
                    data = nhefs, family = binomial("logit"))
summary(cw.denom)
##
## Call:
## glm(formula = cens == 0 \sim qsmk + sex + race + age + I(age^2) +
      as.factor(education) + smokeintensity + I(smokeintensity^2) +
##
      smokeyrs + I(smokeyrs^2) + as.factor(exercise) + as.factor(active) +
      wt71 + I(wt71^2), family = binomial("logit"), data = nhefs)
##
##
## Deviance Residuals:
##
      Min
                1Q
                     Median
                                 3Q
                                         Max
## -2.9959
            0.1571
                     0.2069
                              0.2868
                                      1.0967
##
## Coefficients:
##
                          Estimate Std. Error z value Pr(>|z|)
                        -4.0144661 2.5761058 -1.558 0.11915
## (Intercept)
## qsmk
                        -0.5168674  0.2877162  -1.796  0.07242 .
## sex
                        ## race
                         0.0122715 0.4524887
                                               0.027 0.97836
## age
                         0.2697293 0.1174647
                                               2.296 0.02166 *
                        ## I(age^2)
## as.factor(education)2 0.4407884 0.4193993
                                               1.051 0.29326
## as.factor(education)3 0.1646881 0.3705471
                                               0.444 0.65672
## as.factor(education)4 -0.1384470 0.5697969 -0.243 0.80802
## as.factor(education)5 0.3823818 0.5601808
                                               0.683 0.49486
                        -0.0157119 0.0347319 -0.452 0.65100
## smokeintensity
## I(smokeintensity^2)
                                              0.187 0.85171
                         0.0001133 0.0006058
                        -0.0785973 0.0749576 -1.049 0.29438
## smokeyrs
## I(smokeyrs^2)
                         0.0005569 0.0010318 0.540 0.58938
## as.factor(exercise)1
                         0.9714714 0.3878101
                                               2.505 0.01224 *
## as.factor(exercise)2  0.5839890  0.3723133  1.569  0.11675
```

```
## as.factor(active)1
                       0.2474785 0.3254548
                                               0.760 0.44701
## as.factor(active)2 -0.7065829 0.3964577 -1.782 0.07471 .
## wt71
                        0.0878871 0.0400115
                                               2.197 0.02805 *
## I(wt71^2)
                       -0.0006351 0.0002257 -2.813 0.00490 **
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 533.36 on 1628 degrees of freedom
##
## Residual deviance: 465.36 on 1609 degrees of freedom
## AIC: 505.36
##
## Number of Fisher Scoring iterations: 7
nhefs$pd.c <- predict(cw.denom, nhefs, type="response")</pre>
nhefs$wc <- ifelse(nhefs$cens==0, 1/nhefs$pd.c, NA)
# observations with cens=1 only contribute to censoring models
```

# Program 14.2

- G-estimation of a 1-parameter structural nested mean model
- Brute force search
- Data from NHEFS

#### G-estimation: Checking one possible value of psi

```
## geeglm(formula = qsmk ~ sex + race + age + I(age * age) + as.factor(education) +
      smokeintensity + I(smokeintensity * smokeintensity) + smokeyrs +
##
      I(smokeyrs * smokeyrs) + as.factor(exercise) + as.factor(active) +
##
      wt71 + I(wt71 * wt71) + Hpsi, family = binomial, data = nhefs,
##
      weights = wc, id = seqn, corstr = "independence")
##
##
##
   Coefficients:
##
                                       Estimate
                                                   Std.err
                                                             Wald Pr(>|W|)
## (Intercept)
                                     -2.403e+00 1.329e+00 3.269 0.070604 .
## sex
                                     -5.137e-01 1.536e-01 11.193 0.000821 ***
                                     -8.609e-01 2.099e-01 16.826 4.10e-05 ***
## race
                                      1.152e-01 5.020e-02 5.263 0.021779 *
## age
                                     -7.593e-04 5.296e-04 2.056 0.151619
## I(age * age)
## as.factor(education)2
                                     -2.894e-02 1.964e-01 0.022 0.882859
## as.factor(education)3
                                      8.771e-02 1.726e-01 0.258 0.611329
## as.factor(education)4
                                      6.637e-02 2.698e-01 0.061 0.805645
                                      4.711e-01 2.247e-01 4.395 0.036036 *
## as.factor(education)5
                                     -7.834e-02 1.464e-02 28.635 8.74e-08 ***
## smokeintensity
## I(smokeintensity * smokeintensity) 1.072e-03 2.650e-04 16.368 5.21e-05 ***
## smokeyrs
                                     -7.111e-02 2.639e-02 7.261 0.007047 **
## I(smokeyrs * smokeyrs)
                                      8.153e-04 4.490e-04 3.298 0.069384 .
## as.factor(exercise)1
                                      3.363e-01 1.828e-01 3.384 0.065844 .
## as.factor(exercise)2
                                      3.800e-01 1.889e-01 4.049 0.044187 *
## as.factor(active)1
                                      3.412e-02 1.339e-01 0.065 0.798778
## as.factor(active)2
                                      2.135e-01 2.121e-01 1.012 0.314308
## wt71
                                     -7.661e-03 2.562e-02 0.089 0.764963
## I(wt71 * wt71)
                                      8.655e-05 1.582e-04 0.299 0.584233
## Hpsi
                                     -1.903e-06 8.839e-03 0.000 0.999828
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Correlation structure = independence
## Estimated Scale Parameters:
##
              Estimate Std.err
## (Intercept)
                0.9969 0.06717
## Number of clusters: 1566 Maximum cluster size: 1
```

#### G-estimation: Checking multiple possible values of psi

```
#install.packages("geepack")
grid <- seq(from = 2,to = 5, by = 0.1)
j = 0
Hpsi.coefs <- cbind(rep(NA,length(grid)), rep(NA, length(grid)))
colnames(Hpsi.coefs) <- c("Estimate", "p-value")

for (i in grid){
   psi = i</pre>
```

## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm! ## Warning in eval(family\$initialize): non-integer #successes in a binomial glm!

```
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
```

Estimate p-value [1,] 0.0267219 0.001772 [2,] 0.0248946 0.003580 [3,] 0.0230655 0.006963 [4,]0.0212344 0.013026 [5,] 0.0194009 0.023417 [6,] 0.0175647 0.040430 [7,] 0.0157254 0.067015 [8,] 0.0138827 0.106626 [9,] 0.0120362 0.162877 ## [10,] 0.0101857 0.238979 **##** [11,] 0.0083308 0.337048 ## [12,] 0.0064713 0.457433 **##** [13,] 0.0046069 0.598235 ## [14,] 0.0027374 0.755204 ## [15,] 0.0008624 0.922101 ## [16,] -0.0010181 0.908537 ## [17,] -0.0029044 0.744362 ## [18,] -0.0047967 0.592188 ## [19,] -0.0066950 0.457169 ## [20,] -0.0085997 0.342360 ## [21,] -0.0105107 0.248681

Hpsi.coefs

```
## [22,] -0.0124282 0.175239

## [23,] -0.0143523 0.119841

## [24,] -0.0162831 0.079580

## [25,] -0.0182206 0.051347

## [26,] -0.0201649 0.032218

## [27,] -0.0221160 0.019675

## [28,] -0.0240740 0.011706

## [29,] -0.0260389 0.006792

## [30,] -0.0280106 0.003847

## [31,] -0.0299893 0.002129
```

#### Program 14.3

- G-estimation for 2-parameter structural nested mean model
- Closed form estimator
- Data from NHEFS

#### G-estimation: Closed form estimator linear mean models

## Warning in eval(family\$initialize): non-integer #successes in a binomial glm!

```
summary(logit.est)
```

```
##
## Call:
## glm(formula = qsmk ~ sex + race + age + I(age^2) + as.factor(education) +
##
      smokeintensity + I(smokeintensity^2) + smokeyrs + I(smokeyrs^2) +
      as.factor(exercise) + as.factor(active) + wt71 + I(wt71^2),
##
      family = binomial(), data = nhefs, weights = wc)
##
##
## Deviance Residuals:
     Min
              1Q Median
                              3Q
                                     Max
## -1.529 -0.808 -0.650 1.029
                                   2.417
##
## Coefficients:
                         Estimate Std. Error z value Pr(>|z|)
                        -2.40e+00
## (Intercept)
                                   1.31e+00 -1.83 0.06743 .
                        -5.14e-01 1.50e-01 -3.42 0.00062 ***
## sex
                        -8.61e-01 2.06e-01 -4.18 2.9e-05 ***
## race
```

```
## age
                          1.15e-01
                                     4.95e-02
                                                 2.33 0.01992 *
## I(age^2)
                         -7.59e-04
                                     5.14e-04
                                                -1.48
                                                      0.13953
## as.factor(education)2 -2.89e-02
                                     1.93e-01
                                                -0.15
                                                       0.88079
## as.factor(education)3 8.77e-02
                                     1.73e-01
                                                 0.51
                                                       0.61244
## as.factor(education)4 6.64e-02
                                     2.66e-01
                                                 0.25
                                                       0.80301
## as.factor(education)5 4.71e-01
                                     2.21e-01
                                                 2.13 0.03314 *
## smokeintensity
                         -7.83e-02
                                     1.49e-02
                                                -5.27
                                                       1.4e-07 ***
## I(smokeintensity^2)
                                                 3.85 0.00012 ***
                          1.07e-03
                                     2.78e-04
## smokeyrs
                                     2.71e-02
                                                -2.63 0.00862 **
                         -7.11e-02
## I(smokeyrs^2)
                          8.15e-04
                                     4.45e-04
                                                 1.83 0.06722 .
## as.factor(exercise)1
                          3.36e-01
                                     1.75e-01
                                                 1.92 0.05467
## as.factor(exercise)2
                          3.80e-01
                                     1.82e-01
                                                 2.09 0.03637 *
## as.factor(active)1
                          3.41e-02
                                     1.30e-01
                                                 0.26 0.79337
## as.factor(active)2
                          2.13e-01
                                     2.06e-01
                                                 1.04 0.30033
## wt71
                         -7.66e-03
                                     2.46e-02
                                                -0.31 0.75530
## I(wt71^2)
                          8.66e-05
                                     1.51e-04
                                                 0.57 0.56586
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
##
  (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 1872.2 on 1565 degrees of freedom
## Residual deviance: 1755.6 on 1547 degrees of freedom
     (63 observations deleted due to missingness)
## AIC: 1719
##
## Number of Fisher Scoring iterations: 4
nhefs$pqsmk <- predict(logit.est, nhefs, type = "response")</pre>
describe(nhefs$pqsmk)
## nhefs$pqsmk
                                                               .05
                                                                        .10
##
          n missing distinct
                                  Info
                                           Mean
                                                     Gmd
##
       1629
                   0
                         1629
                                     1
                                         0.2622
                                                  0.1302
                                                           0.1015
                                                                     0.1261
##
        .25
                 .50
                          .75
                                   .90
                                            .95
##
     0.1780
              0.2426
                       0.3251
                                0.4221
                                         0.4965
## lowest : 0.05145 0.05157 0.05438 0.05583 0.05931
## highest: 0.67208 0.68643 0.71391 0.73330 0.78914
summary(nhefs$pqsmk)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                              Max.
   0.0514 0.1780 0.2426 0.2622 0.3251 0.7891
# solve sum(w_c * H(psi) * (qsmk - E[qsmk | L])) = 0
# for a single psi and H(psi) = wt82_71 - psi * qsmk
# this can be solved as psi = sum(w_c * wt82_71 * (qsmk - pqsmk)) / sum(w_c * qsmk * (qsmk - pqsmk))
```

```
nhefs.c <- nhefs[which(!is.na(nhefs$wt82)),]
with(nhefs.c, sum(wc*wt82_71*(qsmk-pqsmk)) / sum(wc*qsmk*(qsmk - pqsmk)))
## [1] 3.446</pre>
```

#### G-estimation: Closed form estimator for 2-parameter model

```
diff = with(nhefs.c, qsmk - pqsmk)
diff2 = with(nhefs.c, wc * diff)

lhs = matrix(0,2,2)
lhs[1,1] = with(nhefs.c, sum(qsmk * diff2))
lhs[1,2] = with(nhefs.c, sum(qsmk * smokeintensity * diff2))
lhs[2,1] = with(nhefs.c, sum(qsmk * smokeintensity * diff2))
lhs[2,2] = with(nhefs.c, sum(qsmk * smokeintensity * smokeintensity * diff2))

rhs = matrix(0,2,1)
rhs[1] = with(nhefs.c, sum(wt82_71 * diff2))
rhs[2] = with(nhefs.c, sum(wt82_71 * smokeintensity * diff2))

psi = t(solve(lhs,rhs))
psi
```

```
## [,1] [,2]
## [1,] 2.859 0.03004
```

# 15. Outcome regression and propensity scores

# Program 15.1

- Estimating the average causal effect within levels of confounders under the assumption of effect-measure modification by smoking intensity ONLY
- Data from NHEFS

#### library(here)

```
##
## Call:
## glm(formula = wt82_71 \sim qsmk + sex + race + age + I(age * age) +
       as.factor(education) + smokeintensity + I(smokeintensity *
       smokeintensity) + smokeyrs + I(smokeyrs * smokeyrs) + as.factor(exercise) +
       as.factor(active) + wt71 + I(wt71 * wt71) + I(qsmk * smokeintensity),
##
##
       data = nhefs)
## Deviance Residuals:
                1Q Median
      Min
                                  3Q
                                           Max
## -42.056 -4.171 -0.343 3.891
                                       44.606
##
## Coefficients:
```

```
##
                                     Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                   -1.5881657 4.3130359 -0.368 0.712756
                                    2.5595941 0.8091486 3.163 0.001590 **
## qsmk
## sex
                                   -1.4302717   0.4689576   -3.050   0.002328 **
## race
                                    0.5601096  0.5818888  0.963  0.335913
## age
                                    0.3596353  0.1633188  2.202  0.027809 *
                                   -0.0061010 0.0017261 -3.534 0.000421 ***
## I(age * age)
## as.factor(education)2
                                    0.7904440 0.6070005 1.302 0.193038
                                    0.5563124  0.5561016  1.000  0.317284
## as.factor(education)3
## as.factor(education)4
                                   1.4915695 0.8322704 1.792 0.073301 .
                                   -0.1949770 0.7413692 -0.263 0.792589
## as.factor(education)5
## smokeintensity
                                    0.0491365 0.0517254 0.950 0.342287
## I(smokeintensity * smokeintensity) -0.0009907 0.0009380 -1.056 0.291097
## smokeyrs
                                    0.1343686 0.0917122 1.465 0.143094
                                   -0.0018664 0.0015437 -1.209 0.226830
## I(smokeyrs * smokeyrs)
## as.factor(exercise)1
                                  0.2959754 0.5351533 0.553 0.580298
## as.factor(exercise)2
                                   ## as.factor(active)1
                                   -0.9475695   0.4099344   -2.312   0.020935 *
                                   ## as.factor(active)2
## wt71
                                    0.0455018 0.0833709 0.546 0.585299
## I(wt71 * wt71)
                                   -0.0009653 0.0005247 -1.840 0.066001 .
## I(qsmk * smokeintensity)
                                    ## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for gaussian family taken to be 53.5683)
##
##
      Null deviance: 97176 on 1565 degrees of freedom
## Residual deviance: 82763 on 1545 degrees of freedom
    (63 observations deleted due to missingness)
## AIC: 10701
##
## Number of Fisher Scoring iterations: 2
# (step 1) build the contrast matrix with all zeros
# this function builds the blank matrix
# install.packages("multcomp") # install packages if necessary
library("multcomp")
## Loading required package: mvtnorm
## Loading required package: survival
## Loading required package: TH.data
## Loading required package: MASS
##
## Attaching package: 'TH.data'
```

```
## The following object is masked from 'package:MASS':
##
##
       geyser
makeContrastMatrix <- function(model, nrow, names) {</pre>
  m <- matrix(0, nrow = nrow, ncol = length(coef(model)))</pre>
  colnames(m) <- names(coef(model))</pre>
  rownames(m) <- names</pre>
  return(m)
}
K1 <- makeContrastMatrix(fit, 2, c('Effect of Quitting Smoking at Smokeintensity of 5',
                                       'Effect of Quitting Smoking at Smokeintensity of 40'))
# (step 2) fill in the relevant non-zero elements
K1[1:2, 'qsmk'] <- 1
K1[1:2, 'I(qsmk * smokeintensity)'] <- c(5, 40)</pre>
# (step 3) check the contrast matrix
                                                        (Intercept) qsmk sex race
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                  0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                                 0
                                                        age I(age * age)
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                        as.factor(education)2
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                        as.factor(education)3
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                             0
## Effect of Quitting Smoking at Smokeintensity of 40
##
                                                        as.factor(education)4
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                             0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                             0
                                                        as.factor(education)5
##
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                             0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                             0
##
                                                        smokeintensity
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                     0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                     0
                                                        I(smokeintensity * smokeintensity)
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                                          0
                                                                                          0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                        smokeyrs
## Effect of Quitting Smoking at Smokeintensity of 5
                                                               0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                               0
                                                        I(smokeyrs * smokeyrs)
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                              0
```

## Effect of Quitting Smoking at Smokeintensity of 40

0

```
##
                                                       as.factor(exercise)1
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                          0
                                                       as.factor(exercise)2
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                          0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                       as.factor(active)1
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                        0
                                                                        0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                       as.factor(active)2 wt71
##
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                        0
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                        0
                                                                             0
##
                                                       I(wt71 * wt71)
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                    0
##
                                                       I(qsmk * smokeintensity)
## Effect of Quitting Smoking at Smokeintensity of 5
                                                                              5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                             40
# (step 4) estimate the contrasts, get tests and confidence intervals for them
estimates1 <- glht(fit, K1)
  summary(estimates1)
##
##
     Simultaneous Tests for General Linear Hypotheses
##
## Fit: glm(formula = wt82_71 ~ qsmk + sex + race + age + I(age * age) +
       as.factor(education) + smokeintensity + I(smokeintensity *
##
       smokeintensity) + smokeyrs + I(smokeyrs * smokeyrs) + as.factor(exercise) +
##
       as.factor(active) + wt71 + I(wt71 * wt71) + I(qsmk * smokeintensity),
##
       data = nhefs)
##
##
## Linear Hypotheses:
                                                            Estimate Std. Error
                                                              2.7929
## Effect of Quitting Smoking at Smokeintensity of 5 == 0
                                                                         0.6683
## Effect of Quitting Smoking at Smokeintensity of 40 == 0
                                                              4.4261
                                                                         0.8478
                                                           z value Pr(>|z|)
## Effect of Quitting Smoking at Smokeintensity of 5 == 0
                                                              4.179 5.84e-05 ***
## Effect of Quitting Smoking at Smokeintensity of 40 == 0
                                                              5.221 3.56e-07 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
 confint(estimates1)
##
     Simultaneous Confidence Intervals
##
##
```

```
## Fit: glm(formula = wt82_71 ~ qsmk + sex + race + age + I(age * age) +
##
      as.factor(education) + smokeintensity + I(smokeintensity *
      smokeintensity) + smokeyrs + I(smokeyrs * smokeyrs) + as.factor(exercise) +
##
      as.factor(active) + wt71 + I(wt71 * wt71) + I(qsmk * smokeintensity),
##
##
      data = nhefs)
##
## Quantile = 2.2281
## 95% family-wise confidence level
##
##
## Linear Hypotheses:
                                                     Estimate lwr
## Effect of Quitting Smoking at Smokeintensity of 5 == 0 2.7929
                                                             1.3039 4.2819
## Effect of Quitting Smoking at Smokeintensity of 40 == 0 4.4261
                                                             2.5372 6.3151
# regression on covariates, not allowing for effect modification
fit2 <- glm(wt82_71 ~ qsmk + sex + race + age + I(age*age) + as.factor(education)
         + smokeintensity + I(smokeintensity*smokeintensity) + smokeyrs
         + I(smokeyrs*smokeyrs) + as.factor(exercise) + as.factor(active)
         + wt71 + I(wt71*wt71), data=nhefs)
summary(fit2)
##
## Call:
## glm(formula = wt82_71 ~ qsmk + sex + race + age + I(age * age) +
      as.factor(education) + smokeintensity + I(smokeintensity *
      smokeintensity) + smokeyrs + I(smokeyrs * smokeyrs) + as.factor(exercise) +
##
##
      as.factor(active) + wt71 + I(wt71 * wt71), data = nhefs)
##
## Deviance Residuals:
##
      Min
               10 Median
                               30
                                      Max
          -4.216 -0.318 3.807
## -42.332
                                    44.668
##
## Coefficients:
##
                                    Estimate Std. Error t value Pr(>|t|)
                                  -1.6586176 4.3137734 -0.384 0.700666
## (Intercept)
                                   ## qsmk
                                  -1.4650496 0.4683410 -3.128 0.001792 **
## sex
                                   ## race
                                   ## age
## I(age * age)
                                  ## as.factor(education)2
## as.factor(education)3
                                   0.5715004 0.5561211 1.028 0.304273
## as.factor(education)4
                                   1.5085173 0.8323778 1.812 0.070134 .
## as.factor(education)5
                                  -0.1708264 0.7413289 -0.230 0.817786
                                   0.0651533 0.0503115 1.295 0.195514
## smokeintensity
## I(smokeintensity * smokeintensity) -0.0010468 0.0009373 -1.117 0.264261
                                   0.1333931 0.0917319 1.454 0.146104
## smokeyrs
```

```
## I(smokeyrs * smokeyrs)
                                     -0.0018270 0.0015438 -1.183 0.236818
## as.factor(exercise)1
                                      0.3206824 0.5349616
                                                             0.599 0.548961
## as.factor(exercise)2
                                      0.3628786 0.5589557
                                                             0.649 0.516300
## as.factor(active)1
                                     -0.9429574   0.4100208   -2.300   0.021593 *
## as.factor(active)2
                                                            -0.377 0.706337
                                     -0.2580374 0.6847219
## wt71
                                      0.0373642 0.0831658
                                                             0.449 0.653297
## I(wt71 * wt71)
                                     -0.0009158 0.0005235
                                                            -1.749 0.080426 .
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
   (Dispersion parameter for gaussian family taken to be 53.59474)
##
##
##
       Null deviance: 97176 on 1565 degrees of freedom
## Residual deviance: 82857 on 1546 degrees of freedom
     (63 observations deleted due to missingness)
## AIC: 10701
##
## Number of Fisher Scoring iterations: 2
```

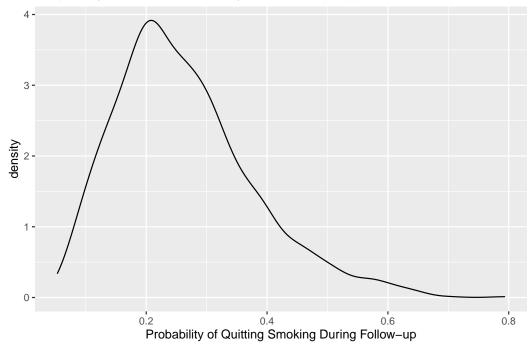
- Estimating and plotting the propensity score
- Data from NHEFS

```
##
## Call:
  glm(formula = qsmk ~ sex + race + age + I(age * age) + as.factor(education) +
      smokeintensity + I(smokeintensity * smokeintensity) + smokeyrs +
      I(smokeyrs * smokeyrs) + as.factor(exercise) + as.factor(active) +
##
##
      wt71 + I(wt71 * wt71), family = binomial(), data = nhefs)
##
## Deviance Residuals:
      Min
                   Median
              1Q
## -1.4646 -0.8044 -0.6460
                         1.0578
                                   2.3550
##
## Coefficients:
##
                                  Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                                 -1.9889022 1.2412792 -1.602 0.109089
## sex
                                 ## race
                                 ## age
                                  0.1030132 0.0488996
                                                     2.107 0.035150 *
```

```
## I(age * age)
                                  -0.0006052 0.0005074 -1.193 0.232973
## as.factor(education)2
                                  ## as.factor(education)3
                                   0.0156987 0.1707139
                                                       0.092 0.926730
## as.factor(education)4
                                  -0.0425260 0.2642761 -0.161 0.872160
## as.factor(education)5
                                   0.3796632 0.2203947
                                                       1.723 0.084952 .
## smokeintensity
                                  -0.0651561  0.0147589  -4.415  1.01e-05 ***
## I(smokeintensity * smokeintensity) 0.0008461 0.0002758 3.067 0.002160 **
## smokeyrs
                                  0.0008384 0.0004435 1.891 0.058669 .
## I(smokeyrs * smokeyrs)
## as.factor(exercise)1
                                   0.2914117  0.1735543  1.679  0.093136 .
## as.factor(exercise)2
                                   0.3550517 0.1799293 1.973 0.048463 *
## as.factor(active)1
                                   0.0108754 0.1298320 0.084 0.933243
## as.factor(active)2
                                   ## wt71
                                  ## I(wt71 * wt71)
                                   0.0001209 0.0001352
                                                        0.895 0.370957
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 1876.3 on 1628 degrees of freedom
##
## Residual deviance: 1766.7 on 1610 degrees of freedom
## AIC: 1804.7
##
## Number of Fisher Scoring iterations: 4
nhefs$ps <- predict(fit3, nhefs, type="response")</pre>
summary(nhefs$ps[nhefs$qsmk==0])
     Min. 1st Qu. Median
                           Mean 3rd Qu.
## 0.05298 0.16949 0.22747 0.24504 0.30441 0.65788
summary(nhefs$ps[nhefs$qsmk==1])
     Min. 1st Qu. Median
                           Mean 3rd Qu.
## 0.06248 0.22046 0.28897 0.31240 0.38122 0.79320
# # plotting the estimated propensity score
# install.packages("ggplot2") # install packages if necessary
# install.packages("dplyr")
library("ggplot2")
library("dplyr")
## Attaching package: 'dplyr'
## The following object is masked from 'package:MASS':
```

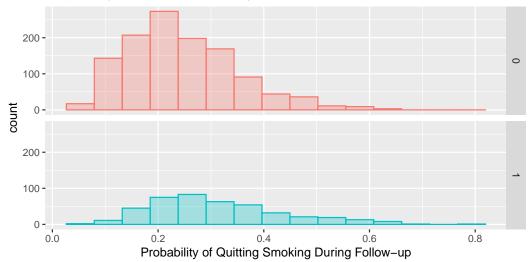
```
##
##
       select
## The following objects are masked from 'package:stats':
##
       filter, lag
##
## The following objects are masked from 'package:base':
##
##
       intersect, setdiff, setequal, union
ggplot(nhefs, aes(x = ps, fill = qsmk)) + geom_density(alpha = 0.2) +
  xlab('Probability of Quitting Smoking During Follow-up') +
  ggtitle('Propensity Score Distribution by Treatment Group') +
  scale_fill_discrete('') +
  theme(legend.position = 'bottom', legend.direction = 'vertical')
```

#### Propensity Score Distribution by Treatment Group



```
scale_fill_discrete('') +
scale_color_discrete('') +
theme(legend.position = 'bottom', legend.direction = 'vertical')
```

#### Propensity Score Distribution by Treatment Group





```
# attempt to reproduce plot from the book
nhefs %>%
  mutate(ps.grp = round(ps/0.05) * 0.05) %
  group_by(qsmk, ps.grp) %>%
  summarize(n = n()) \%
  ungroup() %>%
  mutate(n2 = ifelse(qsmk == 0, yes = n, no = -1*n)) \%
  ggplot(aes(x = ps.grp, y = n2, fill = as.factor(qsmk))) +
  geom_bar(stat = 'identity', position = 'identity') +
  geom_text(aes(label = n, x = ps.grp, y = n2 + ifelse(qsmk == 0, 8, -8))) +
  xlab('Probability of Quitting Smoking During Follow-up') +
  ylab('N') +
  ggtitle('Propensity Score Distribution by Treatment Group') +
  scale_fill_discrete('') +
  scale x continuous(breaks = seq(0, 1, 0.05)) +
  theme(legend.position = 'bottom', legend.direction = 'vertical',
        axis.ticks.y = element_blank(),
       axis.text.y = element_blank())
```

- Stratification on the propensity score
- Data from NHEFS

```
# calculation of deciles
nhefs$ps.dec <- cut(nhefs$ps,</pre>
               breaks=c(quantile(nhefs$ps, probs=seq(0,1,0.1))),
               labels=seq(1:10),
               include.lowest=TRUE)
#install.packages("psych") # install package if required
library("psych")
## Attaching package: 'psych'
## The following objects are masked from 'package:ggplot2':
##
     %+%, alpha
describeBy(nhefs$ps, list(nhefs$ps.dec, nhefs$qsmk))
## Descriptive statistics by group
## : 1
## : 0
  vars n mean sd median trimmed mad min max range skew kurtosis se
## : 2
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
     1 136 0.15 0.01 0.15 0.15 0.01 0.13 0.17 0.04 -0.04 -1.23 0
## -----
## : 3
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 134 0.18 0.01 0.18 0.18 0.01 0.17 0.19 0.03 -0.08 -1.34 0
## -----
## : 4
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 129 0.21 0.01 0.21 0.01 0.19 0.22 0.02 -0.04 -1.13 0
## : 5
## : 0
  vars n mean sd median trimmed mad min max range skew kurtosis se
```

```
1 120 0.23 0.01 0.23 0.23 0.01 0.22 0.25 0.03 0.24
## -----
## : 6
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 117 0.26 0.01 0.26 0.26 0.01 0.25 0.27 0.03 -0.11 -1.29 0
## -----
## : 7
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 120 0.29 0.01 0.29 0.29 0.01 0.27 0.31 0.03 -0.23 -1.19 0
## -----
## : 8
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 112 0.33 0.01 0.33 0.33 0.02 0.31 0.35 0.04 0.15 -1.1 0
## -----
## : 9
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 96 0.38 0.02 0.38 0.38 0.02 0.35 0.42 0.06 0.13 -1.15 0
## -----
## : 10
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 86 0.49 0.06 0.47 0.48 0.05 0.42 0.66 0.24 1.1 0.47 0.01
## -----
## : 1
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis
## X1 1 12 0.1 0.02 0.11 0.1 0.03 0.06 0.13 0.07 -0.5 -1.36 0.01
## -----
## : 2
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
## -----
## : 3
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 29 0.18 0.01 0.18 0.18 0.01 0.17 0.19 0.03 0.01 -1.34 0
## -----
## : 4
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
   1 34 0.21 0.01 0.21 0.21 0.01 0.19 0.22 0.02 -0.31 -1.23 0
## -----
## : 5
## : 1
```

```
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 43 0.23 0.01 0.23 0.23 0.01 0.22 0.25 0.03 0.11 -1.23 0
## : 6
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 45 0.26 0.01 0.26 0.01 0.25 0.27 0.03 0.2
## -----
## : 1
  vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 43 0.29 0.01 0.29 0.29 0.01 0.27 0.31 0.03 0.16 -1.25 0
## -----
## : 8
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 51 0.33 0.01 0.33 0.02 0.31 0.35 0.04 0.11
## : 9
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 67 0.38 0.02 0.38 0.38 0.03 0.35 0.42 0.06 0.19
## -----
## : 10
## : 1
  vars n mean sd median trimmed mad min max range skew kurtosis
# function to create deciles easily
decile <- function(x) {</pre>
 return(factor(quantcut(x, seq(0, 1, 0.1), labels = FALSE)))
}
# regression on PS deciles, allowing for effect modification
for (deciles in c(1:10)) {
 print(t.test(wt82_71~qsmk, data=nhefs[which(nhefs$ps.dec==deciles),]))
}
##
## Welch Two Sample t-test
## data: wt82_71 by qsmk
## t = 0.0060506, df = 11.571, p-value = 0.9953
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -5.283903 5.313210
## sample estimates:
## mean in group 0 mean in group 1
  3.995205
               3.980551
##
```

```
##
##
##
   Welch Two Sample t-test
##
## data: wt82_71 by qsmk
## t = -3.1117, df = 37.365, p-value = 0.003556
\mbox{\tt \#\#} alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -6.849335 -1.448161
## sample estimates:
## mean in group 0 mean in group 1
          2.904679
                          7.053426
##
##
##
  Welch Two Sample t-test
##
## data: wt82_71 by qsmk
## t = -4.5301, df = 35.79, p-value = 6.317e-05
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -9.474961 -3.613990
## sample estimates:
## mean in group 0 mean in group 1
          2.612094
                          9.156570
##
##
##
##
   Welch Two Sample t-test
##
## data: wt82_71 by qsmk
## t = -1.4117, df = 45.444, p-value = 0.1648
\#\# alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -5.6831731 0.9985715
## sample estimates:
## mean in group 0 mean in group 1
          3.474679
                          5.816979
##
##
##
  Welch Two Sample t-test
##
## data: wt82_71 by qsmk
## t = -3.1371, df = 74.249, p-value = 0.002446
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -6.753621 -1.507087
## sample estimates:
## mean in group 0 mean in group 1
##
          2.098800
                          6.229154
##
```

```
##
   Welch Two Sample t-test
## data: wt82_71 by qsmk
## t = -2.1677, df = 50.665, p-value = 0.0349
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -8.7516605 -0.3350127
## sample estimates:
## mean in group 0 mean in group 1
          1.847004
                          6.390340
##
##
##
##
  Welch Two Sample t-test
##
## data: wt82_71 by qsmk
## t = -3.3155, df = 84.724, p-value = 0.001348
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -6.904207 -1.727590
## sample estimates:
## mean in group 0 mean in group 1
          1.560048
                          5.875946
##
##
##
## Welch Two Sample t-test
##
## data: wt82_71 by qsmk
## t = -2.664, df = 75.306, p-value = 0.009441
\#\# alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -6.2396014 -0.9005605
## sample estimates:
## mean in group 0 mean in group 1
##
         0.2846851
                         3.8547661
##
##
## Welch Two Sample t-test
## data: wt82_71 by qsmk
## t = -1.9122, df = 129.12, p-value = 0.05806
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -4.68143608 0.07973698
## sample estimates:
## mean in group 0 mean in group 1
##
        -0.8954482
                        1.4054014
##
##
```

```
Welch Two Sample t-test
##
## data: wt82_71 by qsmk
## t = -1.5925, df = 142.72, p-value = 0.1135
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
   -5.0209284 0.5404697
## sample estimates:
## mean in group 0 mean in group 1
        -0.5043766
##
                         1.7358528
# regression on PS deciles, not allowing for effect modification
fit.psdec <- glm(wt82_71 ~ qsmk + as.factor(ps.dec), data = nhefs)
summary(fit.psdec)
##
## Call:
## glm(formula = wt82_71 ~ qsmk + as.factor(ps.dec), data = nhefs)
##
## Deviance Residuals:
                     Median
##
      Min
                 1Q
                                   3Q
                                          Max
## -43.543
            -3.932
                     -0.085
                                4.233
                                        46.773
##
## Coefficients:
##
                      Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                         3.7505
                                   0.6089
                                            6.159 9.29e-10 ***
                                            7.659 3.28e-14 ***
## qsmk
                        3.5005
                                   0.4571
## as.factor(ps.dec)2
                       -0.7391
                                   0.8611 -0.858
                                                    0.3908
## as.factor(ps.dec)3
                       -0.6182
                                   0.8612 -0.718
                                                    0.4730
## as.factor(ps.dec)4
                       -0.5204
                                   0.8584 -0.606
                                                    0.5444
## as.factor(ps.dec)5
                       -1.4884
                                   0.8590 -1.733
                                                     0.0834 .
## as.factor(ps.dec)6
                       -1.6227
                                   0.8675 -1.871
                                                     0.0616 .
## as.factor(ps.dec)7
                       -1.9853
                                   0.8681 -2.287
                                                     0.0223 *
## as.factor(ps.dec)8
                       -3.4447
                                   0.8749 -3.937 8.61e-05 ***
## as.factor(ps.dec)9
                        -5.1544
                                   0.8848 -5.825 6.91e-09 ***
## as.factor(ps.dec)10 -4.8403
                                   0.8828 -5.483 4.87e-08 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for gaussian family taken to be 58.42297)
##
##
       Null deviance: 97176 on 1565 degrees of freedom
## Residual deviance: 90848 on 1555 degrees of freedom
     (63 observations deleted due to missingness)
## AIC: 10827
##
## Number of Fisher Scoring iterations: 2
```

```
confint.lm(fit.psdec)
```

```
2.5 %
                                     97.5 %
##
## (Intercept)
                       2.556098 4.94486263
## qsmk
                       2.603953 4.39700504
## as.factor(ps.dec)2 -2.428074 0.94982494
## as.factor(ps.dec)3 -2.307454 1.07103569
## as.factor(ps.dec)4 -2.204103 1.16333143
## as.factor(ps.dec)5 -3.173337 0.19657938
## as.factor(ps.dec)6 -3.324345 0.07893027
## as.factor(ps.dec)7 -3.688043 -0.28248110
## as.factor(ps.dec)8 -5.160862 -1.72860113
## as.factor(ps.dec)9 -6.889923 -3.41883853
## as.factor(ps.dec)10 -6.571789 -3.10873731
```

- Standardization using the propensity score
- Data from NHEFS

```
#install.packages("boot") # install package if required
library("boot")
```

```
##
## Attaching package: 'boot'
## The following object is masked from 'package:psych':
##
##
       logit
## The following object is masked from 'package:survival':
##
##
       aml
# standardization by propensity score, agnostic regarding effect modification
std.ps <- function(data, indices) {</pre>
  d <- data[indices,] # 1st copy: equal to original one
  # calculating propensity scores
  ps.fit <- glm(qsmk ~ sex + race + age + I(age*age)</pre>
                + as.factor(education) + smokeintensity
                + I(smokeintensity*smokeintensity) + smokeyrs
                + I(smokeyrs*smokeyrs) + as.factor(exercise)
                + as.factor(active) + wt71 + I(wt71*wt71),
                data=d, family=binomial())
  d$pscore <- predict(ps.fit, d, type="response")</pre>
```

```
# create a dataset with 3 copies of each subject
  d$interv <- -1 # 1st copy: equal to original one
  d0 <- d # 2nd copy: treatment set to 0, outcome to missing
  dO$interv <- 0
  d0$qsmk <- 0
  d0$wt82 71 <- NA
  d1 <- d # 3rd copy: treatment set to 1, outcome to missing
  d1$interv <- 1
  d1$qsmk <- 1
  d1$wt82_71 <- NA
  d.onesample <- rbind(d, d0, d1) # combining datasets
  std.fit <- glm(wt82_71 ~ qsmk + pscore + I(qsmk*pscore), data=d.onesample)
  d.onesample$predicted_meanY <- predict(std.fit, d.onesample)</pre>
  # estimate mean outcome in each of the groups interv=-1, interv=0, and interv=1
  return(c(mean(d.onesample$predicted_meanY[d.onesample$interv==-1]),
           mean(d.onesample$predicted_meanY[d.onesample$interv==0]),
           mean(d.onesample$predicted_meanY[d.onesample$interv==1]),
           mean(d.onesample$predicted_meanY[d.onesample$interv==1])-
             mean(d.onesample$predicted_meanY[d.onesample$interv==0])))
}
# bootstrap
results <- boot(data=nhefs, statistic=std.ps, R=5)
# generating confidence intervals
se <- c(sd(results$t[,1]), sd(results$t[,2]),</pre>
        sd(results$t[,3]), sd(results$t[,4]))
mean <- results$t0</pre>
11 <- mean - qnorm(0.975)*se
ul <- mean + qnorm(0.975)*se
bootstrap <- data.frame(cbind(c("Observed", "No Treatment", "Treatment",</pre>
                                 "Treatment - No Treatment"), mean, se, 11, ul))
bootstrap
                           V1
##
                                           mean
                                                                                 11
                                                                se
## 1
                     Observed 2.63384609228479 0.104454848488536 2.42911835123668
## 2
                 No Treatment 1.71983636149843 0.101258698199119 1.52137295990674
                    Treatment 5.35072300362993 0.415472835100134 4.53641121027892
## 3
## 4 Treatment - No Treatment 3.63088664213151 0.457682731419862 2.73384497220266
## 1 2.83857383333291
```

## 2 1.91829976309011 ## 3 6.16503479698094 ## 4 4.52792831206035

```
# regression on the propensity score (linear term)
model6 <- glm(wt82_71 ~ qsmk + ps, data = nhefs) # p.qsmk
summary(model6)
##
## Call:
## glm(formula = wt82_71 ~ qsmk + ps, data = nhefs)
##
## Deviance Residuals:
##
       Min
                 10
                     Median
                                   3Q
                                           Max
## -43.314
            -4.006
                     -0.068
                                4.244
                                        47.158
##
## Coefficients:
               Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                 5.5945
                            0.4831 11.581 < 2e-16 ***
## qsmk
                 3.5506
                            0.4573
                                    7.765 1.47e-14 ***
## ps
               -14.8218
                            1.7576 -8.433 < 2e-16 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for gaussian family taken to be 58.28455)
##
       Null deviance: 97176 on 1565 degrees of freedom
## Residual deviance: 91099 on 1563 degrees of freedom
     (63 observations deleted due to missingness)
## AIC: 10815
##
## Number of Fisher Scoring iterations: 2
# standarization on the propensity score
# (step 1) create two new datasets, one with all treated and one with all untreated
treated <- nhefs</pre>
  treated$qsmk <- 1
untreated <- nhefs
  untreated$qsmk <- 0
# (step 2) predict values for everyone in each new dataset based on above model
treated$pred.y <- predict(model6, treated)</pre>
untreated$pred.y <- predict(model6, untreated)</pre>
# (step 3) compare mean weight loss had all been treated vs. that had all been untreated
mean1 <- mean(treated$pred.y, na.rm = TRUE)</pre>
mean0 <- mean(untreated$pred.y, na.rm = TRUE)</pre>
mean1
```

## [1] 5.250824

```
## [1] 1.700228
mean1 - mean0
## [1] 3.550596
# (step 4) bootstrap a confidence interval
# number of bootstraps
nboot <- 100
# set up a matrix to store results
boots <- data.frame(i = 1:nboot,</pre>
                     mean1 = NA,
                     mean0 = NA.
                     difference = NA)
# loop to perform the bootstrapping
nhefs <- subset(nhefs, !is.na(ps) & !is.na(wt82_71)) # p.qsmk</pre>
for(i in 1:nboot) {
  # sample with replacement
  sampl <- nhefs[sample(1:nrow(nhefs), nrow(nhefs), replace = TRUE), ]</pre>
  # fit the model in the bootstrap sample
  bootmod <- glm(wt82_71 ~ qsmk + ps, data = sampl) # ps
  # create new datasets
  sampl.treated <- sampl %>%
    mutate(qsmk = 1)
  sampl.untreated <- sampl %>%
    mutate(qsmk = 0)
  # predict values
  sampl.treated$pred.y <- predict(bootmod, sampl.treated)</pre>
  sampl.untreated$pred.y <- predict(bootmod, sampl.untreated)</pre>
  # output results
  boots[i, 'mean1'] <- mean(sampl.treated$pred.y, na.rm = TRUE)</pre>
  boots[i, 'mean0'] <- mean(sampl.untreated$pred.y, na.rm = TRUE)</pre>
  boots[i, 'difference'] <- boots[i, 'mean1'] - boots[i, 'mean0']</pre>
  # once loop is done, print the results
  if(i == nboot) {
    cat('95% CI for the causal mean difference\n')
    cat(mean(boots$difference) - 1.96*sd(boots$difference),
        ١,١,
        mean(boots$difference) + 1.96*sd(boots$difference))
 }
```

```
## 95% CI for the causal mean difference ## 2.629013 , 4.501861
```

A more flexible and elegant way to do this is to write a function to perform the model fitting, prediction, bootstrapping, and reporting all at once.

## 16. Instrumental variables estimation

- Estimating the average causal using the standard IV estimator via the calculation of sample averages
- Data from NHEFS

```
library(here)
#install.packages("readxl") # install package if required
library("readxl")
nhefs <- read_excel(here("data", "NHEFS.xls"))</pre>
# some preprocessing of the data
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)</pre>
summary(nhefs$price82)
##
      Min. 1st Qu. Median
                               Mean 3rd Qu.
                                               Max.
                                                       NA's
     1.452
           1.740
                    1.815
                             1.806
                                                          92
##
                                    1.868
                                              2.103
# for simplicity, ignore subjects with missing outcome or missing instrument
nhefs.iv <- nhefs[which(!is.na(nhefs$wt82) & !is.na(nhefs$price82)),]</pre>
nhefs.iv$highprice <- ifelse(nhefs.iv$price82>=1.5, 1, 0)
table(nhefs.iv$highprice, nhefs.iv$qsmk)
##
##
          0
               1
##
         33
               8
     1 1065 370
##
t.test(wt82_71 ~ highprice, data=nhefs.iv)
##
   Welch Two Sample t-test
## data: wt82_71 by highprice
```

```
## t = -0.10179, df = 41.644, p-value = 0.9194
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -3.130588 2.830010
## sample estimates:
## mean in group 0 mean in group 1
## 2.535729 2.686018
```

#### Program 16.2

- Estimating the average causal effect using the standard IV estimator via two-stage-least-squares regression
- Data from NHEFS

```
#install.packages ("sem") # install package if required
library(sem)
model1 <- tsls(wt82_71 ~ qsmk, ~ highprice, data = nhefs.iv)</pre>
summary(model1)
##
##
   2SLS Estimates
##
## Model Formula: wt82_71 ~ qsmk
##
## Instruments: ~highprice
##
## Residuals:
       Min.
               1st Qu.
                          Median
##
                                      Mean
                                              3rd Qu.
                                                           Max.
## -43.34863 -4.00206 -0.02712
                                   0.00000
                                              4.17040 46.47022
##
                Estimate Std. Error t value Pr(>|t|)
               2.068164
                           5.085098 0.40671 0.68428
## (Intercept)
                2.396270 19.840037 0.12078 0.90388
## Residual standard error: 7.8561141 on 1474 degrees of freedom
confint(model1) # note the wide confidence intervals
##
                    2.5 %
                            97.5 %
## (Intercept)
               -7.898445 12.03477
## qsmk
               -36.489487 41.28203
```

### Program 16.3

• Estimating the average causal using the standard IV estimator via additive marginal structural models

- Data from NHEFS
- G-estimation: Checking one possible value of psi
- See Chapter 14 for program that checks several values and computes 95% confidence intervals

```
nhefs.iv$psi <- 2.396
nhefs.iv$Hpsi <- nhefs.iv$wt82_71-nhefs.iv$psi*nhefs.iv$qsmk
#install.packages("geepack") # install package if required
library("geepack")
g.est <- geeglm(highprice ~ Hpsi, data=nhefs.iv, id=seqn, family=binomial(),</pre>
                corstr="independence")
summary(g.est)
##
## Call:
## geeglm(formula = highprice ~ Hpsi, family = binomial(), data = nhefs.iv,
##
       id = seqn, corstr = "independence")
##
##
   Coefficients:
##
                Estimate
                           Std.err Wald Pr(>|W|)
## (Intercept) 3.555e+00 1.652e-01 463.1
                                            <2e-16 ***
               2.748e-07 2.273e-02
                                                 1
## Hpsi
                                      0.0
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Correlation structure = independence
## Estimated Scale Parameters:
##
               Estimate Std.err
##
## (Intercept)
                      1 0.7607
## Number of clusters:
                         1476 Maximum cluster size: 1
beta <- coef(g.est)</pre>
SE <- coef(summary(g.est))[,2]</pre>
lcl <- beta-qnorm(0.975)*SE</pre>
ucl <- beta+qnorm(0.975)*SE
cbind(beta, lcl, ucl)
##
                               lcl
                                       110]
                    beta
## (Intercept) 3.555e+00 3.23152 3.87917
## Hpsi
               2.748e-07 -0.04456 0.04456
```

- Estimating the average causal using the standard IV estimator with altherative proposed instruments
- Data from NHEFS

```
summary(tsls(wt82_71 ~ qsmk, ~ ifelse(price82 >= 1.6, 1, 0), data = nhefs.iv))
##
##
  2SLS Estimates
##
## Model Formula: wt82_71 ~ qsmk
##
## Instruments: ~ifelse(price82 >= 1.6, 1, 0)
##
## Residuals:
     Min. 1st Qu. Median
##
                              Mean 3rd Qu.
                                              Max.
##
    -55.6 -13.5
                       7.6
                               0.0
                                      12.5
                                              56.4
##
              Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  -7.89
                             42.25 -0.187
                                              0.852
## qsmk
                  41.28
                            164.95
                                    0.250
                                              0.802
##
## Residual standard error: 18.6055 on 1474 degrees of freedom
summary(tsls(wt82_71 ~ qsmk, ~ ifelse(price82 >= 1.7, 1, 0), data = nhefs.iv))
##
##
   2SLS Estimates
##
## Model Formula: wt82_71 ~ qsmk
##
## Instruments: ~ifelse(price82 >= 1.7, 1, 0)
## Residuals:
##
     Min. 1st Qu. Median
                              Mean 3rd Qu.
                                              Max.
     -54.4 -13.4
                      -8.4
                               0.0
                                              75.3
##
                                      18.1
##
              Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  13.16
                             48.08
                                    0.274
                                              0.784
## qsmk
                 -40.91
                            187.74 -0.218
                                              0.828
## Residual standard error: 20.591 on 1474 degrees of freedom
summary(tsls(wt82_71 ~ qsmk, ~ ifelse(price82 >= 1.8, 1, 0), data = nhefs.iv))
##
##
   2SLS Estimates
##
## Model Formula: wt82_71 ~ qsmk
## Instruments: ~ifelse(price82 >= 1.8, 1, 0)
##
```

```
## Residuals:
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
   -49.37
                     -3.44
##
           -8.31
                              0.00
                                      7.27
                                              60.53
##
               Estimate Std. Error t value Pr(>|t|)
##
                  8.086
                             7.288
                                      1.110
## (Intercept)
                                               0.267
## qsmk
                -21.103
                            28.428 -0.742
                                               0.458
##
## Residual standard error: 13.0188 on 1474 degrees of freedom
summary(tsls(wt82_71 ~ qsmk, ~ ifelse(price82 >= 1.9, 1, 0), data = nhefs.iv))
##
##
   2SLS Estimates
##
## Model Formula: wt82_71 ~ qsmk
## Instruments: ~ifelse(price82 >= 1.9, 1, 0)
##
## Residuals:
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
   -47.24 -6.33
##
                    -1.43
                              0.00
                                       5.52
                                              54.36
##
##
               Estimate Std. Error t value Pr(>|t|)
                                      0.983
## (Intercept)
                  5.963
                             6.067
                                               0.326
## qsmk
                -12.811
                            23.667 -0.541
                                               0.588
##
## Residual standard error: 10.3637 on 1474 degrees of freedom
```

- Estimating the average causal using the standard IV estimator
- Conditional on baseline covariates
- Data from NHEFS

```
## Instruments: ~highprice + sex + race + age + smokeintensity + smokeyrs + as.factor(exercise) +
##
      as.factor(active) + wt71
##
## Residuals:
     Min. 1st Qu. Median
##
                           Mean 3rd Qu.
                                          Max.
## -42.23 -4.29 -0.62
                           0.00
                                  3.87
                                         46.74
##
##
                      Estimate Std. Error t value Pr(>|t|)
                                          7.399 2.3e-13 ***
## (Intercept)
                      17.280330
                                 2.335402
                     -1.042295 29.987369 -0.035 0.9723
## qsmk
                                 2.630831 -0.625 0.5320
## sex
                      -1.644393
## race
                     -0.183255 4.650386 -0.039 0.9686
## age
                      -0.163640
                                 0.240548 -0.680 0.4964
## smokeintensity
                      0.005767
                                 0.145504 0.040 0.9684
## smokeyrs
                      0.025836
                                 0.161421 0.160 0.8729
                                 2.171239 0.230 0.8184
## as.factor(exercise)1 0.498748
## as.factor(exercise)2 0.581834
                                 2.183148 0.267 0.7899
## as.factor(active)1 -1.170145
                                 0.607466 -1.926
                                                  0.0543 .
## as.factor(active)2 -0.512284
                                 1.308451 -0.392 0.6955
## wt71
                     -0.097949
                                 0.036271 -2.701
                                                  0.0070 **
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Residual standard error: 7.7162 on 1464 degrees of freedom
```

## 17. Causal survival analysis

## Program 17.1

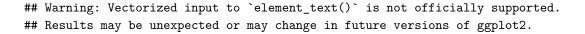
• Nonparametric estimation of survival curves

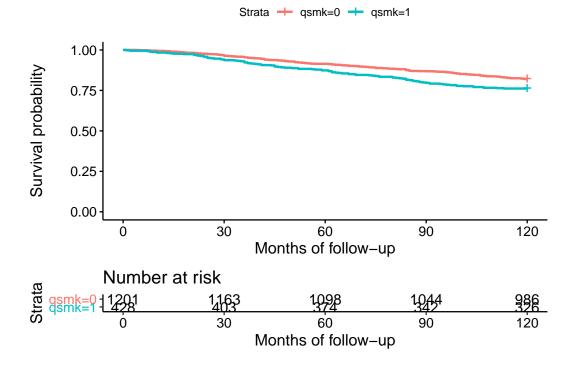
## Loading required package: ggpubr

• Data from NHEFS

```
library(here)
library("readxl")
nhefs <- read_excel(here("data","NHEFS.xls"))</pre>
# some preprocessing of the data
nhefs$survtime <- ifelse(nhefs$death==0, 120,</pre>
                         (nhefs\$yrdth-83)*12+nhefs\$modth) # yrdth ranges from 83 to 92
table(nhefs$death, nhefs$qsmk)
##
##
         0
    0 985 326
##
     1 216 102
summary(nhefs[which(nhefs$death==1),]$survtime)
##
      Min. 1st Qu. Median Mean 3rd Qu.
      1.00 35.00 61.00 61.14 86.75 120.00
##
#install.packages("survival")
#install.packages("ggplot2") # for plots
#install.packages("survminer") # for plots
library("survival")
library("ggplot2")
library("survminer")
```

```
survdiff(Surv(survtime, death) ~ qsmk, data=nhefs)
## Call:
## survdiff(formula = Surv(survtime, death) ~ qsmk, data = nhefs)
##
##
             N Observed Expected (0-E)^2/E (0-E)^2/V
## qsmk=0 1201
                    216
                           237.5
                                       1.95
                                                 7.73
                    102
                            80.5
                                       5.76
                                                 7.73
## qsmk=1 428
##
    Chisq= 7.7 on 1 degrees of freedom, p= 0.005
fit <- survfit(Surv(survtime, death) ~ qsmk, data=nhefs)</pre>
ggsurvplot(fit, data = nhefs, xlab="Months of follow-up",
           ylab="Survival probability",
           main="Product-Limit Survival Estimates", risk.table = TRUE)
```



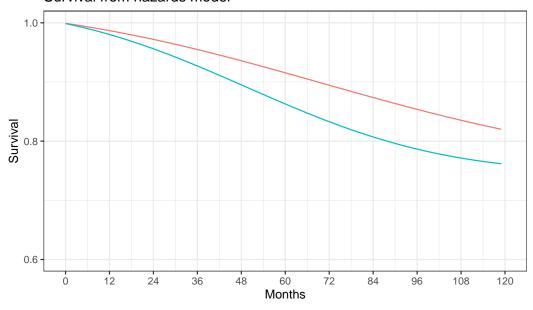


- Parametric estimation of survival curves via hazards model
- Data from NHEFS

```
# creation of person-month data
#install.packages("splitstackshape")
library("splitstackshape")
nhefs.surv <- expandRows(nhefs, "survtime", drop=F)</pre>
nhefs.surv$time <- sequence(rle(nhefs.surv$seqn)$lengths)-1
nhefs.surv$event <- ifelse(nhefs.surv$time==nhefs.surv$survtime-1 &
                            nhefs.surv$death==1, 1, 0)
nhefs.surv$timesq <- nhefs.surv$time^2</pre>
# fit of parametric hazards model
hazards.model <- glm(event==0 ~ qsmk + I(qsmk*time) + I(qsmk*timesq) +
                       time + timesq, family=binomial(), data=nhefs.surv)
summary(hazards.model)
##
## Call:
## glm(formula = event == 0 ~ qsmk + I(qsmk * time) + I(qsmk * timesq) +
       time + timesq, family = binomial(), data = nhefs.surv)
##
##
## Deviance Residuals:
##
      Min
                 1Q
                     Median
                                  3Q
                                           Max
## -3.7253
           0.0546 0.0601
                              0.0625
                                        0.0783
##
## Coefficients:
##
                     Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                   6.996e+00 2.309e-01 30.292
                                                   <2e-16 ***
                   -3.355e-01 3.970e-01 -0.845
                                                   0.3981
## qsmk
## I(qsmk * time) -1.208e-02 1.503e-02 -0.804
                                                   0.4215
## I(qsmk * timesq) 1.612e-04 1.246e-04 1.293
                                                   0.1960
## time
                   -1.960e-02 8.413e-03 -2.329
                                                   0.0198 *
                    1.256e-04 6.686e-05 1.878 0.0604.
## timesq
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 4655.3 on 176763 degrees of freedom
## Residual deviance: 4631.3 on 176758 degrees of freedom
## AIC: 4643.3
##
## Number of Fisher Scoring iterations: 9
# creation of dataset with all time points under each treatment level
qsmk0 <- data.frame(cbind(seq(0, 119),0,(seq(0, 119))^2))
qsmk1 <- data.frame(cbind(seq(0, 119),1,(seq(0, 119))^2))
colnames(qsmk0) <- c("time", "qsmk", "timesq")</pre>
colnames(qsmk1) <- c("time", "qsmk", "timesq")</pre>
```

```
# assignment of estimated (1-hazard) to each person-month */
qsmk0$p.noevent0 <- predict(hazards.model, qsmk0, type="response")</pre>
qsmk1$p.noevent1 <- predict(hazards.model, qsmk1, type="response")</pre>
# computation of survival for each person-month
qsmk0$surv0 <- cumprod(qsmk0$p.noevent0)</pre>
qsmk1$surv1 <- cumprod(qsmk1$p.noevent1)</pre>
# some data management to plot estimated survival curves
hazards.graph <- merge(qsmk0, qsmk1, by=c("time", "timesq"))</pre>
hazards.graph$survdiff <- hazards.graph$surv1-hazards.graph$surv0
# plot
ggplot(hazards.graph, aes(x=time, y=surv)) +
  geom_line(aes(y = surv0, colour = "0")) +
  geom_line(aes(y = surv1, colour = "1")) +
 xlab("Months") +
  scale_x_continuous(limits = c(0, 120), breaks=seq(0,120,12)) +
  scale_y_continuous(limits=c(0.6, 1), breaks=seq(0.6, 1, 0.2)) +
  ylab("Survival") +
  ggtitle("Survival from hazards model") +
  labs(colour="A:") +
  theme_bw() +
  theme(legend.position="bottom")
```

#### Survival from hazards model



A: — 0 — 1

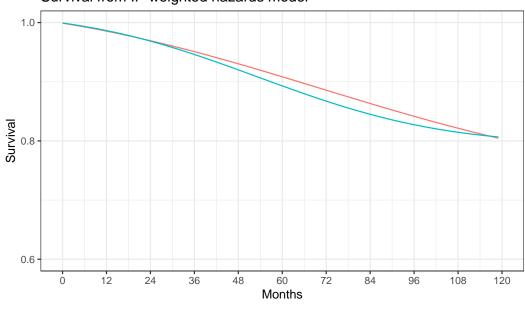
- Estimation of survival curves via IP weighted hazards model
- Data from NHEFS

```
# estimation of denominator of ip weights
p.denom <- glm(qsmk ~ sex + race + age + I(age*age) + as.factor(education)
               + smokeintensity + I(smokeintensity*smokeintensity)
               + smokeyrs + I(smokeyrs*smokeyrs) + as.factor(exercise)
               + as.factor(active) + wt71 + I(wt71*wt71),
               data=nhefs, family=binomial())
nhefs$pd.qsmk <- predict(p.denom, nhefs, type="response")</pre>
# estimation of numerator of ip weights
p.num <- glm(qsmk ~ 1, data=nhefs, family=binomial())</pre>
nhefs$pn.qsmk <- predict(p.num, nhefs, type="response")</pre>
# computation of estimated weights
nhefs$sw.a <- ifelse(nhefs$qsmk==1, nhefs$pn.qsmk/nhefs$pd.qsmk,
                     (1-nhefs$pn.qsmk)/(1-nhefs$pd.qsmk))
summary(nhefs$sw.a)
      Min. 1st Qu. Median
                              Mean 3rd Qu.
##
                                               Max.
   0.3312 0.8640 0.9504 0.9991 1.0755 4.2054
# creation of person-month data
nhefs.ipw <- expandRows(nhefs, "survtime", drop=F)</pre>
nhefs.ipw$time <- sequence(rle(nhefs.ipw$seqn)$lengths)-1</pre>
nhefs.ipw$event <- ifelse(nhefs.ipw$time==nhefs.ipw$survtime-1 &
                            nhefs.ipw$death==1, 1, 0)
nhefs.ipw$timesq <- nhefs.ipw$time^2</pre>
# fit of weighted hazards model
ipw.model <- glm(event==0 ~ qsmk + I(qsmk*time) + I(qsmk*timesq) +
                   time + timesq, family=binomial(), weight=sw.a,
                 data=nhefs.ipw)
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
summary(ipw.model)
##
## Call:
## glm(formula = event == 0 ~ qsmk + I(qsmk * time) + I(qsmk * timesq) +
##
       time + timesq, family = binomial(), data = nhefs.ipw, weights = sw.a)
## Deviance Residuals:
##
       Min
               1Q Median
                                   3Q
                                            Max
```

```
## -7.1859 0.0528 0.0595 0.0640
                                        0.1452
##
## Coefficients:
##
                      Estimate Std. Error z value Pr(>|z|)
                     6.897e+00 2.208e-01 31.242
## (Intercept)
                                                     <2e-16 ***
                    1.794e-01 4.399e-01 0.408 0.6834
## qsmk
## I(qsmk * time)
                    -1.895e-02 1.640e-02 -1.155 0.2481
## I(qsmk * timesq) 2.103e-04 1.352e-04 1.556 0.1198
                    -1.889e-02 8.053e-03 -2.345
                                                     0.0190 *
## time
                    1.181e-04 6.399e-05 1.846 0.0649 .
## timesq
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 4643.9 on 176763 degrees of freedom
## Residual deviance: 4626.2 on 176758 degrees of freedom
## AIC: 4633.5
##
## Number of Fisher Scoring iterations: 9
# creation of survival curves
ipw.qsmk0 <- data.frame(cbind(seq(0, 119),0,(seq(0, 119))^2))</pre>
ipw.qsmk1 \leftarrow data.frame(cbind(seq(0, 119), 1, (seq(0, 119))^2))
colnames(ipw.qsmk0) <- c("time", "qsmk", "timesq")</pre>
colnames(ipw.qsmk1) <- c("time", "qsmk", "timesq")</pre>
# assignment of estimated (1-hazard) to each person-month */
ipw.qsmk0$p.noevent0 <- predict(ipw.model, ipw.qsmk0, type="response")</pre>
ipw.qsmk1$p.noevent1 <- predict(ipw.model, ipw.qsmk1, type="response")</pre>
# computation of survival for each person-month
ipw.qsmk0$surv0 <- cumprod(ipw.qsmk0$p.noevent0)</pre>
ipw.qsmk1$surv1 <- cumprod(ipw.qsmk1$p.noevent1)</pre>
# some data management to plot estimated survival curves
ipw.graph <- merge(ipw.qsmk0, ipw.qsmk1, by=c("time", "timesq"))</pre>
ipw.graph$survdiff <- ipw.graph$surv1-ipw.graph$surv0</pre>
# plot
ggplot(ipw.graph, aes(x=time, y=surv)) +
  geom_line(aes(y = surv0, colour = "0")) +
  geom_line(aes(y = surv1, colour = "1")) +
 xlab("Months") +
  scale_x_continuous(limits = c(0, 120), breaks=seq(0,120,12)) +
  scale_y_continuous(limits=c(0.6, 1), breaks=seq(0.6, 1, 0.2)) +
  ylab("Survival") +
  ggtitle("Survival from IP weighted hazards model") +
```

```
labs(colour="A:") +
theme_bw() +
theme(legend.position="bottom")
```

#### Survival from IP weighted hazards model



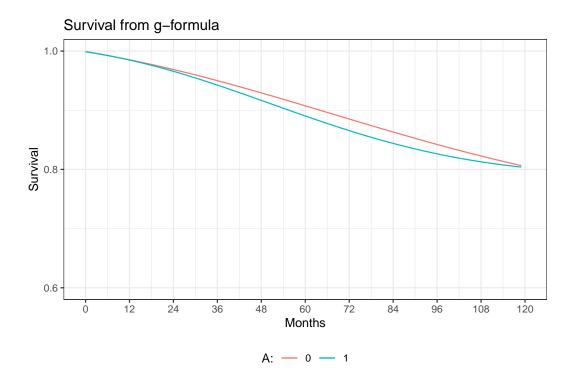
#### A: — 0 — 1

- Estimating of survival curves via g-formula
- Data from NHEFS

```
##
## Call:
## glm(formula = event == 0 ~ qsmk + I(qsmk * time) + I(qsmk * timesq) +
## time + timesq + sex + race + age + I(age * age) + as.factor(education) +
## smokeintensity + I(smokeintensity * smokeintensity) + smkintensity82_71 +
## smokeyrs + I(smokeyrs * smokeyrs) + as.factor(exercise) +
```

```
##
      as.factor(active) + wt71 + I(wt71 * wt71), family = binomial(),
##
      data = nhefs.surv)
##
## Deviance Residuals:
##
      Min
                1Q
                     Median
                                  3Q
                                          Max
## -4.3160
                     0.0395
           0.0244
                              0.0640
                                       0.3303
##
## Coefficients:
##
                                       Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                                      9.272e+00 1.379e+00 6.724 1.76e-11 ***
                                      5.959e-02 4.154e-01
                                                             0.143 0.885924
## qsmk
## I(qsmk * time)
                                     -1.485e-02 1.506e-02 -0.987 0.323824
## I(qsmk * timesq)
                                      1.702e-04 1.245e-04
                                                            1.367 0.171643
## time
                                     -2.270e-02 8.437e-03 -2.690 0.007142 **
                                      1.174e-04 6.709e-05 1.751 0.080020 .
## timesq
## sex
                                      4.368e-01 1.409e-01
                                                             3.101 0.001930 **
                                     -5.240e-02 1.734e-01 -0.302 0.762572
## race
## age
                                     -8.750e-02 5.907e-02 -1.481 0.138536
                                      8.128e-05 5.470e-04
                                                           0.149 0.881865
## I(age * age)
## as.factor(education)2
                                      1.401e-01 1.566e-01 0.895 0.370980
## as.factor(education)3
                                      4.335e-01 1.526e-01 2.841 0.004502 **
## as.factor(education)4
                                      2.350e-01 2.790e-01
                                                             0.842 0.399750
## as.factor(education)5
                                      3.750e-01 2.386e-01 1.571 0.116115
## smokeintensity
                                     -1.626e-03 1.430e-02 -0.114 0.909431
## I(smokeintensity * smokeintensity) -7.182e-05 2.390e-04 -0.301 0.763741
## smkintensity82_71
                                     -1.686e-03 6.501e-03 -0.259 0.795399
## smokeyrs
                                     -1.677e-02 3.065e-02 -0.547 0.584153
## I(smokeyrs * smokeyrs)
                                     -5.280e-05 4.244e-04 -0.124 0.900997
## as.factor(exercise)1
                                     1.469e-01 1.792e-01
                                                           0.820 0.412300
## as.factor(exercise)2
                                     -1.504e-01 1.762e-01 -0.854 0.393177
## as.factor(active)1
                                     -1.601e-01 1.300e-01 -1.232 0.218048
## as.factor(active)2
                                     -2.294e-01 1.877e-01 -1.222 0.221766
## wt71
                                      6.222e-02 1.902e-02
                                                           3.271 0.001073 **
## I(wt71 * wt71)
                                     -4.046e-04 1.129e-04 -3.584 0.000338 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for binomial family taken to be 1)
##
      Null deviance: 4655.3 on 176763 degrees of freedom
## Residual deviance: 4185.7 on 176739 degrees of freedom
## AIC: 4235.7
## Number of Fisher Scoring iterations: 10
# creation of dataset with all time points for
# each individual under each treatment level
gf.qsmk0 <- expandRows(nhefs, count=120, count.is.col=F)</pre>
gf.qsmk0$time <- rep(seq(0, 119), nrow(nhefs))
```

```
gf.qsmk0$timesq <- gf.qsmk0$time^2</pre>
gf.qsmk0$qsmk <- 0
gf.qsmk1 <- gf.qsmk0
gf.qsmk1$qsmk <- 1
gf.qsmk0$p.noevent0 <- predict(gf.model, gf.qsmk0, type="response")</pre>
gf.qsmk1$p.noevent1 <- predict(gf.model, gf.qsmk1, type="response")</pre>
#install.packages("dplyr")
library("dplyr")
##
## Attaching package: 'dplyr'
## The following objects are masked from 'package:stats':
##
##
       filter, lag
## The following objects are masked from 'package:base':
##
##
       intersect, setdiff, setequal, union
gf.qsmk0.surv <- gf.qsmk0 %>% group_by(seqn) %>% mutate(surv0 = cumprod(p.noevent0))
gf.qsmk1.surv <- gf.qsmk1 %>% group_by(seqn) %>% mutate(surv1 = cumprod(p.noevent1))
gf.surv0 <- aggregate(gf.qsmk0.surv, by=list(gf.qsmk0.surv$time), FUN=mean)[c("qsmk", "time", "surv0")]
gf.surv1 <- aggregate(gf.qsmk1.surv, by=list(gf.qsmk1.surv$time), FUN=mean)[c("qsmk", "time", "surv1")]</pre>
gf.graph <- merge(gf.surv0, gf.surv1, by=c("time"))</pre>
gf.graph$survdiff <- gf.graph$surv1-gf.graph$surv0</pre>
# plot
ggplot(gf.graph, aes(x=time, y=surv)) +
  geom_line(aes(y = surv0, colour = "0")) +
  geom_line(aes(y = surv1, colour = "1")) +
  xlab("Months") +
  scale_x_continuous(limits = c(0, 120), breaks=seq(0,120,12)) +
  scale_y_continuous(limits=c(0.6, 1), breaks=seq(0.6, 1, 0.2)) +
  ylab("Survival") +
  ggtitle("Survival from g-formula") +
  labs(colour="A:") +
  theme_bw() +
  theme(legend.position="bottom")
```



- Estimating of median survival time ratio via a structural nested AFT model
- Data from NHEFS

```
# some preprocessing of the data
nhefs <- read_excel(here("data", "NHEFS.xls"))</pre>
nhefs$survtime <- ifelse(nhefs$death==0, NA, (nhefs$yrdth-83)*12+nhefs$modth) # * yrdth ranges from 83
# model to estimate E[A/L]
modelA <- glm(qsmk ~ sex + race + age + I(age*age)</pre>
               + as.factor(education) + smokeintensity
              + I(smokeintensity*smokeintensity) + smokeyrs
               + I(smokeyrs*smokeyrs) + as.factor(exercise)
              + as.factor(active) + wt71 + I(wt71*wt71),
               data=nhefs, family=binomial())
nhefs$p.qsmk <- predict(modelA, nhefs, type="response")</pre>
d <- nhefs[!is.na(nhefs$survtime),] # select only those with observed death time
n \leftarrow nrow(d)
# define the estimating function that needs to be minimized
sumeef <- function(psi){</pre>
  # creation of delta indicator
  if (psi>=0){
```

```
delta <- ifelse(d$qsmk==0 |</pre>
                        (d^{qsmk}=1 & psi \leq log(120/d^{survtime})),
                      1, 0)
  } else if (psi < 0) {</pre>
    delta <- ifelse(d$qsmk==1 |</pre>
                        (d$qsmk==0 & psi > log(d$survtime/120)), 1, 0)
  }
  smat <- delta*(d$qsmk-d$p.qsmk)</pre>
  sval <- sum(smat, na.rm=T)</pre>
  save <- sval/n</pre>
  smat <- smat - rep(save, n)</pre>
  # covariance
  sigma <- t(smat) %*% smat
  if (sigma == 0){
    sigma <- 1e-16
  estimeq <- sval*solve(sigma)*t(sval)</pre>
  return(estimeq)
}
res <- optimize(sumeef, interval = c(-0.2,0.2))
psi1 <- res$minimum
objfunc <- as.numeric(res$objective)</pre>
# Use simple bisection method to find estimates of lower and upper 95% confidence bounds
increm <- 0.1
for_conf <- function(x){</pre>
  return(sumeef(x) - 3.84)
}
if (objfunc < 3.84){
  # Find estimate of where sumeef(x) > 3.84
  # Lower bound of 95% CI
  psilow <- psi1
  testlow <- objfunc
  countlow <- 0
  while (testlow < 3.84 & countlow < 100){
    psilow <- psilow - increm</pre>
   testlow <- sumeef(psilow)</pre>
    countlow <- countlow + 1</pre>
  }
  # Upper bound of 95% CI
  psihigh <- psi1</pre>
```

```
testhigh <- objfunc
counthigh <- 0
while (testhigh < 3.84 & counthigh < 100){
  psihigh <- psihigh + increm</pre>
  testhigh <- sumeef(psihigh)</pre>
  counthigh <- counthigh + 1</pre>
}
# Better estimate using bisection method
if ((testhigh > 3.84) & (testlow > 3.84)){
  # Bisection method
  left <- psi1
  fleft <- objfunc - 3.84
  right <- psihigh
  fright <- testhigh - 3.84
  middle <- (left + right) / 2
  fmiddle <- for_conf(middle)</pre>
  count <- 0
  diff <- right - left
  while (!(abs(fmiddle) < 0.0001 | diff < 0.0001 | count > 100)){
    test <- fmiddle * fleft</pre>
    if (test < 0){</pre>
      right <- middle
      fright <- fmiddle</pre>
    } else {
      left <- middle
      fleft <- fmiddle
    middle <- (left + right) / 2
    fmiddle <- for_conf(middle)</pre>
    count <- count + 1</pre>
    diff <- right - left
  }
  psi_high <- middle</pre>
  objfunc_high <- fmiddle + 3.84
  # lower bound of 95% CI
  left <- psilow</pre>
  fleft <- testlow - 3.84
  right <- psi1
  fright <- objfunc - 3.84
  middle <- (left + right) / 2
  fmiddle <- for_conf(middle)</pre>
  count <- 0
  diff <- right - left
```

```
while(!(abs(fmiddle) < 0.0001 | diff < 0.0001 | count > 100)){
      test <- fmiddle * fleft</pre>
      if (test < 0){</pre>
        right <- middle
        fright <- fmiddle
      } else {
        left <- middle
        fleft <- fmiddle</pre>
      middle <- (left + right) / 2
      fmiddle <- for_conf(middle)</pre>
      diff <- right - left</pre>
      count <- count + 1</pre>
    }
    psi_low <- middle</pre>
    objfunc_low <- fmiddle + 3.84
    psi <- psi1
c(psi, psi_low, psi_high)
```

## [1] -0.05041591 -0.22312099 0.33312901

## Session information: R

For reproducibility.

```
# install.packages("sessioninfo")
sessioninfo::session_info()
  - Session info -----
   setting value
   version R version 4.0.0 (2020-04-24)
##
           Windows 10 x64
##
  os
  system
           x86_64, mingw32
##
##
  ui
           RTerm
   language (EN)
##
  collate English_United Kingdom.1252
##
   ctype
           English_United Kingdom.1252
##
  tz
           Europe/London
##
   date
           2020-05-14
##
## - Packages ------
   package
              * version date
                                 lib source
##
##
   assertthat
                0.2.1
                       2019-03-21 [1] CRAN (R 4.0.0)
   bookdown
                0.18
                       2020-03-05 [1] CRAN (R 4.0.0)
                2.0.2
                       2020-02-28 [1] CRAN (R 4.0.0)
##
   cli
                1.3.4
                       2017-09-16 [1] CRAN (R 4.0.0)
##
   crayon
   digest
                0.6.25 2020-02-23 [1] CRAN (R 4.0.0)
##
   evaluate
                0.14
                       2019-05-28 [1] CRAN (R 4.0.0)
##
##
  fansi
                0.4.1
                       2020-01-08 [1] CRAN (R 4.0.0)
                1.4.0
                       2020-04-03 [1] CRAN (R 4.0.0)
##
   glue
                0.4.0
                       2019-10-04 [1] CRAN (R 4.0.0)
## htmltools
                       2020-02-06 [1] CRAN (R 4.0.0)
##
  knitr
                1.28
                       2014-11-22 [1] CRAN (R 4.0.0)
   magrittr
                1.5
##
##
   Rcpp
                1.0.4.6 2020-04-09 [1] CRAN (R 4.0.0)
                       2020-05-02 [1] CRAN (R 4.0.0)
                0.4.6
## rlang
                       2020-01-20 [1] CRAN (R 4.0.0)
##
  rmarkdown
                2.1
## sessioninfo
                1.1.1
                       2018-11-05 [1] CRAN (R 4.0.0)
                       2020-02-17 [1] CRAN (R 4.0.0)
## stringi
                1.4.6
                1.4.0
                       2019-02-10 [1] CRAN (R 4.0.0)
## stringr
## withr
                2.2.0
                       2020-04-20 [1] CRAN (R 4.0.0)
                       2020-04-13 [1] CRAN (R 4.0.0)
                0.13
## xfun
```

```
## yaml 2.2.1 2020-02-01 [1] CRAN (R 4.0.0)
##
## [1] C:/Users/tom/Documents/R/win-library/4.0
## [2] C:/Program Files/R/R-4.0.0/library
```

# Stata code

# 11. Why model: Stata

- Figures 11.1, 11.2, and 11.3
- Sample averages by treatment level

```
**Figure 11.1**
*create the dataset*
input A Y
1 200
1 150
1 220
1 110
1 50
1 180
```

```
1 90
1 170
0 170
0 30
0 70
0 110
0 80
0 50
0 10
0 20
end
*Save the data*
qui save ./data/fig1, replace
*Build the scatterplot*
scatter Y A, ylab(0(50)250) xlab(0 1) xscale(range(-0.5 1.5))
qui gr export figs/stata-fig-11-1.png, replace
*Output the mean values for Y in each level of A*
bysort A: sum Y
```

A Y

\_\_\_\_\_

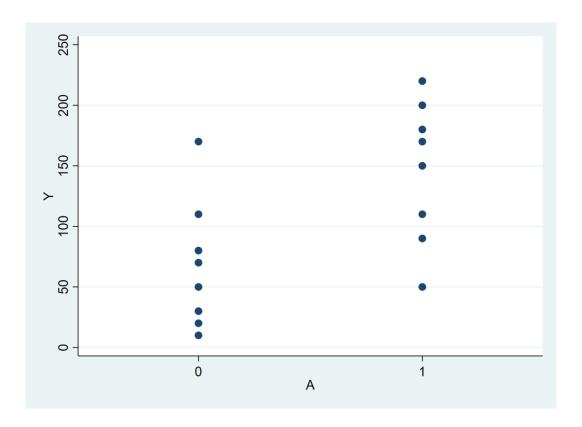
-> A = O

Variable	l Obs	Mean	Std. Dev.	Min	Max	
Y	l 8	67.5	53.11712	10	170	

-----

-> A = 1

Variable	1	0bs	Mean	Std. Dev.	Min	Max
	+					
Y	1	8	146.25	58.2942	50	220

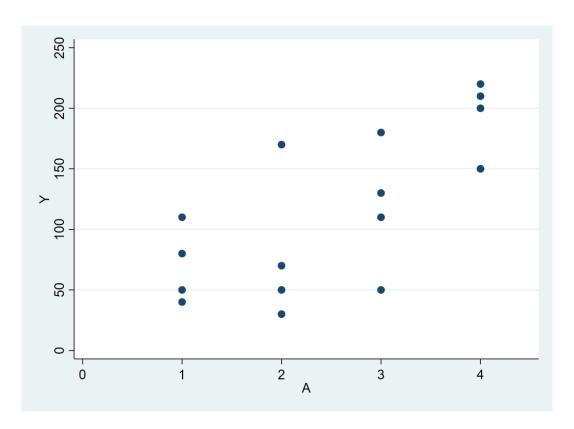


```
*Clear the workspace to be able to use a new dataset*
clear
**Figure 11.2**
input A Y
1 110
1 80
1 50
1 40
2 170
2 30
2 70
2 50
3 110
3 50
3 180
3 130
4 200
4 150
4 220
4 210
end
qui save ./data/fig2, replace
scatter Y A, ylab(0(50)250) xlab(0(1)4) xscale(range(0 4.5))
```

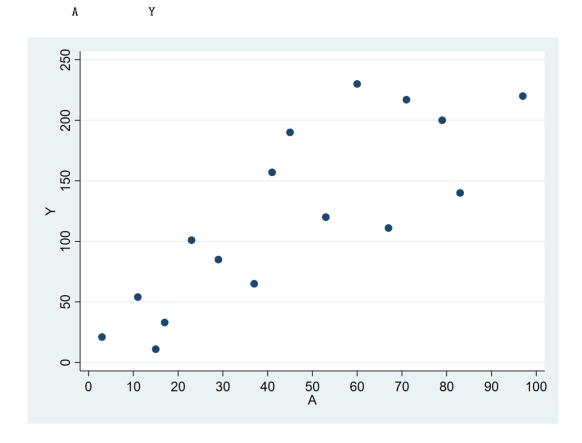
```
qui gr export figs/stata-fig-11-2.png, replace
bysort A: sum Y
```

A Y

A = 1	 		 
Variable			
·		31.62278	
A = 2	 		 
Variable			
		62.18253	
A = 3	 		 
Variable			Max
		53.77422	
A = 4	 		 
Variable			
•		31.09126	



```
clear
**Figure 11.3**
\mathtt{input}\ \mathtt{A}\ \mathtt{Y}
3
    21
11
    54
17 33
23 101
29 85
37 65
41 157
53 120
67 111
79 200
83 140
97
    220
60
   230
71
    217
15 11
45
   190
end
qui save ./data/fig3, replace
scatter Y A, ylab(0(50)250) xlab(0(10)100) xscale(range(0 100))
qui gr export figs/stata-fig-11-3.png, replace
```



- 2-parameter linear model
- $\bullet$  Creates Figure 11.4, parameter estimates with 95% confidence intervals from Section 11.2, and parameter estimates with 95% confidence intervals from Section 11.3

```
**Section 11.2: parametric estimators**
*Reload data
use ./data/fig3, clear

*Plot the data*
scatter Y A, ylab(0(50)250) xlab(0(10)100) xscale(range(0 100))

*Fit the regression model*
regress Y A, noheader cformat(%5.2f)

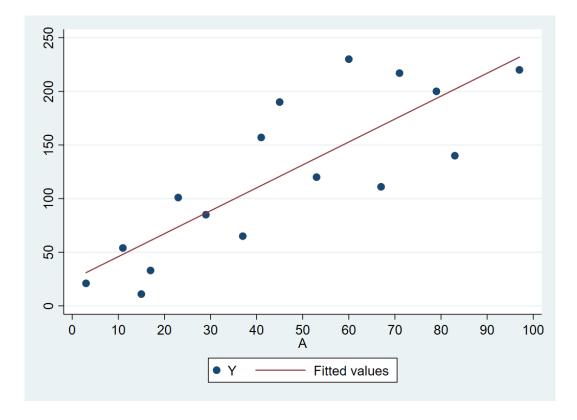
*Output the estimated mean Y value when A = 90*
lincom _b[_cons] + 90*_b[A]

*Plot the data with the regression line: Fig 11.4*
scatter Y A, ylab(0(50)250) xlab(0(10)100) xscale(range(0 100)) || lfit Y A
qui gr export figs/stata-fig-11-4.png, replace
```

			• • •	[95% Conf.	•
A	0.40	5.35	0.000	1.28 -21.20	2.99

(1) 90\*A + cons = 0

Y	•		 [95% Conf.	Interval]
			172.1468	261.6333



```
**Section 11.3: non-parametric estimation*

* Reload the data
use ./data/fig1, clear

*Fit the regression model*
regress Y A, noheader cformat(%5.2f)

*E[Y|A=1]*
di 67.50 + 78.75
```

Y | Coef. Std. Err. t P>|t| [95% Conf. Interval]

+-						
A	78.75	27.88	2.82	0.014	18.95	138.55
_cons	67.50	19.72	3.42	0.004	25.21	109.79

146.25

- $\bullet$  3-parameter linear model
- Creates Figure 11.5 and Parameter estimates for Section 11.4

```
* Reload the data
use ./data/fig3, clear

*Create the product term*
gen Asq = A*A

*Fit the regression model*
regress Y A Asq, noheader cformat(%5.2f)

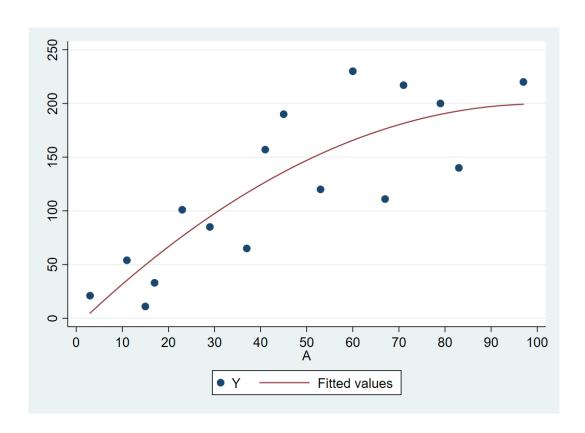
*Output the estimated mean Y value when A = 90*
lincom _b[_cons] + 90*_b[A] + 90*90*_b[Asq]

*Plot the data with the regression line: Fig 11.5*
scatter Y A, ylab(0(50)250) xlab(0(10)100) xscale(range(0 100)) || qfit Y A
qui gr export figs/stata-fig-11-5.png, replace
```

Y	Coef.	Std. Err.	t	P> t	[95% Conf.	Interval]
Α	4.11	1.53	2.68	0.019	0.80	7.41
Asq	-0.02	0.02	-1.33	0.206	-0.05	0.01
_cons	-7.41	31.75	-0.23	0.819	-75.99	61.18

```
(1) 90*A + 8100*Asq + _cons = 0
```

•		• • •	[95% Conf. In	-
			142.7687 2	



# 12. IP Weighting and Marginal Structural Models: Stata

## Program 12.1

library(Statamarkdown)

• Descriptive statistics from NHEFS data (Table 12.1)

```
use ./data/nhefs, clear
/*Provisionally ignore subjects with missing values for follow-up weight*/
/*Sample size after exclusion: N = 1566*/
drop if wt82==.
/* Calculate mean weight change in those with and without smoking cessation*/
label define qsmk 0 "No smoking cessation" 1 "Smoking cessation"
label values qsmk qsmk
by qsmk, sort: egen years = mean(age) if age < .
label var years "Age, years"
by qsmk, sort: egen male = mean(100 * (sex==0)) if sex < .
label var male "Men, %"
by qsmk, sort: egen white = mean(100 * (race==0)) if race < .
label var white "White, %"
by qsmk, sort: egen university = mean(100 * (education == 5)) if education < .
label var university "University, %"
by qsmk, sort: egen kg = mean(wt71) if wt71 < .
label var kg "Weight, kg"
by qsmk, sort: egen cigs = mean(smokeintensity) if smokeintensity < .
```

```
label var cigs "Cigarettes/day"
by qsmk, sort: egen meansmkyrs = mean(smokeyrs) if smokeyrs < .
label var kg "Years smoking"
by qsmk, sort: egen noexer = mean(100 * (exercise == 2)) if exercise < .
label var noexer "Little/no exercise"
by qsmk, sort: egen inactive = mean(100 * (active==2)) if active < .
label var inactive "Inactive daily life"
qui save ./data/nhefs-formatted, replace
(63 observations deleted)
use ./data/nhefs-formatted, clear
/*Output table*/
foreach var of varlist years male white university kg cigs meansmkyrs noexer inactive {
 tabdisp qsmk, cell(`var') format(%3.1f)
}
quit smoking between |
baseline and 1982 | Age, years
-----
No smoking cessation |
  Smoking cessation |
                       46.2
_____
quit smoking between |
baseline and 1982 | Men, %
-----
No smoking cessation |
                       46.6
  Smoking cessation |
                       54.6
quit smoking between |
baseline and 1982 | White, \%
----+----
No smoking cessation |
                       85.4
                        91.1
  Smoking cessation |
quit smoking between |
baseline and 1982 | University, %
No smoking cessation |
                           9.9
                           15.4
  Smoking cessation |
```

   Years smoking
70.3   72.4
   Cigarettes/day
21.2   18.6
   meansmkyrs
24.1   26.0
37.9   40.7
8.9   11.2

- Estimating IP weights for Section 12.2
- Data from NHEFS

```
use ./data/nhefs-formatted, clear
/*Fit a logistic model for the IP weights*/
logit qsmk sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity ///
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active c.wt71##c.wt71
/*Output predicted conditional probability of quitting smoking for each individual*/
predict p_qsmk, pr
/*Generate nonstabilized weights as P(A=1|covariates) if A=1 and 1-P(A=1|covariates) if A=0*/
gen w=.
replace w=1/p_qsmk if qsmk==1
replace w=1/(1-p_qsmk) if qsmk==0
/*Check the mean of the weights; we expect it to be close to 2.0*/
summarize w
/*Fit marginal structural model in the pseudopopulation*/
/*Weights assigned using pweight = w*/
/*Robust standard errors using cluster() option where 'seqn' is the ID variable*/
regress wt82_71 qsmk [pweight=w], cluster(seqn)
Iteration 0: log likelihood = -893.02712
Iteration 1:
            log likelihood = -839.70016
Iteration 2: log likelihood = -838.45045
Iteration 3: log likelihood = -838.44842
Iteration 4: log likelihood = -838.44842
Logistic regression
                                          Number of obs
                                                               1,566
                                          LR chi2(18)
                                                               109.16
                                          Prob > chi2
                                                               0.0000
Log likelihood = -838.44842
                                          Pseudo R2
                                                               0.0611
      qsmk
                 Coef. Std. Err. z P>|z| [95% Conf. Interval]
  sex | -.5274782 .1540497 -3.42 0.001
                                                    -.82941
                                                             -.2255463
       race | -.8392636 .2100668 -4.00 0.000 -1.250987 -.4275404
        age | .1212052 .0512663 2.36 0.018
                                                  .0207251 .2216853
 c.age#c.age | -.0008246 .0005361 -1.54 0.124 -.0018753 .0002262
  education |
        1 | -.4759606 .2262238
                                   -2.10 0.035
                                                  -.9193511
                                                             -.0325701
        2 | -.5047361 .217597
                                  -2.32 0.020
                                                  -.9312184 -.0782538
        3 | -.3895288 .1914353
                                   -2.03
                                          0.042
                                                  -.7647351 -.0143226
        4 | -.4123596 .2772868 -1.49 0.137
                                                 -.9558318 .1311126
smokeinten~y | -.0772704 .0152499
                                   -5.07
                                                  -.1071596 -.0473812
                                          0.000
```

Ι

c.						
smokeinten~y#						
c.						
smokeinten~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
1						
smokeyrs	0735966	.0277775	-2.65	0.008	1280395	0191538
I						
c.smokeyrs#						
c.smokeyrs	.0008441	.0004632	1.82	0.068	0000637	.0017519
1						
exercise						
0	395704	.1872401	-2.11	0.035	7626878	0287201
1	0408635	.1382674	-0.30	0.768	3118627	.2301357
1						
active						
0	176784	.2149721	-0.82	0.411	5981215	. 2445535
1	1448395	.2111472	-0.69	0.493	5586806	.2690015
1						
wt71	0152357	.0263161	-0.58	0.563	0668144	.036343
1						
c.wt71#						
c.wt71	.0001352	.0001632	0.83	0.407	0001846	.000455
1						
_cons	-1.19407	1.398493	-0.85	0.393	-3.935066	1.546925

(1,566 missing values generated)

(403 real changes made)

(1,163 real changes made)

Variable	Obs	Mean	Std. Dev.	Min	Max
w	1,566	1.996284	1.474787	1.053742	16.70009

(sum of wgt is 3,126.18084549904)

Linear regression Number of obs = 1,566 F(1, 1565) = 42.81 Prob > F = 0.0000 R-squared = 0.0435 Root MSE = 8.0713

(Std. Err. adjusted for 1,566 clusters in seqn)

| Robust | Wt82\_71 | Coef. Std. Err. t P>|t| [95% Conf. Interval]

```
qsmk | 3.440535 .5258294 6.54 0.000 2.409131 4.47194

_cons | 1.779978 .2248742 7.92 0.000 1.338892 2.221065
```

- Estimating stabilized IP weights for Section 12.3
- Data from NHEFS

```
use ./data/nhefs-formatted, clear
/*Fit a logistic model for the denominator of the IP weights and predict the conditional probability of
logit qsmk sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity ///
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active c.wt71##c.wt71
predict pd_qsmk, pr
/*Fit a logistic model for the numerator of ip weights and predict Pr(A=1) */
logit qsmk
predict pn_qsmk, pr
/*Generate stabilized weights as f(A)/f(A/L)*/
gen sw a=.
replace sw_a=pn_qsmk/pd_qsmk if qsmk==1
replace sw_a=(1-pn_qsmk)/(1-pd_qsmk) if qsmk==0
/*Check distribution of the stabilized weights*/
summarize sw_a
/*Fit marginal structural model in the pseudopopulation*/
regress wt82_71 qsmk [pweight=sw_a], cluster(seqn)
/*********************
FINE POINT 12.2
Checking positivity
***********************************
/*Check for missing values within strata of covariates, for example: */
tab age qsmk if race==0 & sex==1 & wt82!=.
tab age qsmk if race==1 & sex==1 & wt82!=.
Iteration 0:
              log\ likelihood = -893.02712
Iteration 1:
              log\ likelihood = -839.70016
Iteration 2:
              log\ likelihood = -838.45045
Iteration 3: log likelihood = -838.44842
Iteration 4:
             log likelihood = -838.44842
```

Logistic regression	Number of obs	=	1,566
	LR chi2(18)	=	109.16
	Prob > chi2	=	0.0000
Log likelihood = -838.44842	Pseudo R2	=	0.0611

qsmk	Coef.	Std. Err.	Z	P> z	[95% Conf.	Interval]
sex	5274782	. 1540497	-3.42	0.001	82941	2255463
race	8392636	.2100668	-4.00	0.000	-1.250987	4275404
age	.1212052	.0512663	2.36	0.018	.0207251	.2216853
ا   c.age#c.age 	0008246	.0005361	-1.54	0.124	0018753	.0002262
education						
1	4759606	.2262238	-2.10	0.035	9193511	0325701
2	5047361	.217597	-2.32	0.020	9312184	0782538
3 l	3895288	.1914353	-2.03	0.042	7647351	0143226
4	4123596	.2772868	-1.49	0.137	9558318	.1311126
smokeinten~y   	0772704	.0152499	-5.07	0.000	1071596	0473812
c.						
smokeinten~y#						
c.						
smokeinten~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
smokeyrs   	0735966	.0277775	-2.65	0.008	1280395	0191538
c.smokeyrs#						
c.smokeyrs		.0004632	1.82	0.068	0000637	.0017519
exercise						
0 1	395704	.1872401	-2.11	0.035	7626878	0287201
1 I		.1382674	-0.30	0.768	3118627	.2301357
. !						
active						
0	176784	.2149721	-0.82	0.411	5981215	. 2445535
1	1448395	.2111472	-0.69	0.493	5586806	.2690015
wt71   	0152357	.0263161	-0.58	0.563	0668144	.036343
c.wt71#						
c.wt71		.0001632	0.83	0.407	0001846	.000455
_cons	-1.19407	1.398493	-0.85	0.393	-3.935066	1.546925

Iteration 0: log likelihood = -893.02712
Iteration 1: log likelihood = -893.02712

Logistic regression Number of obs = 1,566

LR chi2(0) = 0.00

Prob > chi2 =

Log likelihood = -893.02712 Pseudo R2 = 0.0000

-			[95% Conf.	Interval]
_cons			-1.173114	946529

(1,566 missing values generated)

(403 real changes made)

(1,163 real changes made)

Variable		Obs Me	ean Std.	Dev. N	Min Max
sw_a	   1,!	566 .9988	 444 .2882	233 .33124	489 4.297662

(sum of wgt is 1,564.19025221467)

Linear regression Number of obs = 1,566

F(1, 1565) = 42.81 Prob > F = 0.0000 R-squared = 0.0359 Root MSE = 7.7972

(Std. Err. adjusted for 1,566 clusters in seqn)

   wt82_71		Robust Std. Err.			Interval]
qsmk	3.440535 1.779978	.5258294	0.000	2.409131 1.338892	4.47194 2.221065

-----

		quit smokin	ng between		
		baseline	and 1982		
O		No smokin	0		
 25	•	24	3	•	27
26	1	1/1	5	1	10

+		+	
74	0	1	1
72	2	2	4
71	0	1	1
70 l	2	1	3
69 I	6	2	8
67	2	0	2
66 I	4	0	4
65	3	2	5
64	7	1	8
63	3	3	6
61   62	5 6	2   5	7 11
60   61	5	4	9
59	5	4	9
58   50	8	4	12
57   50	9	2	11
56	8	7	15
55   56	9	4	13 15
54	17	9	26
53	11	4	15
52	11	3	14
51	9	3	12
50	18	4	22
49	11	3	14
48	19	4	23
47	19	4	23
46	19	4	23
45	12	5	17
44	9	4	13
43	14	3	17
42	16	3	19
41	13	3	16
40	10	4	14
39	19	4	23
38	6	2	8
37	14	1	15
36 l	13	3	16
35	16	5	21
34	22	5 l	27
33	12	3	15
32	14	7	21
31	11	5	16
30 l	14	5	19
29 I	15	4	19
28	20	5 I	25
27	18	2	20

	quit smokin	ng between and 1982	
age	No smokin	Smoking c	Total
25	3	1	4
26	] 3	0	] 3
28	] 3	1	1 4
29	1	0	1
30	4	0	1 4
31	] 3	0	] 3
32	8	0	8
33	1 2	0	1 2
34	1 2	1	] 3
35	] 3	0	] 3
36	J 5	0	J 5
37	] 3	1	4
38	4	2	1 6
39	1	1	1 2
40	1 2	2	1 4
41	] 3	0	] 3
42	] 3	0	] 3
43	4	2	l 6
44	] 3	0	3
45	1	3	1 4
46	J 5	0	J 5
47	] 3	0	3
48	1 4	0	1 4
49	1	1	1 2
50	1 2	0	1 2
51	1 4	0	1 4
52	1	0	1
53	1 2	0	1 2
54	1 2	0	1 2
55	] 3	0	] 3
56	1 2	1	] 3
57	1 2	1	] 3
61	1	1	2
67	1	0	1
68	1	0	1
69	1 2	0	1 2
70	0	1	1
Total	97	 19	116

# Program 12.4

 $\bullet$  Estimating the parameters of a marginal structural mean model with a continuous treatment Data from NHEFS

#### • Section 12.4

```
use ./data/nhefs-formatted, clear
* drop sw_a
/*Analysis restricted to subjects reporting <=25 ciq/day at baseline: N = 1162*/
keep if smokeintensity <=25</pre>
/*Fit a linear model for the denominator of the IP weights and calculate the mean expected smoking inte
regress smkintensity82_71 sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity /
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active c.wt71##c.wt71
quietly predict p_den
/*Generate the denisty of the denomiator expectation using the mean expected smoking intensity and the
/*Note: The regress command in STATA saves the root mean squared error for the immediate regression as
gen dens_den = normalden(smkintensity82_71, p_den, e(rmse))
/*Fit a linear model for the numerator of ip weights, calculate the mean expected value, and generate t
quietly regress smkintensity82_71
quietly predict p_num
gen dens_num = normalden( smkintensity82_71, p_num, e(rmse))
/*Generate the final stabilized weights from the estimated numerator and denominator, and check the wei
gen sw_a=dens_num/dens_den
summarize sw_a
/*Fit a marginal structural model in the pseudopopulation*/
regress wt82_71 c.smkintensity82_71##c.smkintensity82_71 [pweight=sw_a], cluster(seqn)
/*Dutput the estimated mean Y value when smoke intensity is unchanged from baseline to 1982 */
lincom _b[_cons]
/*Output the estimated mean Y value when smoke intensity increases by 20 from baseline to 1982*/
lincom _b[_cons] + 20*_b[smkintensity82_71] +400*_b[c.smkintensity82_71#c.smkintensity82_71]
(404 observations deleted)
     Source | SS df MS Number of obs =
                                                              1,162
----- F(18, 1143) =
                                                               5.39
      Model | 9956.95654
                            18 553.164252 Prob > F
                                                         = 0.0000
   Residual | 117260.18 1,143 102.589834 R-squared
                                                        = 0.0783
------ Adj R-squared = 0.0638
      Total | 127217.137 1,161 109.575484 Root MSE
                                                        = 10.129
smkintens~71 | Coef. Std. Err. t P>|t| [95% Conf. Interval]
sex | 1.087021 .7425694 1.46 0.144 -.3699308
                                                             2.543973
```

race		.8434739	0.28	0.783	-1.422952	1.88691
age   	8099902	. 2555388	-3.17	0.002	-1.311368	3086124
c.age#c.age   	.0066545	.0026849	2.48	0.013	.0013865	.0119224
education						
1 l	1.508097	1.184063	1.27	0.203	8150843	3.831278
2	2.02692	1.133772	1.79	0.074	1975876	4.251428
3	2.240314	1.022556	2.19	0.029	.2340167	4.246611
4	2.528767	1.44702	1.75	0.081	3103458	5.36788
smokeinten~y	3589684	.2246653	-1.60	0.110	799771	.0818342
c.						
smokeinten~y#						
c.						
smokeinten~y	.0019582	.0085753	0.23	0.819	0148668	.0187832
smokeyrs	.3857088	.1416765	2.72	0.007	.1077336	.6636841
c.smokeyrs#						
c.smokeyrs		.0023837	-2.30	0.022	0101641	0008101
exercise						
0 1		.9080421	2.20	0.028	.215288	3.778521
1	.988812	.6929239	1.43	0.154	3707334	2.348357
I						
active						
0	.8451341	1.098573	0.77	0.442	-1.310312	3.000581
1	.800114	1.08438	0.74	0.461	-1.327485	2.927712
wt71	0656882	.136955	-0.48	0.632	3343996	.2030232
c.wt71#						
c.wt71#		.000877	0.65	0.515	0011496	.0022918
			0.00	0.020		
_cons	16.86761	7.109189	2.37	0.018	2.91909	30.81614

Variable	l	0bs	Mean	Std.	Dev.	Min	Max
sw_a	+   1	,162 .9	9968057	.3222	2937 .193	 38336	5.102339

(sum of wgt is 1,158.28818286955)

Linear regression Number of obs = 1,162  $F(2, 1161) = 12.75 \\ Prob > F = 0.0000 \\ R-squared = 0.0233$ 

Root MSE = 7.7864

(Std. Err. adjusted for 1,162 clusters in seqn)

   wt82_71 		Robust Std. Err.	t	P> t	[95% Conf.	Interval]
smkintens~71		.0315762	-3.45	0.001	1709417	0470361
c.   smkintens~71#  c.						
smkintens~71	.0026949	.0024203	1.11	0.266	0020537	.0074436
_cons	2.004525	.295502	6.78	0.000	1.424747	2.584302

(1) cons = 0

wt82_71			[95% Conf.	Interval]
			1.424747	2.584302

( 1) 20\*smkintensity82\_71 + 400\*c.smkintensity82\_71#c.smkintensity82\_71 +
 \_cons = 0

wt82_71		• • •	[95% Conf.	Interval]
			-1.668554	3.474001

- $\bullet\,$  Estimating the parameters of a marginal structural logistic model
- Data from NHEFS
- Section 12.4

```
/*Provisionally ignore subjects with missing values for follow-up weight*/
/*Sample size after exclusion: N = 1566*/
drop if wt82==.
```

```
/*Estimate the stabilized weights for quitting smoking as in PROGRAM 12.3*/
/*Fit a logistic model for the denominator of the IP weights and predict the conditional probability of
logit qsmk sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity ///
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active c.wt71##c.wt71
predict pd_qsmk, pr
/*Fit a logistic model for the numerator of ip weights and predict Pr(A=1) */
logit qsmk
predict pn_qsmk, pr
/*Generate stabilized weights as f(A)/f(A|L)*/
gen sw_a=.
replace sw_a=pn_qsmk/pd_qsmk if qsmk==1
replace sw_a=(1-pn_qsmk)/(1-pd_qsmk) if qsmk==0
summarize sw_a
/*Fit marginal structural model in the pseudopopulation*/
/*NOTE: Stata has two commands for logistic regression, logit and logistic*/
/*Using logistic allows us to output the odds ratios directly*/
/*We can also output odds ratios from the logit command using the or option (default logit output is re
logistic death qsmk [pweight=sw_a], cluster(seqn)
(63 observations deleted)
Iteration 0: log likelihood = -893.02712
Iteration 1: log likelihood = -839.70016
Iteration 2: log likelihood = -838.45045
Iteration 3: log likelihood = -838.44842
Iteration 4: log likelihood = -838.44842
Logistic regression
                                          Number of obs =
                                                               1,566
                                          LR chi2(18)
                                                                109.16
                                          Prob > chi2
                                                         =
                                                                0.0000
Log likelihood = -838.44842
                                          Pseudo R2
                                                                0.0611
      qsmk
                Coef. Std. Err. z P>|z|
                                                  [95% Conf. Interval]
______
                                                   -.82941 -.2255463
       sex | -.5274782 .1540497 -3.42 0.001
       race | -.8392636 .2100668 -4.00 0.000 -1.250987 -.4275404
        age | .1212052 .0512663
                                   2.36 0.018 .0207251
                                                             .2216853
 c.age#c.age | -.0008246 .0005361 -1.54 0.124 -.0018753 .0002262
  education |
        1 | -.4759606 .2262238 -2.10 0.035
                                                  -.9193511 -.0325701
        2 | -.5047361 .217597 -2.32 0.020 -.9312184 -.0782538
        3 | -.3895288 .1914353 -2.03 0.042
                                                  -.7647351 -.0143226
        4 | -.4123596 .2772868
                                   -1.49 0.137
                                                  -.9558318 .1311126
```

smokeinten~y	0772704	.0152499	-5.07	0.000	1071596	0473812
1						
c.						
smokeinten~y#						
c.						
smokeinten~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
1						
smokeyrs   	0735966	.0277775	-2.65	0.008	1280395	0191538
c.smokeyrs#						
c.smokeyrs	.0008441	.0004632	1.82	0.068	0000637	.0017519
1						
exercise						
0	395704	.1872401	-2.11	0.035	7626878	0287201
1	0408635	.1382674	-0.30	0.768	3118627	.2301357
1						
active						
0	176784	.2149721	-0.82	0.411	5981215	. 2445535
1	1448395	.2111472	-0.69	0.493	5586806	.2690015
1						
wt71	0152357	.0263161	-0.58	0.563	0668144	.036343
1						
c.wt71#						
c.wt71	.0001352	.0001632	0.83	0.407	0001846	.000455
1						
_cons	-1.19407	1.398493	-0.85	0.393	-3.935066	1.546925

Iteration 0: log likelihood = -893.02712
Iteration 1: log likelihood = -893.02712

(1,566 missing values generated)

(403 real changes made)

(1,163 real changes made)

	Obs					Ma	ax
	1,566					4.29766	32
Logistic regre	ession			Number	of obs	=	1,566
Togibule regi	3551011			Wald ch			0.04
				Prob >			
Log pseudolike	elihood = -749	0.11596		Pseudo		=	
		(Std. E	Err. adjus		-		-
	 I	Robust					
death	   Odds Ratio					Conf. I	[nterval]
-	1.030578   .2252711	.1621842	0.19	0.848	.7570		1.402931 .2629781
_cons	1 .2252/11	.0111002	-10.00	0.000	.1923	9101	.2029701

Note: \_cons estimates baseline odds.

- Assessing effect modification by sex using a marginal structural mean model
- Data from NHEFS
- Section 12.5

```
wse ./data/nhefs, clear

* drop pd_qsmk pn_qsmk sw_a

/*Check distribution of sex*/
tab sex

/*Fit logistc model for the denominator of IP weights, as in PROGRAM 12.3 */
logit qsmk sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity ///
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active c.wt71##c.wt71
predict pd_qsmk, pr

/*Fit logistic model for the numerator of IP weights, no including sex */
logit qsmk sex
predict pn_qsmk, pr

/*Generate IP weights as before*/
gen sw_a=.
replace sw_a=pn_qsmk/pd_qsmk if qsmk=1
replace sw_a=(1-pn_qsmk)/(1-pd_qsmk) if qsmk==0
```

#### summarize sw a

/\*Fit marginal structural model in the pseudopopulation, including interaction term between quitting sm regress wt82\_71 qsmk##sex [pw=sw\_a], cluster(seqn)

sex	Freq.	Percent	Cum.
0	799   830	49.05 50.95	49.05 100.00
Total	1,629	100.00	

Iteration 0: log likelihood = -938.14308
Iteration 1: log likelihood = -884.53806
Iteration 2: log likelihood = -883.35064
Iteration 3: log likelihood = -883.34876
Iteration 4: log likelihood = -883.34876

Logistic regression Number of obs = 1,629 LR chi2(18) = 109.59 Prob > chi2 = 0.0000 Log likelihood = -883.34876 Pseudo R2 = 0.0584

qsmk | Coef. Std. Err. z P>|z| [95% Conf. Interval] sex | -.5075218 .1482316 -3.42 0.001 -.7980505 -.2169932 race | -.8502312 .2058722 -4.13 0.000 -1.253733 -.4467292 age | .1030132 .0488996 2.11 0.035 .0071718 .1988547 c.age#c.age | -.0006052 .0005074 -1.190.233 -.0015998 .0003893 education | 1 | -.3796632 .2203948 -1.72 0.085 .0523026 -.811629 2 | -.4779835 .2141771 -2.23 0.026 -.8977629 -.0582041 3 | -.3639645 .1885776 -1.93 0.054 -.7335698 .0056409 4 | -.4221892 -.9547574 .2717235 -1.55 0.120 .110379 smokeinten~y | -.0651561 -4.41 0.000 -.0940831 -.0362292 .0147589 c. | smokeinten~y#| c. | smokeinten~y | .0008461 .0002758 3.07 0.002 .0003054 .0013867 smokeyrs | -.0733708 .0269958 -2.720.007 -.1262816 -.02046

c.smokeyrs#						
c.smokeyrs	.0008384	.0004435	1.89	0.059	0000307	.0017076
1						
exercise						
0	3550517	.1799293	-1.97	0.048	7077067	0023967
1	06364	.1351256	-0.47	0.638	3284812	.2012013
1						
active						
0	0683123	.2087269	-0.33	0.743	4774095	.3407849
1 l	057437	.2039967	-0.28	0.778	4572632	.3423892
1						
wt71	0128478	.0222829	-0.58	0.564	0565214	.0308258
1						
c.wt71#						
c.wt71	.0001209	.0001352	0.89	0.371	000144	.0003859
1						
_cons	-1.185875	1.263142	-0.94	0.348	-3.661588	1.289838

Iteration 0: log likelihood = -938.14308
Iteration 1: log likelihood = -933.49896
Iteration 2: log likelihood = -933.49126
Iteration 3: log likelihood = -933.49126

Logistic regression Number of obs = 1,629 LR chi2(1) = 9.30 Prob > chi2 = 0.0023 Log likelihood = -933.49126 Pseudo R2 = 0.0050

qsmk | Coef. Std. Err. z P>|z| [95% Conf. Interval]

sex | -.3441893 .1131341 -3.04 0.002 -.565928 -.1224506

\_cons | -.8634417 .0774517 -11.15 0.000 -1.015244 -.7116391

(1,629 missing values generated)

(428 real changes made)

(1,201 real changes made)

Variable	l Obs	Mean	Std. Dev.	Min	Max
	+				
sw_a	1,629	.9991318	.2636164	.2901148	3.683352

Linear regression	Number of obs	=	1,566
	F(3, 1565)	=	16.31
	Prob > F	=	0.0000
	R-squared	=	0.0379
	Root MSE	=	7.8024

(Std. Err. adjusted for 1,566 clusters in seqn)

   wt82_71 		Robust Std. Err.	t	P> t	[95% Conf.	Interval]
qsmk#sex   1 1	161224	1.036143	-0.16	0.876	-2.1936	1.871152
_cons	1.759045	.3102511	5.67	0.000	1.150494	2.367597

- Estimating IP weights to adjust for selection bias due to censoring
- Data from NHEFS
- Section 12.6

```
use ./data/nhefs, clear
/*Analysis including all individuals regardless of missing wt82 status: N=1629*/
/*Generate censoring indicator: C = 1 if wt82 missing*/
gen byte cens = (wt82 == .)
/*Check distribution of censoring by quitting smoking and baseline weight*/
tab cens qsmk, column
bys cens: summarize wt71
/*Fit logistic regression model for the denominator of IP weight for A*/
logit qsmk sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity ///
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active c.wt71##c.wt71
predict pd_qsmk, pr
/*Fit logistic regression model for the numerator of IP weights for A*/
logit qsmk
predict pn_qsmk, pr
/*Fit logistic regression model for the denominator of IP weights for C, including quitting smoking*/
logit cens qsmk sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity ///
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active c.wt71##c.wt71
```

```
predict pd_cens, pr
/*Fit logistic regression model for the numerator of IP weights for C, including quitting smoking */
logit cens qsmk
predict pn_cens, pr
/*Generate the stabilized weights for A (sw_a)*/
gen sw_a=.
replace sw_a=pn_qsmk/pd_qsmk if qsmk==1
replace sw_a=(1-pn_qsmk)/(1-pd_qsmk) if qsmk==0
/*Generate the stabilized weights for C (sw_c)*/
/*NOTE: the conditional probability estimates generated by our logistic models for C represent the cond
/*We want weights for the conditional probability of bing uncensored, Pr(C=0|A,L)*/
gen sw c=.
replace sw_c=(1-pn_cens)/(1-pd_cens) if cens==0
/*Generate the final stabilized weights and check distribution*/
gen sw=sw_a*sw_c
summarize sw
/*Fit marginal structural model in the pseudopopulation*/
regress wt82_71 qsmk [pw=sw], cluster(seqn)
| Key
| frequency |
| column percentage |
+----+
         | quit smoking between
        baseline and 1982
    cens | 0 1 |
-----
                     403 | 1,566
       0 | 1,163
       96.84
                     94.16 |
      1 |
            38
                      25 |
                                63
              3.16 5.84 |
       3.87
_____
    Total |
            1,201
                      428 |
                               1,629
       | 100.00 100.00 | 100.00
-> cens = 0
   Variable | Obs Mean Std. Dev. Min Max
```

wt71 | 1,566 70.83092 15.3149 39.58 151.73

-----

-> cens = 1

Variable	0bs	Mean	Std. Dev.	Min	Max
wt71	63	76.55079	23.3326	36.17	169.19

Iteration 0: log likelihood = -938.14308
Iteration 1: log likelihood = -884.53806
Iteration 2: log likelihood = -883.35064
Iteration 3: log likelihood = -883.34876
Iteration 4: log likelihood = -883.34876

Logistic regression Number of obs = 1,629 LR chi2(18) = 109.59 Prob > chi2 = 0.0000 Log likelihood = -883.34876 Pseudo R2 = 0.0584

qsmk	Coef.	Std. Err.	z	P> z	[95% Conf.	Interval]
sex	5075218	.1482316	-3.42	0.001	7980505	2169932
race	8502312	.2058722	-4.13	0.000	-1.253733	4467292
age	.1030132	.0488996	2.11	0.035	.0071718	.1988547
1						
c.age#c.age	0006052	.0005074	-1.19	0.233	0015998	.0003893
1						
education						
1	3796632	.2203948	-1.72	0.085	811629	.0523026
2	4779835	.2141771	-2.23	0.026	8977629	0582041
3	3639645	.1885776	-1.93	0.054	7335698	.0056409
4	4221892	.2717235	-1.55	0.120	9547574	.110379
1						
smokeinten~y	0651561	.0147589	-4.41	0.000	0940831	0362292
1						
c.						
smokeinten~y#						
c.						
smokeinten~y	.0008461	.0002758	3.07	0.002	.0003054	.0013867
1						
smokeyrs	0733708	.0269958	-2.72	0.007	1262816	02046
c.smokeyrs#						
c.smokeyrs	.0008384	.0004435	1.89	0.059	0000307	.0017076
I						

exercise						
0	3550517	.1799293	-1.97	0.048	7077067	0023967
1	06364	.1351256	-0.47	0.638	3284812	.2012013
I						
active						
0	0683123	.2087269	-0.33	0.743	4774095	.3407849
1 l	057437	.2039967	-0.28	0.778	4572632	.3423892
I						
wt71	0128478	.0222829	-0.58	0.564	0565214	.0308258
I						
c.wt71#						
c.wt71	.0001209	.0001352	0.89	0.371	000144	.0003859
I						
_cons	-1.185875	1.263142	-0.94	0.348	-3.661588	1.289838
Iteration 0: Iteration 1:	log likeliho					
Logistic regre	ession				of obs =	
				LR chi2		0.00
		,		Prob >		
Log likelihood	l = -938.14308	3		Prob > Pseudo		0.0000
Log likelihood	d = -938.14308	3				
			z	Pseudo		0.0000
qsmk	Coef.	Std. Err.		Pseudo P> z	R2 =	0.0000  Interval]
qsmk	Coef.  -1.031787  log likeliho log likeliho log likeliho log likeliho log likeliho	Std. Err0562947	-18.33 	Pseudo P> z  0.000	R2 =  [95% Conf.  -1.142122	0.0000  Interval]  9214511
qsmk	Coef.  -1.031787  log likeliho log likeliho log likeliho log likeliho log likeliho	Std. Err0562947	-18.33 	Pseudo P> z  0.000  Number	R2 =  [95% Conf.  -1.142122	0.0000  Interval]
qsmk	Coef.  -1.031787  log likeliho log likeliho log likeliho log likeliho log likeliho	Std. Err.  .0562947  .000 = -266.6  .000 = -238.4  .000 = -232.8  .000 = -232.6  .000 = -232.6  .000 = -232.6	-18.33 	Pseudo P> z  0.000  Number	R2 =  [95% Conf.  -1.142122   of obs = 2(19) = chi2 =	0.0000  Interval]  9214511 

qsmk	.5168674	.2877162	1.80	0.072	0470459	1.080781
sex	.0573131	.3302775	0.17	0.862	590019	.7046452
race	0122715	.4524888	-0.03	0.978	8991332	.8745902
age	2697293	.1174647	-2.30	0.022	4999559	0395027
	l					
c.age#c.age	.0028837	.0011135	2.59	0.010	.0007012	.0050661
	l					
education	l					
1	.3823818	.5601808	0.68	0.495	7155523	1.480316
2	0584066	.5749586	-0.10	0.919	-1.185305	1.068491
3	.2176937	.5225008	0.42	0.677	8063891	1.241776
4	.5208288	.6678735	0.78	0.435	7881792	1.829837
	l					
smokeinten~y	.0157119	.0347319	0.45	0.651	0523614	.0837851
	l					
с.	l					
smokeinten~y#	l					
С.	l					
smokeinten~y	0001133	.0006058	-0.19	0.852	0013007	.0010742
	l					
smokeyrs	.0785973	.0749576	1.05	0.294	0683169	.2255116
	l					
c.smokeyrs#	l					
c.smokeyrs	0005569	.0010318	-0.54	0.589	0025791	.0014653
exercise						
0	.583989	.3723133	1.57	0.117	1457317	1.31371
1	3874824	.3439133	-1.13	0.260	-1.06154	.2865754
	1					
active	1					
0	7065829	.3964577	-1.78	0.075	-1.483626	.0704599
1	9540614	.3893181	-2.45	0.014	-1.717111	1910119
wt71	0878871	.0400115	-2.20	0.028	1663082	0094659
	l					
c.wt71#	l					
c.wt71	.0006351	.0002257	2.81	0.005	.0001927	.0010775
	l					
_cons	3.754678	2.651222	1.42	0.157	-1.441622	8.950978

```
Iteration 0: log likelihood = -266.67873 Iteration 1: log likelihood = -264.00252 Iteration 2: log likelihood = -263.88028 Iteration 3: log likelihood = -263.88009 Iteration 4: log likelihood = -263.88009
```

Logistic regression Number of obs = 1,629 LR chi2(1) = 5.60 Prob > chi2 = 0.0180 Log likelihood = -263.88009 Pseudo R2 = 0.0105

cens | Coef. Std. Err. z P>|z| [95% Conf. Interval]

qsmk | .6411113 .2639262 2.43 0.015 .1238255 1.158397

cons | -3.421172 .1648503 -20.75 0.000 -3.744273 -3.098071

(1,629 missing values generated)

(428 real changes made)

(1,201 real changes made)

(1,629 missing values generated)

(1,566 real changes made)

(63 missing values generated)

Variable	l Obs	Mean	Std. Dev.	Min	Max
SW	+   1,566	.9962351	. 2819583	. 3546469	4.093113

(sum of wgt is 1,560.10419079661)

(Std. Err. adjusted for 1,566 clusters in seqn)

# 13. Standardization and the parametric G-formula: Stata

#### Program 13.1

- Estimating the mean outcome within levels of treatment and confounders: Data from NHEFS
- Section 13.2

```
use ./data/nhefs-formatted, clear

/* Estimate the the conditional mean outcome within strata of quitting
smoking and covariates, among the uncensored */
glm wt82_71 qsmk sex race c.age##c.age ib(last).education ///
    c.smokeintensity##c.smokeintensity c.smokeyrs##c.smokeyrs ///
    ib(last).exercise ib(last).active c.wt71##c.wt71 ///
    qsmk##c.smokeintensity
predict meanY
summarize meanY

/*Look at the predicted value for subject ID = 24770*/
list meanY if seqn == 24770

/*Observed mean outcome for comparison */
summarize wt82_71
```

note: 1.qsmk omitted because of collinearity

note: smokeintensity omitted because of collinearity

Iteration 0:	log likeliho	ood = -5328.	5765			
Generalized li	near models			No. o	f obs =	1,566
Optimization					ual df =	
opoimización	• 1.2				parameter =	· ·
Deviance	= 82763.0	)2862			) Deviance =	
Pearson	= 82763.0				) Pearson =	
1 001 001	02.001			(=/ ==	, 10012011	00.000
Variance funct	ion: V(u) = 1	L		ΓGaus	sian]	
Link function					tity]	
	<b>3</b> · ·				<b>y</b> -	
				AIC	=	6.832154
Log likelihood	= -5328.57	76456		BIC	=	71397.58
I		MIO				
wt82_71	Coef.	Std. Err.	z	P> z	[95% Conf.	Interval]
qsmk	2.559594	.8091486	3.16	0.002	.973692	4.145496
sex	-1.430272	.4689576	-3.05	0.002	-2.349412	5111317
race	.5601096	.5818888	0.96	0.336	5803714	1.700591
age	.3596353	.1633188	2.20	0.028	.0395364	.6797342
c.age#c.age	006101	.0017261	-3.53	0.000	0094841	0027178
I						
education						
1	.194977	.7413692	0.26	0.793	-1.25808	1.648034
2	.9854211	.7012116	1.41	0.160	3889285	2.359771
3	.7512894	.6339153	1.19	0.236	4911617	1.993741
4	1.686547	.8716593	1.93	0.053	0218744	3.394967
I						
smokeinten~y	.0491365	.0517254	0.95	0.342	0522435	.1505165

smokeinten~y#|

c. |

1 | -.0579374

c. | .000938 smokeinten~y | -.0009907 -1.06 0.291 -.0028292 .0008479 smokeyrs | .1343686 .0917122 1.47 0.143 -.045384 .3141212 c.smokeyrs#| c.smokeyrs | -.0018664 .0015437 -1.21 0.227 -.0048921 .0011592 exercise | -0.63 0.527 0 | -.3539128 .5588587 -1.449256 .7414301

.4316468

132

0.893

-.9039497

.7880749

-0.13

```
active |
       0 | .2613779 .6845577
                              0.38
                                     0.703 -1.08033
                                                      1.603086
       1 | -.6861916 .6739131
                                     0.309
                               -1.02
                                            -2.007037
                                                       .6346539
      wt71 |
            .0455018
                     .0833709
                              0.55
                                    0.585
                                            -.1179022
                                                       .2089058
    c.wt71#|
    c.wt71 | -.0009653
                    .0005247
                               -1.84 0.066
                                            -.0019937
                                                       .0000631
      qsmk |
Smoking c.. |
                 0 (omitted)
smokeinten~y |
                 0 (omitted)
      qsmk#|
       c. |
smokeinten~y |
Smoking c.. |
            .0466628 .0351448 1.33 0.184 -.0222197 .1155453
     6.911444
```

(option mu assumed; predicted mean wt82\_71)

Variable	Obs	Mean	Std. Dev.	Min	Max
	1,566	2.6383	3.034683	-10.87582	9.876489
+   mea 	mY				
Variable	Obs	Mean	Std. Dev.	Min	Max
wt82_71	1,566	2.6383	7.879913	-41.28047	48.53839

- Standardizing the mean outcome to the baseline confounders
- Data from Table 2.2
- Section 13.3

```
clear
input str10 ID L A Y

"Rheia" 0 0 0

"Kronos" 0 0 1

"Demeter" 0 0 0

"Hades" 0 0 0
```

```
"Hestia" 0 1 0
"Poseidon" 0 1 0
"Hera"
          0 1 0
"Zeus"
           0 1 1
"Artemis" 1 0 1
"Apollo"
           1 0 1
"Leto"
          1 0 0
"Ares"
           1 1 1
"Athena" 1 1 1
"Hephaestus" 1 1 1
"Aphrodite" 1 1 1
"Cyclope"
           1 1 1
"Persephone" 1 1 1
"Hermes"
           1 1 0
"Hebe"
           1 1 0
"Dionysus" 1 1 0
end
/* i. Data set up for standardization:
 - create 3 copies of each subject first,
 - duplicate the dataset and create a variable `interv` which indicates
which copy is the duplicate (interv =1) */
expand 2, generate(interv)
/* Next, duplicate the original copy (interv = 0) again, and create
another variable 'interv2' to indicate the copy */
expand 2 if interv == 0, generate(interv2)
/* Now, change the value of 'interv' to -1 in one of the copies so that
there are unique values of interv for each copy */
replace interv = -1 if interv2 ==1
drop interv2
/* Check that the data has the structure you want:
 - there should be 1566 people in each of the 3 levels of interv*/
tab interv
/* Two of the copies will be for computing the standardized result
for these two copies (interv = 0 and interv = 1), set the outcome to
missing and force qsmk to either 0 or 1, respectively.
You may need to edit this part of the code for your outcome and exposure variables */
replace Y = . if interv != -1
replace A = 0 if interv == 0
replace A = 1 if interv == 1
/* Check that the data has the structure you want:
for interv = -1, some people quit and some do not;
for interv = 0 or 1, noone quits or everyone quits, respectively */
by interv, sort: summarize A
```

```
*ii.Estimation in original sample*
*Now, we do a parametric regression with the covariates we want to adjust for*
*You may need to edit this part of the code for the variables you want.*
*Because the copies have missing Y, this will only run the regression in the original copy*
*The double hash between A & L creates a regression model with A and L and a product term between A and
regress Y A##L
*Ask Stata for expected values - Stata will give you expected values for all copies, not just the origin
predict predY, xb
*Now ask for a summary of these values by intervention*
*These are the standardized outcome estimates: you can subtract them to get the standardized difference
by interv, sort: summarize predY
*iii.OPTIONAL: Output standardized point estimates and difference*
*The summary from the last command gives you the standardized estimates*
*We can stop there, or we can ask Stata to calculate the standardized difference and display all the re
*The code below can be used as-is without changing any variable names*
*The option "quietly" asks Stata not to display the output of some intermediate calculations*
*You can delete this option if you want to see what is happening step-by-step*
quietly summarize predY if(interv == -1)
matrix input observe = (-1, r(mean)')
quietly summarize predY if(interv == 0)
matrix observe = (observe \0, r(mean)')
quietly summarize predY if(interv == 1)
matrix observe = (observe \1, r(mean)')
matrix observe = (observe \lambda, observe[3,2]-observe[2,2])
*Add some row/column descriptions and print results to screen*
matrix rownames observe = observed E(Y(a=0)) E(Y(a=1)) difference
matrix colnames observe = interv value
matrix list observe
*to interpret these results:*
*row 1, column 2, is the observed mean outcome value in our original sample*
*row 2, column 2, is the mean outcome value if everyone had not quit smoking*
*row 3, column 2, is the mean outcome value if everyone had quit smoking*
*row 4, column 2, is the mean difference outcome value if everyone had quit smoking compared to if ever
             ID
                                    Α
                                               Y
(20 observations created)
(20 observations created)
(20 real changes made)
     interv |
                   Freq.
                             Percent
                                            Cum.
```

+-						
-1	20	33.33	33.33			
0						
1	20	33.33	100.00			
+- Total	60	100.00				
(40 real chang	ges made, 40 t	o missing)				
(13 real chang	ges made)					
(7 real change	s made)					
-> interv = -1						
	Obs			Min	Max	
A				0	1	
-> interv = 0						
Variable	Obs	Mean	Std. Dev.	Min	Max	
A	20	0	0	0	0	
Variable	Obs	Mean	Std. Dev.	Min	Max	
+						
A	20	1	0	1	1	
	SS					
				F(3, 16) Prob > F		
				R-squared		
				Adj R-squared		
Total	5	19	.263157895	Root MSE	=	.51031
	Coef.	Std. Err.	t P>	t  [95% C		 cerval]
   A#L     1	-5.83e-17	. 4959325	-0.00 1.	000 -1.051	33 :	1.05133

<del>-</del>		0.98 0.342		)48
-> interv = -1				
Variable		Std. Dev.		
		. 209427		
-> interv = 0				
Variable		Std. Dev.	Max	
•		.209427	. 6666667	
Variable		Std. Dev.		
predY		.209427		

# observe[4,2]

	interv	value
observed	-1	.50000001
E(Y(a=0))	0	.50000001
E(Y(a=1))	1	.50000001
difference		0

# Program 13.3

- Standardizing the mean outcome to the baseline confounders:
- Data from NHEFS

#### • Section 13.3

```
use ./data/nhefs-formatted, clear
*i.Data set up for standardization: create 3 copies of each subject*
*first, duplicate the dataset and create a variable 'interv' which indicates which copy is the duplicat
expand 2, generate(interv)
*next, duplicate the original copy (interv = 0) again, and create another variable 'interv2' to indicat
expand 2 if interv == 0, generate(interv2)
*now, change the value of 'interv' to -1 in one of the copies so that there are unique values of interv
replace interv = -1 if interv2 ==1
drop interv2
*check that the data has the structure you want: there should be 1566 people in each of the 3 levels of
tab interv
*two of the copies will be for computing the standardized result*
*for these two copies (interv = 0 and interv = 1), set the outcome to missing and force qsmk to either
*you may need to edit this part of the code for your outcome and exposure variables*
replace wt82_71 = . if interv != -1
replace qsmk = 0 if interv == 0
replace qsmk = 1 if interv == 1
*check that the data has the structure you want: for interv = -1, some people quit and some do not; for
by interv, sort: summarize qsmk
*ii.Estimation in original sample*
*Now, we do a parametric regression with the covariates we want to adjust for*
*You may need to edit this part of the code for the variables you want.*
*Because the copies have missing wt82_71, this will only run the regression in the original copy*
regress wt82_71 qsmk sex race c.age##c.age ib(last).education c.smokeintensity##c.smokeintensity c.smok
*Ask Stata for expected values - Stata will give you expected values for all copies, not just the origin
predict predY, xb
*Now ask for a summary of these values by intervention*
*These are the standardized outcome estimates: you can subtract them to get the standardized difference
by interv, sort: summarize predY
/* iii.OPTIONAL: Output standardized point estimates and difference
- The summary from the last command gives you the
standardized estimates
- We can stop there, or we can ask Stata to calculate the
standardized difference and display all the results
in a simple table
- The code below can be used as-is without changing any
variable names
```

```
- The option `quietly` asks Stata not to display the output of
some intermediate calculations
- You can delete this option if you want to see what is
happening step-by-step */
quietly summarize predY if(interv == -1)
matrix input observe = (-1, r(mean)')
quietly summarize predY if(interv == 0)
matrix observe = (observe \0, r(mean)')
quietly summarize predY if(interv == 1)
matrix observe = (observe \1, r(mean)')
matrix observe = (observe \., observe[3,2]-observe[2,2])
* Add some row/column descriptions and print results to screen
matrix rownames observe = observed E(Y(a=0)) E(Y(a=1)) difference
matrix colnames observe = interv value
matrix list observe
/* To interpret these results:
- row 1, column 2, is the observed mean outcome value
in our original sample
- row 2, column 2, is the mean outcome value
if everyone had not quit smoking
- row 3, column 2, is the mean outcome value
if everyone had quit smoking
- row 4, column 2, is the mean difference outcome value
if everyone had quit smoking compared to if everyone
had not quit smoking */
/* Addition due to way Statamarkdown works
i.e. each code chunk is a separate Stata session */
mata observe = st_matrix("observe")
mata mata matsave ./data/observe observe, replace
*drop the copies*
drop if interv != -1
gen meanY_b =.
qui save ./data/nhefs_std, replace
(1,566 observations created)
(1,566 observations created)
```

(1,566 real changes made)

Cum.	Percent	Freq.	interv
33.33	33.33	1,566	-1
66.67	33.33	1,566	0
100.00	33.33	1,566	1

```
Total | 4,698 100.00
(3,132 real changes made, 3,132 to missing)
(403 real changes made)
(1,163 real changes made)
\rightarrow interv = -1
  Variable | Obs
                   Mean Std. Dev.
                                Min
______
    qsmk | 1,566 .2573436 .4373099
                                   0
-> interv = 0
  Variable | Obs Mean Std. Dev. Min Max
           1,566
                     0
                             0
                                    0
    qsmk |
-> interv = 1
  Variable | Obs
                  Mean Std. Dev. Min
                   1
                          0
    qsmk | 1,566
                                   1
   Source | SS df MS Number of obs = 1,566
------ F(20, 1545) =
                                            13.45
                   20 720.6279 Prob > F
    Model | 14412.558
                                         = 0.0000
  Residual | 82763.0286 1,545 53.5683033 R-squared
                                         = 0.1483
----- Adj R-squared = 0.1373
   Total | 97175.5866 1,565 62.0930266 Root MSE
                                             7.319
wt82_71 | Coef. Std. Err. t P>|t| [95% Conf. Interval]
                                   [95% Conf. Interval]
                                   .9724486
    qsmk | 2.559594 .8091486
                        3.16 0.002
                                            4.14674
     race | .5601096 .5818888 0.96 0.336 -.5812656 1.701485
     age | .3596353 .1633188 2.20 0.028 .0392854 .6799851
c.age#c.age | -.006101 .0017261 -3.53 0.000 -.0094868 -.0027151
```

education |

1   2   3   4	.9854211 .7512894	.7413692 .7012116 .6339153 .8716593	0.26 1.41 1.19 1.93	0.793 0.160 0.236 0.053	-1.259219 390006 4921358 0232138	1.649173 2.360848 1.994715 3.396307
smokeinten~y   	.0491365	.0517254	0.95	0.342	052323	.1505959
c.   smokeinten~y#						
c.						
smokeinten~y		.000938	-1.06	0.291	0028306	.0008493
smokeyrs	.1343686	.0917122	1.47	0.143	045525	.3142621
c.smokeyrs#						
c.smokeyrs		.0015437	-1.21	0.227	0048944	.0011616
exercise						
0 1		.5588587	-0.63	0.527	-1.450114	.7422889
1 I	0579374	.4316468	-0.13	0.893	904613	.7887381
- · 	700,00,1	. 1010100	0.120	0.000	1001010	
active						
0	.2613779	.6845577	0.38	0.703	-1.081382	1.604138
1 l	6861916	.6739131	-1.02	0.309	-2.008073	.6356894
I						
wt71	.0455018	.0833709	0.55	0.585	1180303	.2090339
I						
c.wt71#						
c.wt71	0009653	.0005247	-1.84	0.066	0019945	.0000639
I						
qsmk#						
c.						
smokeinten~y						
Smoking c	.0466628	.0351448	1.33	0.184	0222737	.1155993
60ng	-1.690608	4.388883	-0.39	0.700	-10.2994	6.918188
_cons	1.090000	4.300003	-0.39	0.700	-10.2994	0.910100

-----

<sup>-&</sup>gt; interv = -1

Variable		Mean		Max	
	1,566			9.876489	

-> interv = 0

Variable	Obs	Mean	Std. Dev.		Max
·			2.826271 -11		
Variable	0bs	Mean	Std. Dev.	Min	Max
predY	1,566	5.273587	2.920532 -9.	091126	11.0506

### observe[4,2]

	interv	value
observed	-1	2.6382998
E(Y(a=0))	0	1.7562131
E(Y(a=1))	1	5.2735873
difference	_	3.5173742

```
(saving observe[4,2])
file ./data/observe.mmat saved
```

(3,132 observations deleted)

(1,566 missing values generated)

# Program 13.4

- $\bullet$  Computing the 95% confidence interval of the standardized means and their difference: Data from NHEFS
- Section 13.3

```
*Run program 13.3 to obtain point estimates, and then the code below*
capture program drop bootstdz
```

```
program define bootstdz, rclass
use ./data/nhefs std, clear
preserve
* Draw bootstrap sample from original observations
bsample
/* Create copies with each value of qsmk in bootstrap sample.
First, duplicate the dataset and create a variable `interv` which
indicates which copy is the duplicate (interv =1)*/
expand 2, generate(interv_b)
/* Next, duplicate the original copy (interv = 0) again, and create
another variable `interv2` to indicate the copy*/
expand 2 if interv_b == 0, generate(interv2_b)
/* Now, change the value of interv to -1 in one of the copies so that
there are unique values of interv for each copy*/
replace interv_b = -1 if interv2_b ==1
drop interv2_b
/* Two of the copies will be for computing the standardized result.
For these two copies (interv = 0 and interv = 1), set the outcome to
missing and force qsmk to either 0 or 1, respectively*/
replace wt82_71 = . if interv_b != -1
replace qsmk = 0 if interv_b == 0
replace qsmk = 1 if interv_b == 1
* Run regression
regress wt82_71 qsmk sex race c.age##c.age ib(last).education ///
  c.smokeintensity##c.smokeintensity c.smokeyrs##c.smokeyrs ///
  ib(last).exercise ib(last).active c.wt71##c.wt71 ///
  qsmk#c.smokeintensity
/* Ask Stata for expected values.
Stata will give you expected values for all copies, not just the
original ones*/
predict predY_b, xb
summarize predY_b if interv_b == 0
return scalar boot_0 = r(mean)
summarize predY_b if interv_b == 1
return scalar boot_1 = r(mean)
return scalar boot_diff = return(boot_1) - return(boot_0)
drop meanY_b
restore
end
```

```
/* Then we use the `simulate` command to run the bootstraps as many
times as we want.
Start with reps(10) to make sure your code runs, and then change to
reps(1000) to generate your final CIs.*/
simulate EY_a0=r(boot_0) EY_a1 = r(boot_1) ///
  difference = r(boot_diff), reps(10) seed(1): bootstdz
/* Next, format the point estimate to allow Stata to calculate our
standard errors and confidence intervals*/
* Addition: read back in the observe matrix
mata mata matuse ./data/observe, replace
mata st_matrix("observe", observe)
matrix pe = observe[2..4, 2]'
matrix list pe
/* Finally, the bstat command generates valid 95% confidence intervals
under the normal approximation using our bootstrap results.
The default results use a normal approximation to calcutlate the
confidence intervals.
Note, n contains the original sample size of your data before censoring*/
bstat, stat(pe) n(1629)
      command: bootstdz
       EY_a0: r(boot_0)
       EY_a1: r(boot_1)
   difference: r(boot_diff)
Simulations (10)
---+-- 1 ---+-- 2 ---+-- 3 ---+-- 4 ---+-- 5
. . . . . . . . . .
(loading observe[4,2])
pe[1,3]
          r2
                    r3
c2 1.7562131 5.2735873 3.5173742
Bootstrap results
                                                Number of obs
                                                                         1,629
                                                Replications
                                                                            10
                 Observed Bootstrap
                                                              Normal-based
```

					[95% Conf.	
EY_a0	1.756213 5.273587	.2179372 .5759346	8.06 9.16	0.000	1.329064	2.183362 6.402398

# 14. G-estimation of Structural Nested Models: Stata

# Program 14.1

library(Statamarkdown)

- Ranks of extreme observations
- Data from NHEFS
- Section 14.4

```
/*For Stata 15 or later, first install the extremes function using this code:*/
* ssc install extremes

*Data preprocessing***

use ./data/nhefs, clear
gen byte cens = (wt82 == .)

/*Ranking of extreme observations*/
extremes wt82_71 seqn

/*Estimate unstabilized censoring weights for use in g-estimation models*/
glm cens qsmk sex race c.age##c.age ib(last).education ///
c.smokeintensity##c.smokeintensity c.smokeyrs##c.smokeyrs ///
ib(last).exercise ib(last).active c.wt71##c.wt71 ///
, family(binomial)
predict pr_cens
gen w_cens = 1/(1-pr_cens)
```

```
replace w_cens = . if cens == 1 /*observations with cens = 1 contribute to censoring models but not out
summarize w_cens
/*Analyses restricted to N=1566*/
drop if wt82 == .
summarize wt82_71
save ./data/nhefs-wcens, replace
 | obs: wt82_71 seqn |
 |-----|
 | 1329. -41.28046982 23321 |
 | 527. -30.50192161 13593 |
 | 1515. -30.05007421 24363 |
 | 204. -29.02579305 5412 |
 | 1067. -25.97055814 21897 |
 | 205. 34.01779932 5415 |
 | 1145. 36.96925111 22342 |
 | 64. 37.65051215 1769 |
 | 260. 47.51130337 6928 |
 | 1367. 48.53838568 23522 |
Iteration 0: \log likelihood = -292.45812
Iteration 1: log likelihood = -233.5099
Iteration 2: log likelihood = -232.68635
Iteration 3: log likelihood = -232.68
Iteration 4: log likelihood = -232.67999
Generalized linear models
                                           No. of obs = 1,629
                                           Residual df =
Optimization : ML
                                                               1,609
                                           Scale parameter =
Deviance = 465.3599898
                                           (1/df) Deviance = .2892231
Pearson
           = 1654.648193
                                           (1/df) Pearson = 1.028371
Variance function: V(u) = u*(1-u)
                                           [Bernoulli]
Link function : g(u) = ln(u/(1-u))
                                           [Logit]
                                           AIC
                                                       = .3102271
Log likelihood = -232.6799949
                                           BIC
                                                       = -11434.36
         OIM
      cens | Coef. Std. Err. z P>|z| [95% Conf. Interval]
```

qsmk	.5168674	.2877162	1.80	0.072	0470459	1.080781
sex	.0573131	.3302775	0.17	0.862	590019	.7046452
race	0122715	.4524888	-0.03	0.978	8991332	.8745902
age	2697293	.1174647	-2.30	0.022	4999558	0395027
I						
c.age#c.age	.0028837	.0011135	2.59	0.010	.0007012	.0050661
I						
education						
1 l	.3823818	.5601808	0.68	0.495	7155523	1.480316
2	0584066	.5749586	-0.10	0.919	-1.185305	1.068491
3	.2176937	.5225008	0.42	0.677	8063891	1.241776
4	.5208288	.6678735	0.78	0.435	7881792	1.829837
I						
smokeinten~y	.0157119	.0347319	0.45	0.651	0523614	.0837851
١						
c.						
smokeinten~y#						
c.						
smokeinten~y	0001133	.0006058	-0.19	0.852	0013007	.0010742
١						
smokeyrs	.0785973	.0749576	1.05	0.294	068317	.2255116
١						
c.smokeyrs#						
c.smokeyrs	0005569	.0010318	-0.54	0.589	0025791	.0014653
I						
exercise						
0	.583989	.3723133	1.57	0.117	1457317	1.31371
1 l	3874824	.3439133	-1.13	0.260	-1.06154	.2865753
I						
active						
0	7065829	.3964577	-1.78	0.075	-1.483626	.0704599
1	9540614	.3893181	-2.45	0.014	-1.717111	1910119
I						
wt71	0878871	.0400115	-2.20	0.028	1663082	0094659
I						
c.wt71#						
c.wt71	.0006351	.0002257	2.81	0.005	.0001927	.0010775
I						
_cons	3.754678	2.651222	1.42	0.157	-1.441622	8.950978

(option mu assumed; predicted mean cens)

## (63 real changes made, 63 to missing)

Variable	l Obs	Mean	Std. Dev.	Min	Max
w_cens	+   1,566	1.039197	.05646	1.001814	1.824624

(63 observations deleted)

file ./data/nhefs-wcens.dta saved

## Program 14.2

- G-estimation of a 1-parameter structural nested mean model
- Brute force search
- Data from NHEFS
- Section 14.5

```
use ./data/nhefs-wcens, clear
/*Generate test value of Psi = 3.446*/
gen psi = 3.446
/*Generate H(Psi) for each individual using test value of Psi and their own values of weight change and
gen Hpsi = wt82_71 - psi * qsmk
/*Fit a model for smoking status, given confounders and H(Psi) value, with censoring weights and displa
logit qsmk sex race c.age##c.age ib(last).education ///
  c.smokeintensity##c.smokeintensity c.smokeyrs##c.smokeyrs ///
  ib(last).exercise ib(last).active c.wt71##c.wt71 Hpsi ///
  [pw = w_cens], cluster(seqn)
di _b[Hpsi]
/*G-estimation*/
/*Checking multiple possible values of psi*/
cap noi drop psi Hpsi
local seq_start = 2
local seq_end = 5
local seq_by = 0.1 // Setting seq_by = 0.01 will yield the result 3.46
local seq_len = (`seq_end'-`seq_start')/`seq_by' + 1
matrix results = J(`seq_len', 4, 0)
qui gen psi = .
qui gen Hpsi = .
local j = 0
forvalues i = `seq_start'(`seq_by')`seq_end' {
```

```
local j = 'j' + 1
   qui replace psi = `i'
   qui replace Hpsi = wt82_71 - psi * qsmk
   quietly logit qsmk sex race c.age##c.age ///
     ib(last).education c.smokeintensity##c.smokeintensity ///
     c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active ///
     c.wt71##c.wt71 Hpsi ///
     [pw = w_cens], cluster(seqn)
   matrix p_mat = r(table)
   matrix p_mat = p_mat["pvalue","qsmk:Hpsi"]
   local p = p_mat[1,1]
   local b = _b[Hpsi]
   di "coeff", %6.3f `b', "is generated from psi", %4.1f `i'
   matrix results['j',1]= 'i'
   matrix results['j',2]= 'b'
   matrix results['j',3] = abs('b')
   matrix results['j',4]= 'p'
}
matrix colnames results = "psi" "B(Hpsi)" "AbsB(Hpsi)" "pvalue"
mat li results
mata
res = st_matrix("results")
for(i=1; i<= rows(res); i++) {</pre>
 if (res[i,3] == colmin(res[,3])) res[i,1]
}
end
* Setting seq_by = 0.01 will yield the result 3.46
Iteration 0:
             log pseudolikelihood = -936.10067
Iteration 1:
             log pseudolikelihood = -879.13942
Iteration 2:
             log pseudolikelihood = -877.82647
Iteration 3:
             log pseudolikelihood = -877.82423
Iteration 4:
            log pseudolikelihood = -877.82423
Logistic regression
                                            Number of obs =
                                                                  1,566
                                            Wald chi2(19)
                                                                  106.13
                                            Prob > chi2
                                                                  0.0000
Log pseudolikelihood = -877.82423
                                            Pseudo R2
                                                                  0.0623
                            (Std. Err. adjusted for 1,566 clusters in seqn)
_____
           Robust
       qsmk |
                  Coef.
                         Std. Err.
                                     z P>|z| [95% Conf. Interval]
        sex | -.5137324 .1536024 -3.34 0.001 -.8147876 -.2126772
       race | -.8608912 .2099415 -4.10 0.000 -1.272369 -.4494133
        age | .1151589 .0502116
                                    2.29 0.022
                                                    .016746 .2135718
```

c.age#c.age	0007593	.0005297	-1.43	0.152	0017976	.000279
education						
1	4710855	.2247701	-2.10	0.036	9116268	0305441
2		.2208583	-2.26	0.024	9328974	0671487
3 I		.195914	-1.96	0.050	7673632	.0006056
4 I	4047116	.2836068	-1.43	0.154	9605707	.1511476
smokeinten~y	0783425	.014645	-5.35	0.000	1070461	0496389
I						
c.						
smokeinten~y#						
c.						
smokeinten~y	.0010722	.0002651	4.04	0.000	.0005526	.0015917
I						
smokeyrs	0711097	.026398	-2.69	0.007	1228488	0193705
I						
c.smokeyrs#						
c.smokeyrs	.0008153	.0004491	1.82	0.069	000065	.0016955
exercise						
0	3800465	.1889205	-2.01	0.044	7503238	0097692
1	0437043	. 1372725	-0.32	0.750	3127534	. 2253447
I						
active						
0	2134552	.2122025	-1.01	0.314	6293645	.2024541
1	1793327	.207151	-0.87	0.387	5853412	.2266758
I						
wt71	0076607	.0256319	-0.30	0.765	0578983	.0425769
l						
c.wt71#						
c.wt71	.0000866	.0001582	0.55	0.584	0002236	.0003967
I						
Hpsi	-1.90e-06	.0088414	-0.00	1.000	0173307	.0173269
_cons	-1.338367	1.359613	-0.98	0.325	-4.00316	1.326426

<sup>-1.905</sup>e-06

coeff 0.027 is generated from psi 2.0

```
coeff 0.025 is generated from psi
coeff 0.023 is generated from psi
coeff 0.021 is generated from psi
coeff 0.019 is generated from psi
coeff 0.018 is generated from psi
coeff 0.016 is generated from psi
coeff 0.014 is generated from psi
coeff 0.012 is generated from psi
coeff 0.010 is generated from psi
                                   2.9
coeff 0.008 is generated from psi
coeff 0.006 is generated from psi
                                   3.1
coeff 0.005 is generated from psi
coeff 0.003 is generated from psi
coeff 0.001 is generated from psi
coeff -0.001 is generated from psi
coeff -0.003 is generated from psi
coeff -0.005 is generated from psi
                                   3.7
coeff -0.007 is generated from psi
coeff -0.009 is generated from psi
coeff -0.011 is generated from psi
coeff -0.012 is generated from psi
coeff -0.014 is generated from psi
coeff -0.016 is generated from psi
coeff -0.018 is generated from psi
coeff -0.020 is generated from psi
coeff -0.022 is generated from psi
coeff -0.024 is generated from psi 4.7
coeff -0.026 is generated from psi
coeff -0.028 is generated from psi
coeff -0.030 is generated from psi 5.0
```

#### results[31,4]

	psi	B(Hpsi)	AbsB(Hpsi)	pvalue
r1	2	.02672188	.02672188	.00177849
r2	2.1	.02489456	.02489456	.00359089
r3	2.2	.02306552	.02306552	.00698119
r4	2.3	.02123444	.02123444	.01305479
r5	2.4	.01940095	.01940095	.02346121
r6	2.5	.01756472	.01756472	.04049437
r7	2.6	.0157254	.0157254	.06710192
r8	2.7	.01388267	.01388267	.10673812
r9	2.8	.0120362	.0120362	.16301154
r10	2.9	.01018567	.01018567	.23912864
r11	3	.00833081	.00833081	.33720241
r12	3.1	.00647131	.00647131	.45757692
r13	3.2	.0046069	.0046069	.59835195
r14	3.3	.00273736	.00273736	.75528009

```
r15
           3.4
                 .00086243
                             .00086243
                                        .92212566
r16
           3.5 -.00101809
                             .00101809
                                        .90856559
r17
           3.6 -.00290439
                             .00290439
                                         .7444406
r18
           3.7 -.00479666
                             .00479666
                                        .59230593
r19
           3.8 -.00669505
                             .00669505
                                        .45731304
r20
           3.9 -.00859969
                             .00859969
                                         .3425138
r21
             4 -.01051072
                             .01051072
                                         .2488326
           4.1 -.01242824
r22
                            .01242824
                                        .17537691
           4.2 -.01435235
r23
                             .01435235
                                         .1199593
           4.3 -.01628313
                            .01628313
r24
                                        .07967563
           4.4 -.01822063
                             .01822063
                                        .05142147
r25
r26
           4.5 -.02016492
                            .02016492
                                        .03227271
           4.6 -.02211603
r27
                             .02211603
                                        .01971433
r28
           4.7 -.02407401
                            .02407401
                                        .01173271
           4.8 -.02603888
                            .02603888
                                        .00680955
r29
r30
           4.9 -.02801063
                             .02801063
                                        .00385828
             5 -.02998926
r31
                             .02998926
                                        .00213639
----- mata (type end to exit) -----
: res = st_matrix("results")
: for(i=1; i<= rows(res); i++) {
   if (res[i,3] == colmin(res[,3])) res[i,1]
> }
: end
```

# Program 14.3

- G-estimation for 2-parameter structural nested mean model
- Closed form estimator
- Data from NHEFS
- Section 14.6

```
/*create weights*/
logit qsmk sex race c.age##c.age ib(last).education ///
    c.smokeintensity##c.smokeintensity c.smokeyrs##c.smokeyrs ///
    ib(last).exercise ib(last).active c.wt71##c.wt71 ///
    [pw = w_cens], cluster(seqn)
predict pr_qsmk
summarize pr_qsmk

/* Closed form estimator linear mean models **/
* ssc install tomata
putmata *, replace
```

```
mata: diff = qsmk - pr_qsmk
mata: part1 = w_cens :* wt82_71 :* diff
mata: part2 = w_cens :* qsmk :* diff
mata: psi = sum(part1)/sum(part2)
/*** Closed form estimator for 2-parameter model **/
diff = qsmk - pr_qsmk
diff2 = w_cens :* diff
lhs = J(2,2,0)
lhs[1,1] = sum(qsmk :* diff2)
lhs[1,2] = sum( qsmk :* smokeintensity :* diff2 )
lhs[2,1] = sum( qsmk :* smokeintensity :* diff2)
lhs[2,2] = sum( qsmk :* smokeintensity :* smokeintensity :* diff2 )
rhs = J(2,1,0)
rhs[1] = sum(wt82_71 :* diff2)
rhs[2] = sum(wt82_71 :* smokeintensity :* diff2 )
psi = (lusolve(lhs, rhs))'
psi = (invsym(lhs'lhs)*lhs'rhs)'
psi
end
Iteration 0: log pseudolikelihood = -936.10067
Iteration 1: log pseudolikelihood = -879.13943
Iteration 2: log pseudolikelihood = -877.82647
Iteration 3: log pseudolikelihood = -877.82423
Iteration 4:
            log pseudolikelihood = -877.82423
                                         Number of obs
Logistic regression
                                                             1,566
                                         Wald chi2(18) =
                                                            106.13
                                         Prob > chi2
                                                             0.0000
Log pseudolikelihood = -877.82423
                                         Pseudo R2
                                                             0.0623
                          (Std. Err. adjusted for 1,566 clusters in seqn)
______
         - 1
                        Robust
               Coef. Std. Err. z P>|z| [95% Conf. Interval]
      qsmk |
______
       sex | -.5137295 .1533507 -3.35 0.001 -.8142913 -.2131677
      race | -.8608919 .2099555 -4.10 0.000 -1.272397 -.4493867
                                 2.29 0.022
       age | .1151581 .0503079
                                                .0165564
                                                           .2137598
 c.age#c.age | -.0007593 .00053 -1.43 0.152 -.0017981 .0002795
  education |
```

1   2   3   4	4710854 5000247 3833802 4047148	.2247796 .220776 .1954991 .2833093	-2.10 -2.26 -1.96 -1.43	0.036 0.024 0.050 0.153	9116454 9327378 7665515 9599908	0305255 0673116 0002089 .1505613
smokeinten~y	0783426	.0146634	-5.34	0.000	1070824	0496029
c.						
smokeinten~y#						
c.						
smokeinten~y	.0010722	.0002655	4.04	0.000	.0005518	.0015925
1						
smokeyrs	0711099	.0263523	-2.70	0.007	1227596	0194602
1						
c.smokeyrs#						
c.smokeyrs	.0008153	.0004486	1.82	0.069	0000639	.0016945
1						
exercise						
0	3800461	.1890123	-2.01	0.044	7505034	0095887
1	0437044	.137269	-0.32	0.750	3127467	. 225338
1						
active						
0	2134564	.2121759	-1.01	0.314	6293135	.2024007
1	1793322	.2070848	-0.87	0.386	5852109	.2265466
I						
wt71	0076609	.0255841	-0.30	0.765	0578048	.042483
1						
c.wt71#						
c.wt71	.0000866	.0001572	0.55	0.582	0002216	.0003947
1						
_cons	-1.338358	1.359289	-0.98	0.325	-4.002516	1.3258

(option pr assumed; Pr(qsmk))

Variable	Obs	Mean	Std. Dev.	Min	Max
pr_qsmk	+   1,566	. 2607709	.1177584	.0514466	.7891403

(68 vectors posted)



: diff = qsmk - pr\_qsmk

: diff2 = w\_cens :\* diff

```
: lhs = J(2,2, 0)
: lhs[1,1] = sum( qsmk :* diff2)
: lhs[1,2] = sum( qsmk :* smokeintensity :* diff2 )
: lhs[2,1] = sum( qsmk :* smokeintensity :* diff2)
: lhs[2,2] = sum( qsmk :* smokeintensity :* smokeintensity :* diff2 )
: rhs = J(2,1,0)
: rhs[1] = sum(wt82_71 :* diff2)
: rhs[2] = sum(wt82_71 :* smokeintensity :* diff2 )
: psi = (lusolve(lhs, rhs))'
: psi
 1 | 2.859470362 .0300412816 |
   +----+
: psi = (invsym(lhs'lhs)*lhs'rhs)'
: psi
 1 | 2.859470362 .0300412816 |
: end
```

# 15. Outcome regression and propensity scores: Stata

# Program 15.1

- Estimating the average causal effect within levels of confounders under the assumption of effect-measure modification by smoking intensity ONLY
- Data from NHEFS
- Section 15.1

```
/* Generate smoking intensity among smokers product term */
gen qsmkintensity = qsmk*smokeintensity

* Regression on covariates, allowing for some effect modification
regress wt82_71 qsmk qsmkintensity ///
    c.smokeintensity##c.smokeintensity sex race c.age##c.age ///
    ib(last).education c.smokeyrs##c.smokeyrs ///
    ib(last).exercise ib(last).active c.wt71##c.wt71

/* Display the estimated mean difference between quitting and
    not quitting value when smoke intensity = 5 cigarettes/ day */
lincom 1*_b[qsmk] + 5*1*_b[qsmkintensity]

/* Display the estimated mean difference between quitting and
    not quitting value when smoke intensity = 40 cigarettes/ day */
```

```
lincom 1*_b[qsmk] + 40*1*_b[qsmkintensity]

/* Regression on covariates, with no product terms */
regress wt82_71 qsmk c.smokeintensity##c.smokeintensity ///
sex race c.age##c.age ///
ib(last).education c.smokeyrs##c.smokeyrs ///
ib(last).exercise ib(last).active c.wt71##c.wt71
```

Source	SS	df	MS		mber of obs =	,
Model	14412.558	20	720.627		,	= 13.45 = 0.0000
Residual			53.568303			0.1483
					-	0.1400
Total	97175.5866	1.565	62.093026		ot MSE =	
		_,				
		G+ 1 F		DS 1+1		T+
wt82_71	Coef.	Std. Err.	t 	P> t	[95% Conf.	Interval]
qsmk	2.559594	.8091486	3.16	0.002	.9724486	4.14674
qsmkintens~y	.0466628	.0351448	1.33	0.184	0222737	.1155993
smokeinten~y	.0491365	.0517254	0.95	0.342	052323	.1505959
I						
c.						
smokeinten~y#						
c.						
smokeinten~y	0009907	.000938	-1.06	0.291	0028306	.0008493
sex	-1.430272	. 4689576	-3.05	0.002	-2.350132	5104111
race		.5818888	0.96	0.336		1.701485
age		.1633188	2.20	0.028	.0392854	.6799851
c.age#c.age	006101	.0017261	-3.53	0.000	0094868	0027151
education						
1	.194977	.7413692	0.26	0.793	-1.259219	1.649173
2	.9854211	.7012116	1.41	0.160	390006	2.360848
3	.7512894	.6339153	1.19	0.236	4921358	1.994715
4	1.686547	.8716593	1.93	0.053	0232138	3.396307
smokeyrs	. 1343686	.0917122	1.47	0.143	045525	.3142621
c.smokeyrs#  c.smokeyrs	0018664	.0015437	-1.21	0.227	0048944	.0011616
C.SMOKEYIS	0010004	.0015457	-1.21	0.221	0040944	.0011616
exercise						
0	3539128	. 5588587	-0.63	0.527	-1.450114	.7422889
1		.4316468	-0.13	0.893		.7887381
- ' 					. 30 2020	
active						

0	.2613779	.6845577	0.38	0.703	-1.081382	1.604138
1	6861916	.6739131	-1.02	0.309	-2.008073	.6356894
1						
wt71	.0455018	.0833709	0.55	0.585	1180303	.2090339
1						
c.wt71#						
c.wt71	0009653	.0005247	-1.84	0.066	0019945	.0000639
1						
_cons	-1.690608	4.388883	-0.39	0.700	-10.2994	6.918188

# (1) qsmk + 5\*qsmkintensity = 0

wt82_71			[95% Conf.	Interval]
			1.482117	4.1037

# (1) qsmk + 40\*qsmkintensity = 0

wt82_71	Coef.	Std. Err.					
	4.426108	.8477818	5.22	0.000	2.763183		6.089032
	SS						
+				- F(19,	1546)	=	14.06
Model	14318.1239	19	753.5854	7 Prob	> F	=	0.0000
Residual	82857.4627	1,546	53.594736	5 R-squ	ared	=	0.1473
+				- Adj R	-squared	=	0.1369
Total	97175.5866	1,565	62.093026	6 Root	MSE	=	7.3208
wt82_71	Coef.	Std. Err.	t	P> t	[95% Con	f.	Interval]
wt82_71	Coef.	Std. Err.	t 	P> t	[95% Con	f. 	Interval]
wt82_71   + qsmk	Coef. 3.462622	Std. Err. 4384543	t 7.90	P> t  	[95% Con  2.602594	f.	Interval]  4.32265
wt82_71	Coef. 3.462622	Std. Err. 4384543	t 7.90	P> t  	[95% Con  2.602594	f.	Interval]  4.32265
wt82_71   + qsmk	Coef. 3.462622	Std. Err. 4384543	t 7.90	P> t  	[95% Con  2.602594	f.	Interval]  4.32265
wt82_71   	Coef. 3.462622	Std. Err. 4384543	t 7.90	P> t  	[95% Con  2.602594	f.	Interval]  4.32265
wt82_71   	Coef. 3.462622	Std. Err. 4384543	t 7.90	P> t  	[95% Con  2.602594	f.	Interval]  4.32265
<pre>wt82_71   qsmk   smokeinten~y  </pre>	Coef. 3.462622 .0651533	Std. Err.  .4384543 .0503115	t 7.90 1.29	P> t   0.000 0.196	[95% Con  2.602594 0335327	f.	Interval] 4.32265 .1638392
<pre>wt82_71   qsmk   smokeinten~y  </pre>	Coef. 3.462622 .0651533	Std. Err.  .4384543 .0503115	t 7.90 1.29	P> t   0.000 0.196	[95% Con  2.602594 0335327	f.	Interval] 4.32265 .1638392
<pre>wt82_71  </pre>	Coef. 3.462622 .0651533	Std. Err. .4384543 .0503115	t 7.90 1.29	P> t   0.000 0.196	[95% Con  2.602594 0335327	f.	Interval] 4.32265 .1638392
<pre>wt82_71  </pre>	Coef. 3.462622 .0651533	Std. Err. .4384543 .0503115	7.90 1.29 -1.12	P> t   0.000 0.196 0.264 0.002	[95% Con  2.602594 0335327 0028853 -2.3837	f.	Interval] 4.32265 .1638392 .00079185463989

age	.3626624	.1633431	2.22	0.027	.0422649	.6830599
c.age#c.age	0061377	.0017263	-3.56	0.000	0095239	0027515
education						
1	.1708264	.7413289	0.23	0.818	-1.28329	1.624943
2	.9893527	.7013784	1.41	0.159	3864007	2.365106
3	.7423268	.6340357	1.17	0.242	501334	1.985988
4	1.679344	.8718575	1.93	0.054	0308044	3.389492
1						
smokeyrs	.1333931	.0917319	1.45	0.146	0465389	.3133252
c.smokeyrs#						
c.smokeyrs	001827	.0015438	-1.18	0.237	0048552	.0012012
exercise						
0	3628786	.5589557	-0.65	0.516	-1.45927	.7335129
1	0421962	.4315904	-0.10	0.922	8887606	.8043683
1						
active						
0	.2580374	.6847219	0.38	0.706	-1.085044	1.601119
1	68492	.6740787	-1.02	0.310	-2.007125	.6372851
wt71	.0373642	.0831658	0.45	0.653	1257655	.200494
1						
c.wt71#						
c.wt71	0009158	.0005235	-1.75	0.080	0019427	.0001111
1						
_cons	-1.724603	4.389891	-0.39	0.694	-10.33537	6.886166

# Prorgam 15.2

- Estimating and plotting the propensity score
- Data from NHEFS
- Section 15.2

```
/*Fit a model for the exposure, quitting smoking*/
logit qsmk sex race c.age##c.age ib(last).education ///
    c.smokeintensity##c.smokeintensity ///
    c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active ///
    c.wt71##c.wt71

/*Estimate the propensity score, P(Qsmk|Covariates)*/
predict ps, pr
```

```
/*Check the distribution of the propensity score*/
bys qsmk: summarize ps
/*Return extreme values of propensity score:
 note, for Stata versions 15 and above, start by installing extremes*/
* ssc install extremes
extremes ps seqn
bys qsmk: extremes ps seqn
save ./data/nhefs-ps, replace
/*Plotting the estimated propensity score*/
histogram ps, width(0.05) start(0.025) ///
 frequency fcolor(none) lcolor(black) ///
 lpattern(solid) addlabel ///
 addlabopts(mlabcolor(black) mlabposition(12) ///
 mlabangle(zero)) ///
 ytitle(No. Subjects) ylabel(#4) ///
 xtitle(Estimated Propensity Score) xlabel(#15) ///
 by(, title(Estimated Propensity Score Distribution) ///
 subtitle(By Quit Smoking Status)) ///
 by(, legend(off)) ///
 by(qsmk, style(compact) colfirst) ///
 subtitle(, size(small) box bexpand)
qui gr export ./figs/stata-fig-15-2.png, replace
Iteration 0:
             log\ likelihood = -893.02712
Iteration 1: log likelihood = -839.70016
Iteration 2: log likelihood = -838.45045
Iteration 3: log likelihood = -838.44842
Iteration 4: log likelihood = -838.44842
Logistic regression
                                          Number of obs
                                                               1,566
                                          LR chi2(18)
                                                               109.16
                                          Prob > chi2
                                                               0.0000
Log likelihood = -838.44842
                                          Pseudo R2
                                                               0.0611
           ______
                 Coef.
                        Std. Err.
                                                  [95% Conf. Interval]
                                  z P>|z|
      asmk |
______
       sex | -.5274782 .1540497
                                  -3.42 0.001
                                                   -.82941
                                                            -.2255463
       race | -.8392636 .2100668 -4.00 0.000 -1.250987 -.4275404
              .1212052 .0512663 2.36 0.018 .0207251
                                                             .2216853
       age |
 c.age#c.age | -.0008246
                       .0005361
                                  -1.54 0.124 -.0018753 .0002262
  education |
        1 | -.4759606 .2262238
                                  -2.10 0.035 -.9193511 -.0325701
        2 | -.5047361 .217597
                                  -2.32 0.020 -.9312184 -.0782538
```

3	3895288	.1914353	-2.03	0.042	7647351	0143226
4	4123596	.2772868	-1.49	0.137	9558318	.1311126
smokeinten~y	0772704	.0152499	-5.07	0.000	1071596	0473812
c.						
smokeinten~y#						
c.						
smokeinten~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
smokeyrs	0735966	.0277775	-2.65	0.008	1280395	0191538
<pre>c.smokeyrs#  c.smokeyrs  </pre>	.0008441	.0004632	1.82	0.068	0000637	.0017519
c.smokeyis	.0006441	.0004632	1.02	0.000	0000637	.0017519
exercise						
0	395704	.1872401	-2.11	0.035	7626878	0287201
1	0408635	.1382674	-0.30	0.768	3118627	.2301357
active	470704	04.40704	0.00	0 111	5004045	0445505
0	176784	.2149721	-0.82	0.411	5981215	. 2445535
1	1448395	.2111472	-0.69	0.493	5586806	.2690015
wt71	0152357	.0263161	-0.58	0.563	0668144	.036343
+71#						
c.wt71#	0001250	0001630	0.00	0 407	0001046	000455
c.wt71   	.0001352	.0001632	0.83	0.407	0001846	.000455
_cons	-1.19407	1.398493	-0.85	0.393	-3.935066	1.546925

\_\_\_\_\_

<sup>-&</sup>gt; qsmk = No smoking cessation

Variable	0bs	Mean	Std. Dev.	Min	Max
ps	1,163	.2392928	.1056545	.0510008	.6814955

-----

## -> qsmk = Smoking cessation

Variable	Obs	Mean	Std. Dev.	Min	Max
ps	403	.3094353	.1290642	.0598799	.7768887

+-			+
1	obs:	ps	seqn
-			
1	65.	.0510008	22941
1	364.	.0527126	1769
1	427.	.0558418	21140
1	578.	.0558752	2522
1	1139.	.0567372	12639
+-			+
+-			+
1	1365.	.6659576	22272
1	13.	.6814955	22773
1	1346.	.7166381	14983
1	1289.	.7200644	24817
1	1495.	.7768887	24949
+-			+

\_\_\_\_\_\_

### -> qsmk = No smoking cessation

+		+
obs:	ps	seqn
65.	.0510008	22941
1 364.	.0527126	1769
427.	.0558418	21140
578.	.0558752	2522
1139.	.0567372	12639
+		+

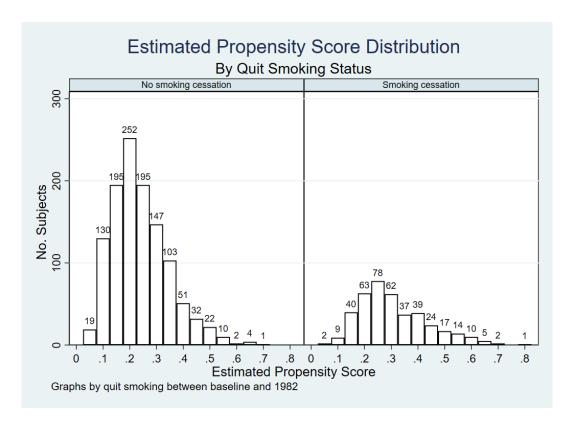
+----+

## -> qsmk = Smoking cessation

165

```
| 1263.
          .0821677
                      22904 |
 1511.
          .1021875
                      24584 |
 1215.
           .635695
                      17738
| 1365.
          .6659576
                      22272 |
| 1346.
          .7166381
                      14983 |
| 1289.
          .7200644
                      24817
| 1495.
          .7768887
                      24949 |
```

file ./data/nhefs-ps.dta saved



# Program 15.3

- Stratification and outcome regression using deciles of the propensity score
- Data from NHEFS
- Section 15.3
- Note: Stata decides borderline cutpoints differently from SAS, so, despite identically distributed propensity scores, the results of regression using decides are not an exact match with the book.

```
use ./data/nhefs-ps, clear
```

```
/*Calculation of deciles of ps*/
xtile ps_dec = ps, nq(10)
by ps_dec, sort: summarize ps
/*Stratification on PS deciles, allowing for effect modification*/
/*Note: stata compares qsmk 0 vs qsmk 1, so the coefficients are reversed relative to the book*/
by ps_dec: ttest wt82_71, by(qsmk)
/*Regression on PS deciles, with no product terms*/
regress wt82_71 qsmk ib(last).ps_dec
-> ps_dec = 1
  Variable | Obs Mean Std. Dev. Min Max
______
      ps | 157 .0976251 .0185215 .0510008 .1240482
\rightarrow ps_dec = 2
  Variable | Obs Mean Std. Dev. Min Max
______
      ps | 157 .1430792 .0107751 .1241923 .1603558
\rightarrow ps_dec = 3
  Variable | Obs Mean Std. Dev. Min Max
      ps | 156 .1750423 .008773 .1606041 .1893271
______
\rightarrow ps_dec = 4
  Variable | Obs Mean Std. Dev. Min Max
-----
             157 .2014066 .0062403 .189365 .2121815
\rightarrow ps_dec = 5
  Variable | Obs Mean Std. Dev. Min Max
______
      ps | 156 .2245376 .0073655 .2123068 .237184
\rightarrow ps_dec = 6
  Variable | Obs Mean Std. Dev. Min Max
```

ps	157	.2515298	.0078777	. 2377578	.2655718	
-> ps_dec = 7						
Variable	Obs	Mean	Std. Dev.	Min	Max	
ps	157	. 2827476	.0099986	. 2655724	.2994968	
Variable			Std. Dev.	Min	Max	
ps		.3204104	.0125102	. 2997581	.3438773	
-> ps_dec = 9						
Variable	Obs	Mean	Std. Dev.	Min	Max	
ps	157	.375637	.0221347	.3439862	.4174631	
-> ps_dec = 10						
Variable	Obs	Mean	Std. Dev.	Min	Max	
ps	156	.5026508	.0733494	.4176717	.7768887	
Two-sample t t	<del>-</del>		} 			
	Obs	Mean Std.	Err. Std.	Dev. [9	5% Conf. In	terval]
No smoki   Smoking	146 3.7 11 3.94	74236 .653 19703 2.33	31341 7.89 32995 7.73	1849 2.4	451467 5	.033253
combined	157 3.75	66887 .627				. 995484
diff	207	3431 2.46	64411 	-5.0	075509 4	
		- mean(Smoki	ng)			-0.0841

Ha: diff < 0 Ha: diff != 0 Ha: diff > 0 Pr(T < t) = 0.4665 Pr(|T| > |t|) = 0.9331 Pr(T > t) = 0.5335

 $\rightarrow$  ps\_dec = 2

Two-sample t test with equal variances

Group		Mean		Std. Dev.	[95% Conf.	Interval]
No smoki Smoking	l 134 l 23	2.813019 7.726944	.589056 1.260784	6.818816 6.046508	1.647889 5.112237	3.978149 10.34165
combined	l 157	3.532893	.5519826	6.916322	2.442569	4.623217
diff		-4.913925			-7.907613	
diff =		smoki) - mea	n(Smoking)	degree	t s of freedom	= -3.2425 = 155

-----

 $\rightarrow$  ps\_dec = 3

Two-sample t test with equal variances

Group		Mean	Std. Err.		[95% Conf	. Interval]
No smoki Smoking	128   28	3.25684 7.954974	.5334655 1.418184	6.035473 7.504324	2.201209 5.045101	4.312472 10.86485
combined	156	4.100095	.5245749	6.551938	3.063857	5.136334
diff		-4.698134	1.318074		-7.301973	-2.094294

-----

 $\rightarrow$  ps\_dec = 4

Two-sample t test with equal variances

\_\_\_\_\_\_

Group		Mean	Std. Err.	Std. Dev.	[95% Conf.	Interval]
No smoki	121	3.393929		5.794362		
Smoking   +			1.543143 9.258861			8.808819
	157	3.917223				4.986266
diff					-4.807663	. 2433778
diff = m Ho: diff = 0		moki) - mean	(Smoking)	degrees	t of freedom	= -1.7850 = 155
Ha: diff			Ha: diff !=			iff > 0
Pr(T < t) =	0.0381	Pr(	T  >  t ) =	0.0762	Pr(T > t	) = 0.9619
 Group   		Mean	Std. Err.		[95% Conf.	Interval]
-					[95% Conf.	Interval]
		1.368438				
Smoking		5.195421	1.388723	8.44727	2.378961	8.011881
		2.27612				3.671489
diff		-3.826983			-7.061407	592559
diff = m Ho: diff = 0		moki) - mean	(Smoking)	degrees	t of freedom	= -2.3374 = 154
Ha: diff	< 0		Ha: diff !=	0	Ha: d	iff > 0
Pr(T < t) = 0.0104 $Pr( T  >  t ) = 0.0207$					Pr(T > t	) = 0.9896
-> ps_dec =	6					
Two-sample t	test wi	th equal var	iances			

Group	Obs	Mean	Std. Err.		[95% Conf.	Interval]
No smoki Smoking	112   45	2.25564 7.199088	.6850004 1.724899	7.249362 11.57097	.8982664 3.722782	3.613014 10.67539
combined	l 157	3.672552	.7146582	8.954642	2.260897	5.084207
diff	 	-4.943447	1.535024		-7.975714	

diff = mean(No smoki) - mean(Smoking) t = -3.2204Ho: diff = 0degrees of freedom = 155 Ha: diff < 0 Ha: diff != 0 Ha: diff > 0 Pr(T < t) = 0.0008 Pr(|T| > |t|) = 0.0016 Pr(T > t) = 0.9992 $\rightarrow$  ps\_dec = 7 Two-sample t test with equal variances Group | Obs Mean Std. Err. Std. Dev. [95% Conf. Interval] No smoki | 116 .7948483 .7916172 8.525978 -.773193 41 6.646091 1.00182 6.414778 4.621337 8.670844 Smoking | 2.32288 .6714693 8.413486 combined | 157 .9965349 \_\_\_\_\_\_ -5.851242 1.45977 diff | -8.734853 -2.967632 \_\_\_\_\_ diff = mean(No smoki) - mean(Smoking) t = -4.0083Ho: diff = 0degrees of freedom = 155 Ha: diff < 0 Ha: diff != 0 Ha: diff > 0 Pr(T < t) = 0.0000 Pr(|T| > |t|) = 0.0001 Pr(T > t) = 1.0000\_\_\_\_\_\_  $\rightarrow$  ps\_dec = 8 Two-sample t test with equal variances \_\_\_\_\_\_ Group | Obs Mean Std. Err. Std. Dev. [95% Conf. Interval] No smoki | 107 1.063848 .5840159 6.041107 -.0940204 2.221716 Smoking | 49 3.116263 1.113479 7.794356 .8774626 5.355063 \_\_\_\_\_\_\_\_\_\_\_ 156 1.708517 .5352016 6.684666 .6512864 \_\_\_\_\_\_ diff | -2.052415 1.144914 -4.31418 \_\_\_\_\_ diff = mean(No smoki) - mean(Smoking) Ho: diff = 0degrees of freedom = Ha: diff < 0 Ha: diff != 0 Ha: diff > 0 Pr(T < t) = 0.0375 Pr(|T| > |t|) = 0.0750 Pr(T > t) = 0.9625

 $\rightarrow$  ps\_dec = 9

Two-sample t test with equal variances

					Std.	Dev.	 [95% Conf	. Interval]
No smoki					7.63	 7396	 -1.544716	1.486134
	57	.911264	7 .99	969309				2.908357
·					7.58	6766	8838316	1.508201
diff		940555	54 1	.26092			-3.43136	1.550249
	mean(No s	smoki) - m	nean(Smo	king)	3			= -0.7459
Ho: diff =	U				a	egrees	of freedom	= 155
Ha: di	ff < 0		Ha:	diff !=	0		Ha:	diff > 0 t) = 0.7716
Pr(T < t)	= 0.2284	F	r( T  >	t ) =	0.4568		Pr(T >	t) = 0.7716
	= 10							
Two-sample	t test wi	ith equal	varianc	es				
Group	0bs	Mea	n Sto	d. Err. 	Std.	Dev.	[95% Conf	. Interval]
					8.25	0872	-2.604646	1.067638
	76 				9.18	0992	.2973737	4.493267
combined		.772846	3 .70	071067		1759	6239631	2.169656
							-5.921957	405692
diff =	mean(No s	smoki) - m	nean(Smo	 king)			 t	= -2.2661
Ho: diff =	0				d	egrees	of freedom	= 154
Ha: di	ff < 0		Ha:	diff !=	0		Ha:	diff > 0
Pr(T < t)	= 0.0124	P	r( T  >	t ) =	0.0248		Pr(T >	t) = 0.9876
	ce						r of obs	= 1,566
	+					,		= 9.87
								= 0.0000 = 0.0597
								= 0.0536
Tota	al   9717	75.5866	1,565	62.093	0266	Root 1	MSE	= 7.6657
wt82_	71	Coef. S	Std. Err	. t	P>	t	[95% Conf	. Interval]
	•							4.255368

```
ps_dec |
   1 |
          4.384269
                   .8873947
                                4.94
                                       0.000
                                                 2.643652
                                                            6.124885
   2 |
          3.903694
                    .8805212
                                4.43
                                       0.000
                                                 2.17656
                                                            5.630828
                                4.96
   3 l
           4.36015
                   .8793345
                                       0.000
                                                 2.635343
                                                            6.084956
   4 |
          4.010061
                                4.59
                    .8745966
                                       0.000
                                                 2.294548
                                                            5.725575
   5 |
                                2.68
          2.342505
                   .8754878
                                       0.008
                                                 .6252438
                                                            4.059766
   6 |
          3.572955
                    .8714389
                                4.10
                                       0.000
                                                 1.863636
                                                            5.282275
   7 |
           2.30881
                                2.65 0.008
                   .8727462
                                                 .5969261
                                                            4.020693
   8 |
        1.516677
                                1.74
                    .8715796
                                       0.082
                                                -.1929182
                                                            3.226273
   9 | -.0439923
                   .8684465
                               -0.05
                                       0.960
                                               -1.747442
                                                            1.659457
 _cons | -.8625798
                   .6530529
                               -1.32 0.187
                                               -2.143537
                                                            .4183773
```

#### Program 15.4

- Standardization and outcome regression using the propensity score
- Data from NHEFS
- Section 15.3

```
use ./data/nhefs-formatted, clear
/*Estimate the propensity score*/
logit qsmk sex race c.age##c.age ib(last).education ///
  c.smokeintensity##c.smokeintensity ///
  c.smokeyrs##c.smokeyrs ib(last).exercise ///
  ib(last).active c.wt71##c.wt71
predict ps, pr
/*Expand the dataset for standardization*/
expand 2, generate(interv)
expand 2 if interv == 0, generate(interv2)
replace interv = -1 if interv2 ==1
drop interv2
tab interv
replace wt82_71 = . if interv != -1
replace qsmk = 0 if interv == 0
replace qsmk = 1 if interv == 1
by interv, sort: summarize qsmk
/*Regression on the propensity score, allowing for effect modification*/
regress wt82_71 qsmk##c.ps
predict predY, xb
by interv, sort: summarize predY
quietly summarize predY if(interv == -1)
matrix input observe = (-1, r(mean)')
quietly summarize predY if(interv == 0)
```

```
matrix observe = (observe \0, r(mean)')
quietly summarize predY if(interv == 1)
matrix observe = (observe \1, r(mean)')
matrix observe = (observe \., observe[3,2]-observe[2,2])
matrix rownames observe = observed E(Y(a=0)) E(Y(a=1)) difference
matrix colnames observe = interv value
matrix list observe
/*bootstrap program*/
drop if interv != -1
gen meanY_b =.
qui save ./data/nhefs_std, replace
capture program drop bootstdz
program define bootstdz, rclass
use ./data/nhefs std, clear
preserve
bsample
/*Create 2 new copies of the data.
Set the outcome AND the exposure to missing in the copies*/
expand 2, generate(interv_b)
expand 2 if interv_b == 0, generate(interv2_b)
qui replace interv_b = -1 if interv2_b ==1
qui drop interv2_b
qui replace wt82_71 = . if interv_b != -1
qui replace qsmk = . if interv_b != -1
/*Fit the propensity score in the original data
(where qsmk is not missing) and generate predictions for everyone*/
logit qsmk sex race c.age##c.age ib(last).education ///
  c.smokeintensity##c.smokeintensity ///
    c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active ///
    c.wt71##c.wt71
predict ps_b, pr
/*Set the exposure to 0 for everyone in copy 0,
and 1 to everyone for copy 1*/
qui replace qsmk = 0 if interv_b == 0
qui replace qsmk = 1 if interv_b == 1
/*Fit the outcome regression in the original data
(where wt82_71 is not missing) and
generate predictions for everyone*/
regress wt82_71 qsmk##c.ps
predict predY_b, xb
/*Summarize the predictions in each set of copies*/
```

```
summarize predY_b if interv_b == 0
return scalar boot 0 = r(mean)
summarize predY_b if interv_b == 1
return scalar boot_1 = r(mean)
return scalar boot_diff = return(boot_1) - return(boot_0)
qui drop meanY_b
restore
end
/*Then we use the `simulate` command to run the bootstraps
as many times as we want.
Start with reps(10) to make sure your code runs,
and then change to reps(1000) to generate your final CIs*/
simulate EY_a0=r(boot_0) EY_a1 = r(boot_1) ///
 difference = r(boot_diff), reps(500) seed(1): bootstdz /
matrix pe = observe[2..4, 2]'
matrix list pe
bstat, stat(pe) n(1629)
estat bootstrap, p
Iteration 0: log likelihood = -893.02712
Iteration 1: log likelihood = -839.70016
Iteration 2: log likelihood = -838.45045
Iteration 3: log likelihood = -838.44842
Iteration 4: log likelihood = -838.44842
Logistic regression
                                          Number of obs
                                                               1,566
                                          LR chi2(18)
                                                               109.16
                                          Prob > chi2
                                                               0.0000
Log likelihood = -838.44842
                                          Pseudo R2
                                                               0.0611
      qsmk |
               Coef. Std. Err. z P>|z| [95% Conf. Interval]
  ._________
       sex | -.5274782 .1540497 -3.42 0.001
                                                   -.82941
                                                            -.2255463
      race | -.8392636 .2100668 -4.00 0.000 -1.250987 -.4275404
       age | .1212052 .0512663 2.36 0.018
                                                 .0207251 .2216853
 c.age#c.age | -.0008246
                       .0005361 -1.54 0.124 -.0018753 .0002262
  education |
        1 | -.4759606
                                   -2.10 0.035
                                                  -.9193511
                       .2262238
                                                            -.0325701
        2 | -.5047361 .217597
                                   -2.32 0.020
                                                 -.9312184 -.0782538
        3 | -.3895288 .1914353
                                   -2.03
                                          0.042
                                                  -.7647351 -.0143226
        4 | -.4123596 .2772868 -1.49 0.137
                                                 -.9558318 .1311126
smokeinten~y | -.0772704
                                  -5.07 0.000
                       .0152499
                                                  -.1071596 -.0473812
```

c.						
smokeinten~y#						
c.						
smokeinten~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
1						
smokeyrs	0735966	.0277775	-2.65	0.008	1280395	0191538
1						
c.smokeyrs#						
c.smokeyrs	.0008441	.0004632	1.82	0.068	0000637	.0017519
1						
exercise						
0	395704	.1872401	-2.11	0.035	7626878	0287201
1	0408635	.1382674	-0.30	0.768	3118627	.2301357
1						
active						
0	176784	.2149721	-0.82	0.411	5981215	. 2445535
1	1448395	.2111472	-0.69	0.493	5586806	.2690015
1						
wt71	0152357	.0263161	-0.58	0.563	0668144	.036343
1						
c.wt71#						
c.wt71	.0001352	.0001632	0.83	0.407	0001846	.000455
1						
_cons	-1.19407	1.398493	-0.85	0.393	-3.935066	1.546925

(1,566 observations created)

(1,566 observations created)

(1,566 real changes made)

interv	1	Freq.	Percent	Cum.
-1 0 1	į	1,566 1,566 1,566	33.33 33.33 33.33	33.33 66.67 100.00
Total		4,698	100.00	

(3,132 real changes made, 3,132 to missing)

(403 real changes made)

(1,163 real changes made)

\_\_\_\_\_

->	interv	=	-1
----	--------	---	----

Variable	0bs	Mean	Std.	Dev.	Min	N	ſax
qsmk	1,566	. 2573436	. 4373	3099	0		1
-> interv = 0							
	0bs			Dev.	Min	N	lax
•	1,566			0	0		0
-> interv = 1							
Variable		Mean	Std.	Dev.	Min	N	ſax
·	1,566	1		0	1		1
	SS				Number of ob F(3, 1562)		
•					Prob > F		
	91888.2723						
					Adj R-square		
Total	97175.5866	1,565	62.09302	266	Root MSE		
wt82_71	Coef.	Std. Err.	t	P>	t  [95%	Conf.	Interval]
qsmk							
Smoking c	4.036457	1.13904	3.54	0.	000 1.80	)225	6.270665
					000 -16.50		
qsmk#c.ps							
Smoking c	-2.038829	3.649684	-0.56	0.	576 -9.197	7625	5.119967
_cons	4.935432	.5570216	8.86	0.	000 3.842	2843	6.028021

-> interv = -1

Variable	Obs	Mean	Std. Dev.	Min	Max
	 1,566	2.6383	1.838063	-3.4687	8.111371

-> interv = 0					
Variable	0bs	Mean		Min	Max
 predY	1,566			-4.645079	4.306496
F	_,				
-> interv = 1					
Variable	Obs	Mean	Std. Dev.	Min	Max
+					
predY	1,566	5.273676	1.670225	-2.192565	8.238971
observe[4,2]					
	interv	value			
	-1 2.63				
E(Y(a=0))	0 1.76	18979			
E(Y(a=1))	1 5.27	36757			
difference	. 3.51	17778			
(3,132 observa	tions deleted	.)			
(1,566 missing	values gener	ated)			
command:	•				
EY_a0:	_				
EY_a1:	_				
difference:	r(boot_diff	)			
<b>4</b>	00)				
Simulations (5		•	_		
+ 1				F.0	
				50	
				100	
				150	
	• • • • • • • • • • • • • • • • • • • •	• • • • • • • • • • • • • • • • • • • •		200	
• • • • • • • • • • • • • • • • • • • •		• • • • • • • • • • • • • • • • • • • •	• • • • • • • • • • • • • • • • • • • •	250	

 350
 400
 450
 500

pe[1,3]

E(Y(a=0)) E(Y(a=1)) difference value 1.7618979 5.2736757 3.5117778

Bootstrap result	S			Number	of obs	=	1,629
				Replica	tions	=	500
	Observed	Bootstrap			Nor	 mal	 -based
i	Coef.	Std. Err.	z	P> z	[95% Co	nf.	Interval]
+							
EY_a0	1.761898	.2273005	7.75	0.000	1.31639	7	2.207399
EY_a1	5.273676	.4636993	11.37	0.000	4.36484	2	6.18251
difference	3.511778	.5247816	6.69	0.000	2.48322	5	4.540331

Bootstrap result	cs		Num	ber of obs	=	1,629
			Rep	lications	=	500
			_			
1	Observed		Bootstrap			
1	Coef.	Bias	Std. Err.	[95% Conf.	<pre>Interval]</pre>	
EY_a0	1.7618979	0079171	.22730051	1.307729	2.191437	(P)
EY_a1	5.2736757	0001726	.46369929	4.340613	6.210879	(P)
difference	3.5117778	.0077445	.5247816	2.531495	4.523983	(P)

<sup>(</sup>P) percentile confidence interval

# 16. Instrumental variables estimation: Stata

#### Program 16.1

library(Statamarkdown)

- Estimating the average causal effect using the standard IV estimator via the calculation of sample averages
- Data from NHEFS
- Section 16.2

```
use ./data/nhefs-formatted, clear

summarize price82

/* ignore subjects with missing outcome or missing instrument for simplicity*/
foreach var of varlist wt82 price82 {
    drop if `var'==. }

/*Create categorical instrument*/
gen byte highprice = (price82 > 1.5 & price82 < .)

save ./data/nhefs-highprice, replace

/*Calculate P[Z/A=a]*/
tab highprice qsmk, row</pre>
```

```
/*Calculate P[Y/Z=z]*/
ttest wt82_71, by(highprice)
/*Final IV estimate, OPTION 1: Hand calculations*/
/*Numerator: num = E[Y|Z=1] - E[Y|Z=0] = 2.686 - 2.536 = 0.150*/
/*Denominator: denom = P[A=1|Z=1] - P[A=1|Z=0] = 0.258 - 0.195 = 0.063 */
/*IV estimator: E[Ya=1] - E[Ya=0] = (E[Y|Z=1] - E[Y|Z=0])/(P[A=1|Z=1] - P[A=1|Z=0]) = 0.150/0.063 = 2.397*/
display "Numerator, E[Y|Z=1] - E[Y|Z=0] = ", 2.686 - 2.536
display "Denominator: denom = P[A=1|Z=1] - P[A=1|Z=0] =", 0.258 - 0.195
display "IV estimator =", 0.150/0.063
/*OPTION 2 2: automated calculation of instrument*/
/*Calculate P[A=1/Z=z], for each value of the instrument,
and store in a matrix*/
quietly summarize qsmk if (highprice==0)
matrix input pa = (`r(mean)')
quietly summarize qsmk if (highprice==1)
matrix pa = (pa , r(mean)')
matrix list pa
/*Calculate P[Y|Z=z], for each value of the instrument,
and store in a second matrix*/
quietly summarize wt82_71 if (highprice==0)
matrix input ey = (`r(mean)')
quietly summarize wt82_71 if (highprice==1)
matrix ey = (ey , r(mean)')
matrix list ey
/*Using Stata's built-in matrix manipulation feature (Mata),
calculate numerator, denominator and IV estimator*/
*Numerator: num = E[Y|Z=1] - E[Y|Z=0]*mata
*Denominator: denom = P[A=1|Z=1] - P[A=1|Z=0]*
*IV estimator: iv_est = IV estimate of E[Ya=1] - E[Ya=0] *
mata
pa = st_matrix("pa")
ey = st_matrix("ey")
num = ey[1,2] - ey[1,1]
denom = pa[1,2] - pa[1,1]
iv_est = num / denom
num
denom
st_numscalar("iv_est", iv_est)
end
di scalar(iv_est)
```

```
(0 observations deleted)
(90 observations deleted)

file ./data/nhefs-highprice.dta cannot be modified or erased; likely cause is read-only directory or file
r(608);
end of do-file
r(608);
```

#### Program 16.2

- Estimating the average causal effect using the standard IV estimator via two-stage-least-squares regression
- Data from NHEFS
- Section 16.2

```
use ./data/nhefs-highprice, clear
/* ivregress fits the model in two stages:
- first model: qsmk = highprice
- second model: wt82_71 = predicted_qsmk */
ivregress 2sls wt82_71 (qsmk = highprice)
Instrumental variables (2SLS) regression
                                                  Number of obs
                                                                        1,476
                                                  Wald chi2(1)
                                                                         0.01
                                                  Prob > chi2
                                                                        0.9038
                                                  R-squared
                                                                        0.0213
                                                  Root MSE
                                                                        7.8508
```

wt82_71	Std. Err.			Interval]
-			-36.46313 -7.89169	

Instrumented: qsmk
Instruments: highprice

#### Program 16.3

- Estimating the average causal effect using the standard IV estimator via an additive marginal structural model
- Data from NHEFS
- Checking one possible value of psi.

- See Chapter 14 for program that checks several values and computes 95% confidence intervals
- Section 16.2

```
use ./data/nhefs-highprice, clear
gen psi = 2.396
gen hspi = wt82_71 - psi*qsmk
logit highprice hspi
Iteration 0:
           log\ likelihood = -187.34948
Iteration 1:
           log likelihood = -187.34948
Logistic regression
                                      Number of obs
                                                          1,476
                                      LR chi2(1)
                                                           0.00
                                      Prob > chi2
                                                         1.0000
Log likelihood = -187.34948
                                      Pseudo R2
                                                         0.0000
______
             Coef. Std. Err. z P>|z| [95% Conf. Interval]
```

\_\_\_\_\_

hspi | 2.75e-07 .0201749 0.00 1.000

\_cons | 3.555347 .1637931 21.71 0.000

#### Program 16.4

• Estimating the average causal effect using the standard IV estimator based on alternative proposed instruments

-.0395419

3.234319

.0395424

3.876376

- Data from NHEFS
- Section 16.5

```
use ./data/nhefs-highprice, clear

/*Instrument cut-point: 1.6*/
replace highprice = .
replace highprice = (price82 >1.6 & price82 < .)

ivregress 2sls wt82_71 (qsmk = highprice)

/*Instrument cut-point: 1.7*/
replace highprice = .
replace highprice = (price82 >1.7 & price82 < .)

ivregress 2sls wt82_71 (qsmk = highprice)</pre>
```

```
/*Instrument cut-point: 1.8*/
replace highprice = .
replace highprice = (price82 >1.8 & price82 < .)</pre>
ivregress 2sls wt82_71 (qsmk = highprice)
/*Instrument cut-point: 1.9*/
replace highprice = .
replace highprice = (price82 >1.9 & price82 < .)</pre>
ivregress 2sls wt82_71 (qsmk = highprice)
(1,476 real changes made, 1,476 to missing)
(1,476 real changes made)
Instrumental variables (2SLS) regression
                                 Number of obs =
                                                1,476
                                 Wald chi2(1) =
                                                0.06
                                 Prob > chi2 =
                                                0.8023
                                 R-squared
                                 Root MSE = 18.593
._____
   wt82_71 | Coef. Std. Err. z P>|z| [95% Conf. Interval]
-----
    qsmk | 41.28124 164.8417 0.25 0.802 -281.8026 364.365
    Instrumented: qsmk
Instruments: highprice
(1,476 real changes made, 1,476 to missing)
(1,476 real changes made)
Instrumental variables (2SLS) regression
                                 Number of obs =
                                                1,476
                                 Wald chi2(1) =
                                                0.05
                                 Prob > chi2 = 0.8274
                                 R-squared
                                           =
                                         = 20.577
                                 Root MSE
______
   wt82_71 | Coef. Std. Err. z P>|z| [95% Conf. Interval]
-----
    qsmk | -40.91185 187.6162 -0.22 0.827 -408.6328
                                               326.8091
    _cons | 13.15927 48.05103 0.27 0.784 -81.01901 107.3375
```

Instrumented: qsmk
Instruments: highprice

(1,476 real changes made, 1,476 to missing)

(1,476 real changes made)

Instrumental variables (2SLS) regression Number of obs = 1,476 
Wald chi2(1) = 0.55 
Prob > chi2 = 0.4576 
R-squared = .

Root MSE = 13.01

wt82_71		Std. Err.				_
qsmk	-21.10342	28.40885	-0.74	0.458	-76.78374 -6.188657	34.57691

Instrumented: qsmk
Instruments: highprice

(1,476 real changes made, 1,476 to missing)

(1,476 real changes made)

Instrumental variables (2SLS) regression Number of obs = 1,476

Wald chi2(1) = 0.29Prob > chi2 = 0.5880

R-squared = . Root MSE = 10.357

Instrumented: qsmk

Instruments: highprice

#### Program 16.5

- Estimating the average causal effect using the standard IV estimator conditional on baseline covariates
- Data from NHEFS
- Section 16.5

```
use ./data/nhefs-highprice, clear
replace highprice = .
replace highprice = (price82 >1.5 & price82 < .)</pre>
ivregress 2sls wt82_71 sex race c.age c.smokeintensity ///
 c.smokeyrs i.exercise i.active c.wt7 ///
 (qsmk = highprice)
(1,476 real changes made, 1,476 to missing)
(1,476 real changes made)
Instrumental variables (2SLS) regression
                                           Number of obs =
                                                              1,476
                                           Wald chi2(11) = 135.18
                                           Prob > chi2
                                                              0.0000
                                           R-squared
                                                        =
                                                               0.0622
                                           Root MSE
                                                               7.6848
    wt82_71 |
                Coef.
                        Std. Err.
                                   z P>|z|
                                                 [95% Conf. Interval]
      qsmk | -1.042295 29.86522 -0.03 0.972 -59.57705
                                                             57.49246
       sex | -1.644393 2.620115 -0.63 0.530 -6.779724
                                                             3.490938
      race | -.1832546  4.631443  -0.04  0.968  -9.260716  8.894207
       age | -.16364 .2395678 -0.68 0.495 -.6331844 .3059043
smokeinten~y | .0057669 .144911
                                  0.04
                                          0.968 -.2782534 .2897872
   smokeyrs |
             .0258357
                        .1607639 0.16
                                          0.872
                                                 -.2892558
                                                             .3409271
          -
   exercise |
             .4987479 2.162395
                                  0.23 0.818
        1 |
                                                 -3.739469
                                                             4.736964
        2 |
             .5818337 2.174255
                                   0.27
                                          0.789
                                                 -3.679628
                                                             4.843296
          - 1
     active |
                                  -1.93 0.053 -2.355908
        1 | -1.170145 .6049921
                                                             .0156176
        2 | -.5122842 1.303121
                                          0.694 -3.066355
                                  -0.39
                                                             2.041787
           wt71 | -.0979493
                                   -2.71
                         .036123
                                          0.007
                                                  -.168749
                                                            -.0271496
                         2.32589
                                  7.43 0.000
      _cons | 17.28033
                                                  12.72167 21.83899
```

Instrumented: qsmk

Instruments: sex race age smokeintensity smokeyrs 1.exercise 2.exercise

## 17. Causal survival analysis: Stata

#### Program 17.1

library(Statamarkdown)

- Nonparametric estimation of survival curves
- Data from NHEFS
- Section 17.1

```
use ./data/nhefs-formatted, clear

/*Some preprocessing of the data*/
gen survtime = .
replace survtime = 120 if death == 0
replace survtime = (yrdth - 83)*12 + modth if death ==1
* yrdth ranges from 83 to 92*

tab death qsmk

/*Kaplan-Meier graph of observed survival over time, by quitting smoking*/
*For now, we use the stset function in Stata*
stset survtime, failure(death=1)
sts graph, by(qsmk) xlabel(0(12)120)
qui gr export ./figs/stata-fig-17-1.png, replace

(1,566 missing values generated)

(1,275 real changes made)
```

#### (291 real changes made)

			death
	ng between	quit smoki	between
	and 1982	baseline	1983 and
Total	Smoking c	No smokin	1992
			+
1,275	312	963	0
291	91	200	1
			+
1,566	403	1,163	Total

failure event: death == 1
obs. time interval: (0, survtime]

exit on or before: failure

\_\_\_\_\_

1,566 total observations
0 exclusions

-----

1,566 observations remaining, representing

291 failures in single-record/single-failure data

171,076 total analysis time at risk and under observation

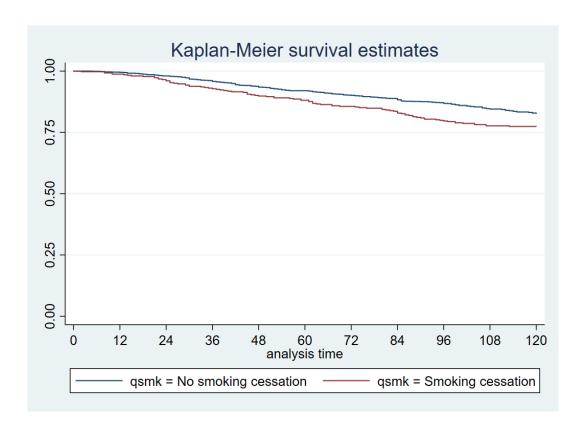
at risk from t = 0

0

earliest observed entry t =

last observed exit t = 120

failure \_d: death == 1
analysis time \_t: survtime



#### Program 17.2

- Parametric estimation of survival curves via hazards model
- Data from NHEFS
- Section 17.1
- Generates Figure 17.4

```
/**Create person-month dataset for survival analyses**/
/* We want our new dataset to include 1 observation per person
per month alive, starting at time = 0.
Individuals who survive to the end of follow-up will have
119 time points
Individuals who die will have survtime - 1 time points*/

use ./data/nhefs-formatted, clear

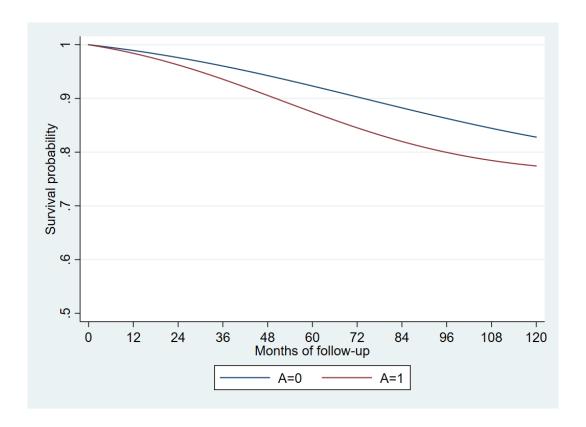
gen survtime = .
replace survtime = 120 if death == 0
replace survtime = (yrdth - 83)*12 + modth if death ==1

*expand data to person-time*
gen time = 0
expand survtime if time == 0
bysort seqn: replace time = _n - 1
```

```
*Create event variable*
gen event = 0
replace event = 1 if time == survtime - 1 & death == 1
tab event
*Create time-squared variable for analyses*
gen timesq = time*time
*Save the dataset to your working directory for future use*
qui save ./data/nhefs_surv, replace
/**Hazard ratios**/
use ./data/nhefs_surv, clear
*Fit a pooled logistic hazards model *
logistic event qsmk qsmk#c.time qsmk#c.time#c.time ///
  c.time c.time#c.time
/**Survival curves: run regression then do:**/
*Create a dataset with all time points under each treatment level*
*Re-expand data with rows for all timepoints*
drop if time != 0
expand 120 if time ==0
bysort seqn: replace time = _n - 1
/*Create 2 copies of each subject, and set outcome to missing
and treatment -- use only the newobs*/
expand 2 , generate(interv)
replace qsmk = interv
/*Generate predicted event and survival probabilities
for each person each month in copies*/
predict pevent_k, pr
gen psurv_k = 1-pevent_k
keep seqn time qsmk interv psurv_k
*Within copies, generate predicted survival over time*
*Remember, survival is the product of conditional survival probabilities in each interval*
sort seqn interv time
gen_t = time + 1
gen psurv = psurv_k if _t ==1
bysort seqn interv: replace psurv = psurv_k*psurv[_t-1] if _t >1
*Display 10-year standardized survival, under interventions*
*Note: since time starts at 0, month 119 is 10-year survival*
by interv, sort: summarize psurv if time == 119
```

```
*Graph of standardized survival over time, under interventions*
/*Note, we want our graph to start at 100% survival,
so add an extra time point with P(surv) = 1*/
expand 2 if time ==0, generate(newtime)
replace psurv = 1 if newtime == 1
gen time2 = 0 if newtime ==1
replace time2 = time + 1 if newtime == 0
/*Separate the survival probabilities to allow plotting by
intervention on qsmk*/
separate psurv, by(interv)
*Plot the curves*
twoway (line psurv0 time2, sort) ///
 (line psurv1 time2, sort) if interv > -1 ///
  , ylabel(0.5(0.1)1.0) xlabel(0(12)120) ///
 ytitle("Survival probability") xtitle("Months of follow-up") ///
 legend(label(1 "A=0") label(2 "A=1"))
qui gr export ./figs/stata-fig-17-2.png, replace
(1,566 missing values generated)
(1,275 real changes made)
(291 real changes made)
(169,510 observations created)
(169510 real changes made)
(291 real changes made)
     event | Freq. Percent
                                       Cum.
-----
         0 | 170,785
                           99.83
                                       99.83
              291
                            0.17
         1 |
                                     100.00
     Total | 171,076
                          100.00
file ./data/nhefs_surv.dta cannot be modified or erased; likely cause is
   read-only directory or file
r(608);
end of do-file
```

r(608);



#### Program 17.3

- Estimation of survival curves via IP weighted hazards model
- Data from NHEFS
- Section 17.4
- $\bullet~$  Generates Figure 17.6

```
keep seqn event qsmk time sex race age education ///
   smokeintensity smkintensity82_71 smokeyrs ///
   exercise active wt71
preserve

*Estimate weights*
logit qsmk sex race c.age##c.age ib(last).education ///
   c.smokeintensity##c.smokeintensity ///
   c.smokeyrs##c.smokeyrs ib(last).exercise ///
   ib(last).active c.wt71##c.wt71 if time == 0
predict p_qsmk, pr

logit qsmk if time ==0
predict num, pr
gen sw=num/p_qsmk if qsmk==1
replace sw=(1-num)/(1-p_qsmk) if qsmk==0
```

```
summarize sw
*IP weighted survival by smoking cessation*
logit event qsmk qsmk#c.time qsmk#c.time#c.time ///
  c.time c.time#c.time [pweight=sw] , cluster(seqn)
*Create a dataset with all time points under each treatment level*
*Re-expand data with rows for all timepoints*
drop if time != 0
expand 120 if time ==0
bysort seqn: replace time = _n - 1
/*Create 2 copies of each subject, and set outcome
to missing and treatment -- use only the newobs*/
expand 2 , generate(interv)
replace qsmk = interv
/*Generate predicted event and survival probabilities
for each person each month in copies*/
predict pevent_k, pr
gen psurv_k = 1-pevent_k
keep seqn time qsmk interv psurv_k
*Within copies, generate predicted survival over time*
/*Remember, survival is the product of conditional survival
probabilities in each interval*/
sort seqn interv time
gen_t = time + 1
gen psurv = psurv_k if _t ==1
bysort seqn interv: replace psurv = psurv_k*psurv[_t-1] if _t >1
*Display 10-year standardized survival, under interventions*
*Note: since time starts at 0, month 119 is 10-year survival*
by interv, sort: summarize psurv if time == 119
quietly summarize psurv if(interv==0 & time ==119)
matrix input observe = (0, r(mean)')
quietly summarize psurv if(interv==1 & time ==119)
matrix observe = (observe \1, r(mean)')
matrix observe = (observe \( \)3, observe [2,2] -observe [1,2])
matrix list observe
*Graph of standardized survival over time, under interventions*
/*Note: since our outcome model has no covariates,
we can plot psurv directly.
If we had covariates we would need to stratify or average across the values*/
expand 2 if time ==0, generate(newtime)
replace psurv = 1 if newtime == 1
gen time2 = 0 if newtime ==1
```

```
replace time2 = time + 1 if newtime == 0
separate psurv, by(interv)
twoway (line psurv0 time2, sort) ///
  (line psurv1 time2, sort) if interv > -1 ///
  , ylabel(0.5(0.1)1.0) xlabel(0(12)120) ///
 ytitle("Survival probability") xtitle("Months of follow-up") ///
 legend(label(1 "A=0") label(2 "A=1"))
qui gr export ./figs/stata-fig-17-3.png, replace
*remove extra timepoint*
drop if newtime == 1
drop time2
restore
**Bootstraps**
qui save ./data/nhefs_std1 , replace
capture program drop bootipw_surv
program define bootipw_surv , rclass
use ./data/nhefs std1 , clear
preserve
bsample, cluster(seqn) idcluster(newseqn)
logit qsmk sex race c.age##c.age ib(last).education ///
  c.smokeintensity##c.smokeintensity ///
    c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active ///
   c.wt71##c.wt71 if time == 0
predict p_qsmk, pr
logit qsmk if time ==0
predict num, pr
gen sw=num/p_qsmk if qsmk==1
replace sw=(1-num)/(1-p_qsmk) if qsmk==0
logit event qsmk qsmk#c.time qsmk#c.time#c.time ///
 c.time c.time#c.time [pweight=sw], cluster(newseqn)
drop if time != 0
expand 120 if time ==0
bysort newseqn: replace time = _n - 1
expand 2 , generate(interv_b)
replace qsmk = interv_b
predict pevent_k, pr
gen psurv_k = 1-pevent_k
keep newseqn time qsmk interv_b psurv_k
```

```
sort newsegn interv b time
gen _t = time + 1
gen psurv = psurv_k if _t ==1
bysort newseqn interv_b: ///
 replace psurv = psurv_k*psurv[_t-1] if _t >1
drop if time != 119
bysort interv_b: egen meanS_b = mean(psurv)
keep newseqn qsmk meanS_b
drop if newseqn != 1 /* only need one pair */
drop newseqn
return scalar boot_0 = meanS_b[1]
return scalar boot_1 = meanS_b[2]
return scalar boot_diff = return(boot_1) - return(boot_0)
restore
end
set rmsg on
simulate PrY_a0 = r(boot_0) PrY_a1 = r(boot_1) ///
 difference=r(boot_diff), reps(10) seed(1): bootipw_surv
set rmsg off
matrix pe = observe[1..3, 2]'
bstat, stat(pe) n(1629)
Iteration 0:
             log likelihood = -893.02712
Iteration 1: log likelihood = -839.70016
Iteration 2: log likelihood = -838.45045
Iteration 3: log likelihood = -838.44842
Iteration 4: log likelihood = -838.44842
                                          Number of obs =
Logistic regression
                                                               1,566
                                          LR chi2(18)
                                                              109.16
                                          Prob > chi2
                                                               0.0000
Log likelihood = -838.44842
                                          Pseudo R2
                                                               0.0611
               Coef. Std. Err. z P>|z| [95% Conf. Interval]
      qsmk |
______
       sex | -.5274782 .1540497 -3.42 0.001
                                                   -.82941 -.2255463
       race | -.8392636 .2100668 -4.00 0.000 -1.250987 -.4275404
             .1212052 .0512663 2.36 0.018 .0207251
       age |
                                                             .2216853
 c.age#c.age | -.0008246 .0005361 -1.54 0.124 -.0018753 .0002262
  education |
        1 | -.4759606 .2262238
                                   -2.10 0.035 -.9193511 -.0325701
```

2   3	5047361 3895288	.217597 .1914353	-2.32 -2.03	0.020 0.042	9312184 7647351	0782538 0143226
4   	4123596	.2772868	-1.49	0.137	9558318	.1311126
smokeinten~y	0772704	.0152499	-5.07	0.000	1071596	0473812
c. l						
smokeinten~y#						
c.						
smokeinten~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
smokeyrs	0735966	.0277775	-2.65	0.008	1280395	0191538
c.smokeyrs#						
c.smokeyrs	.0008441	.0004632	1.82	0.068	0000637	.0017519
exercise						
0	395704	.1872401	-2.11	0.035	7626878	0287201
1	0408635	.1382674	-0.30	0.768	3118627	.2301357
active						
0	176784	.2149721	-0.82	0.411	5981215	.2445535
1	1448395	.2111472	-0.69	0.493	5586806	.2690015
i						
wt71	0152357	.0263161	-0.58	0.563	0668144	.036343
c.wt71#						
c.wt71	.0001352	.0001632	0.83	0.407	0001846	.000455
_cons	-1.19407	1.398493	-0.85	0.393	-3.935066	1.546925

Iteration 0: log likelihood = -893.02712
Iteration 1: log likelihood = -893.02712

Logistic regression Number of obs = 1,566LR chi2(0) = -0.00

Prob > chi2 = ...

 $\label{eq:log_likelihood} \mbox{Log likelihood} = -893.02712 \qquad \qquad \mbox{Pseudo R2} \qquad = \quad -0.0000$ 

#### (128,481 missing values generated)

#### (128,481 real changes made)

Variable	0bs	Mean	Std. Dev.	Min	Max
	<b></b>				
sw	171,076	1.000509	. 2851505	.3312489	4.297662

Iteration 0: log pseudolikelihood = -2136.3671
Iteration 1: log pseudolikelihood = -2127.0974
Iteration 2: log pseudolikelihood = -2126.8556
Iteration 3: log pseudolikelihood = -2126.8554

Logistic regression Number of obs = 171,076 Wald chi2(5) = 22.74 Prob > chi2 = 0.0004 Log pseudolikelihood = -2126.8554 Pseudo R2 = 0.0045

(Std. Err. adjusted for 1,566 clusters in seqn)

1		Robust				
event	Coef.	Std. Err.	z	P> z	[95% Conf.	Interval]
qsmk	1301273	.4186673	-0.31	0.756	9507002	.6904456
qsmk#c.time						
Smoking c	.01916	.0151318	1.27	0.205	0104978	.0488178
qsmk#c.time#						
c.time						
Smoking c	0002152	.0001213	-1.77	0.076	0004528	.0000225
time	.0208179	.0077769	2.68	0.007	.0055754	.0360604
c.time#						
c.time	0001278	.0000643	-1.99	0.047	0002537	-1.84e-06
_cons	-7.038847	.2142855	-32.85	0.000	-7.458839	-6.618855

(169,510 observations deleted)

(186,354 observations created)

(186354 real changes made)

(187,920 observations created)

(187,920 real changes made)

(372,708 missing values generated)

(372708 real changes made)

\_\_\_\_\_

-> interv = 0

\_\_\_\_\_\_

-> interv = 1

observe[3,2]

c1 c2 r1 0 .8161003 r2 1 .81167841 r3 3 -.00442189

(3,132 observations created)

(3,132 real changes made)

(375,840 missing values generated)

(375,840 real changes made)

storage display value variable name type format label variable label

-----

psurv0	float	%9.0g	psurv,	interv == 0
psurv1	float	%9.0g	psurv,	interv == 1

(3,132 observations deleted)

r; t=0.00 8:35:20

command: bootipw\_surv
PrY\_a0: r(boot\_0)
PrY\_a1: r(boot\_1)
difference: r(boot\_diff)

Simulations (10)

. . . . . . . . . .

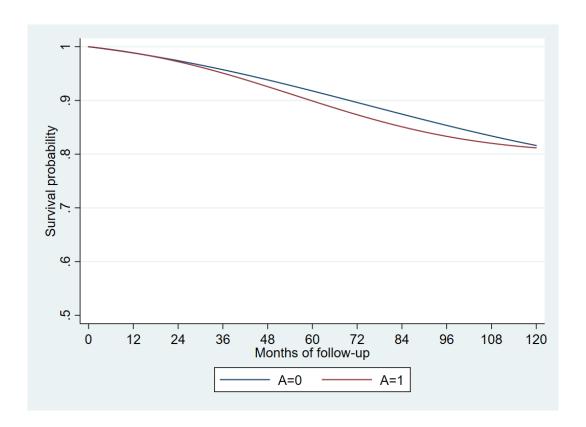
r; t=24.21 8:35:44

Bootstrap results Number of obs = 1,629Replications = 10

-----

	Coef.	Bootstrap Std. Err.		• •	[95% Conf.	-based Interval]
PrY_a0	.8161003 .8116784	.0093124 .0237581	87.64 34.16	0.000	.7978484 .7651133 0485224	.8343522 .8582435 .0396786

201



#### Program 17.4

- Estimating of survival curves via g-formula
- Data from NHEFS
- Section 17.5
- $\bullet~$  Generates Figure 17.7

```
keep seqn event qsmk time sex race age education ///
smokeintensity smkintensity82_71 smokeyrs exercise ///
active wt71
preserve

quietly logistic event qsmk qsmk#c.time ///
qsmk#c.time#c.time time c.time#c.time ///
sex race c.age##c.age ib(last).education ///
c.smokeintensity##c.smokeintensity ///
c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active ///
c.wt71##c.wt71 , cluster(seqn)
drop if time != 0
expand 120 if time ==0
bysort seqn: replace time = _n - 1
expand 2 , generate(interv)
```

```
replace qsmk = interv
predict pevent_k, pr
gen psurv_k = 1-pevent_k
keep seqn time qsmk interv psurv_k
sort seqn interv time
gen_t = time + 1
gen psurv = psurv_k if _t ==1
bysort seqn interv: replace psurv = psurv_k*psurv[_t-1] if _t >1
by interv, sort: summarize psurv if time == 119
keep qsmk interv psurv time
bysort interv : egen meanS = mean(psurv) if time == 119
by interv: summarize meanS
quietly summarize meanS if(qsmk==0 & time ==119)
matrix input observe = ( 0, r(mean)')
quietly summarize meanS if(qsmk==1 & time ==119)
matrix observe = (observe \1, `r(mean)')
matrix observe = (observe \2, observe[2,2]-observe[1,2])
*Add some row/column descriptions and print results to screen*
matrix rownames observe = P(Y(a=0)=1) P(Y(a=1)=1) difference
matrix colnames observe = interv survival
*Graph standardized survival over time, under interventions*
/*Note: unlike in Program 17.3, we now have covariates
so we first need to average survival across strata*/
bysort interv time : egen meanS_t = mean(psurv)
*Now we can continue with the graph*
expand 2 if time ==0, generate(newtime)
replace meanS_t = 1 if newtime == 1
gen time2 = 0 if newtime ==1
replace time2 = time + 1 if newtime == 0
separate meanS_t, by(interv)
twoway (line meanS_t0 time2, sort) ///
  (line meanS_t1 time2, sort) ///
  , ylabel(0.5(0.1)1.0) xlabel(0(12)120) ///
 ytitle("Survival probability") xtitle("Months of follow-up") ///
 legend(label(1 "A=0") label(2 "A=1"))
gr export ./figs/stata-fig-17-4.png, replace
*remove extra timepoint*
drop if newtime == 1
restore
*Bootstraps*
```

```
qui save ./data/nhefs_std2 , replace
capture program drop bootstdz_surv
program define bootstdz_surv , rclass
use ./data/nhefs_std2 , clear
preserve
bsample, cluster(seqn) idcluster(newseqn)
logistic event qsmk qsmk#c.time qsmk#c.time#c.time ///
 time c.time#c.time ///
   sex race c.age##c.age ib(last).education ///
   c.smokeintensity##c.smokeintensity c.smkintensity82_71 ///
   c.smokeyrs##c.smokeyrs ib(last).exercise ib(last).active ///
   c.wt71##c.wt71
drop if time != 0
/*only predict on new version of data */
expand 120 if time ==0
bysort newseqn: replace time = _n - 1
expand 2 , generate(interv_b)
replace qsmk = interv_b
predict pevent_k, pr
gen psurv_k = 1-pevent_k
keep newseqn time qsmk psurv_k
sort newseqn qsmk time
gen_t = time + 1
gen psurv = psurv_k if _t ==1
bysort newseqn qsmk: replace psurv = psurv_k*psurv[_t-1] if _t >1
drop if time != 119  /* keep only last observation */
keep newseqn qsmk psurv
/* if time is in data for complete graph add time to bysort */
bysort qsmk : egen meanS_b = mean(psurv)
keep newseqn qsmk meanS_b
drop if newseqn != 1 /* only need one pair */
drop newseqn
return scalar boot_0 = meanS_b[1]
return scalar boot_1 = meanS_b[2]
return scalar boot_diff = return(boot_1) - return(boot_0)
restore
end
set rmsg on
simulate PrY_a0 = r(boot_0) PrY_a1 = r(boot_1) ///
 difference=r(boot_diff), reps(10) seed(1): bootstdz_surv
set rmsg off
matrix pe = observe[1..3, 2]'
```

```
bstat, stat(pe) n(1629)
(169,510 observations deleted)
(186,354 observations created)
(186354 real changes made)
(187,920 observations created)
(187,920 real changes made)
(372,708 missing values generated)
(372708 real changes made)
-> interv = 0
  Variable | Obs Mean Std. Dev. Min
     psurv |
             1,566 .8160697 .2014345 .014127 .9903372
-> interv = 1
  Variable | Obs Mean Std. Dev. Min Max
    psurv | 1,566 .811763 .2044758 .0123403 .9900259
(372708 missing values generated)
 ______
-> interv = 0
  Variable | Obs Mean Std. Dev. Min
                                                  Max
     meanS | 1,566 .8160697
                               0 .8160697 .8160697
```

#### -> interv = 1

Min Max	Dev.	Std.	Mean	0bs		Variable
					+	
.8117629 .8117629	0 .81		.8117629	,566	1	meanS

(3,132 observations created)

(3,132 real changes made)

(375,840 missing values generated)

(375,840 real changes made)

variable name	storage type	display format	value label	variable label
meanS_t0 meanS_t1	float float			<pre>meanS_t, interv == 0 meanS_t, interv == 1</pre>

(file ./figs/stata-fig-17-4.png written in PNG format)

(3,132 observations deleted)

r; t=0.00 8:35:52

command: bootstdz\_surv
PrY\_a0: r(boot\_0)
PrY\_a1: r(boot\_1)
difference: r(boot\_diff)

Simulations (10)

. . . . . . . . . .

r; t=27.27 8:36:20

Bootstrap results Number of obs = 1,629Replications =

     	Coef.	Bootstrap Std. Err.			[95% Conf.	-based Interval]
PrY_a0	.8160697 .8117629	.0087193 .0292177 .0307674	93.59 27.78 -0.14	0.000 0.000 0.889	.7989802 .7544973 0646099	.8331593 .8690286 .0559963

### Session information: Stata

library(Statamarkdown)

```
For reproducibility.
about
Revision 03 Feb 2020
Copyright 1985-2017 StataCorp LLC
Total physical memory:
                       33429480 KB
Available physical memory: 20132500 KB
94-user 2-core Stata network perpetual license:
      Serial number: 501506248171
       Licensed to: Tom Palmer
                    University of Bristol
# install.packages("sessioninfo")
sessioninfo::session_info()
- Session info -----
setting value
version R version 4.0.0 (2020-04-24)
        Windows 10 x64
system x86_64, mingw32
ui
        RTerm
language (EN)
collate English_United Kingdom.1252
ctype
        English_United Kingdom.1252
tz
        Europe/London
        2020-05-14
date
- Packages ------
package
            * version date
                                lib source
             0.2.1 2019-03-21 [1] CRAN (R 4.0.0)
assertthat
bookdown
             0.18 2020-03-05 [1] CRAN (R 4.0.0)
```

```
cli
                2.0.2
                        2020-02-28 [1] CRAN (R 4.0.0)
crayon
                1.3.4
                        2017-09-16 [1] CRAN (R 4.0.0)
                0.6.25
                        2020-02-23 [1] CRAN (R 4.0.0)
digest
evaluate
                0.14
                        2019-05-28 [1] CRAN (R 4.0.0)
fansi
                0.4.1
                        2020-01-08 [1] CRAN (R 4.0.0)
                        2020-04-03 [1] CRAN (R 4.0.0)
glue
                1.4.0
                        2019-10-04 [1] CRAN (R 4.0.0)
htmltools
                0.4.0
                1.28
                        2020-02-06 [1] CRAN (R 4.0.0)
knitr
                1.5
                        2014-11-22 [1] CRAN (R 4.0.0)
magrittr
                1.0.4.6 2020-04-09 [1] CRAN (R 4.0.0)
Rcpp
                0.4.6
                        2020-05-02 [1] CRAN (R 4.0.0)
rlang
                2.1
                        2020-01-20 [1] CRAN (R 4.0.0)
rmarkdown
sessioninfo
                1.1.1
                        2018-11-05 [1] CRAN (R 4.0.0)
                        2020-04-26 [1] Github (Hemken/Statamarkdown@506cfc9)
Statamarkdown * 0.4.5
                        2020-02-17 [1] CRAN (R 4.0.0)
stringi
                1.4.6
                        2019-02-10 [1] CRAN (R 4.0.0)
                1.4.0
stringr
                2.2.0
                        2020-04-20 [1] CRAN (R 4.0.0)
withr
                0.13
                        2020-04-13 [1] CRAN (R 4.0.0)
xfun
                2.2.1
                        2020-02-01 [1] CRAN (R 4.0.0)
yaml
```

- [1] C:/Users/tom/Documents/R/win-library/4.0
- [2] C:/Program Files/R/R-4.0.0/library

## Bibliography

Miguel A Hernán and James M Robins. Causal Inference: What If. Boca Raton: Chapman & Hall/CRC, 2020.