

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

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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.

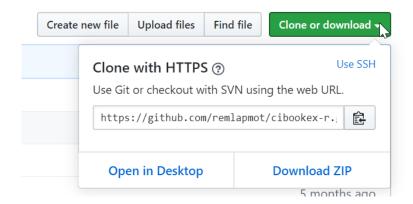
This repo is rendered at https://remlapmot.github.io/cibookex-r/. Click the download button above for the pdf and eBook versions.



### 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

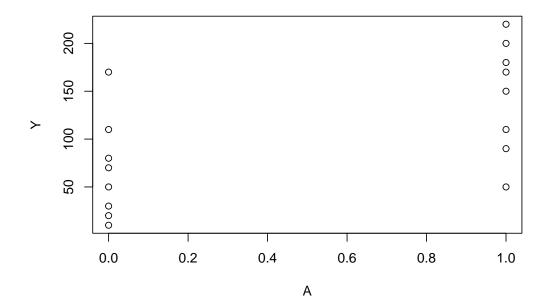
## 11. Why model?

### 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, 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)

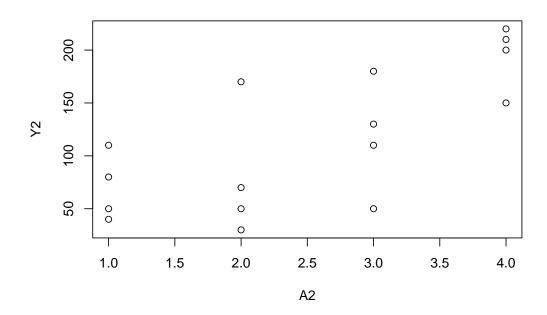
plot(A, Y)
```



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

## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 10.0 27.5 60.0 67.5 87.5 170.0

summary(Y[A == 1])
```



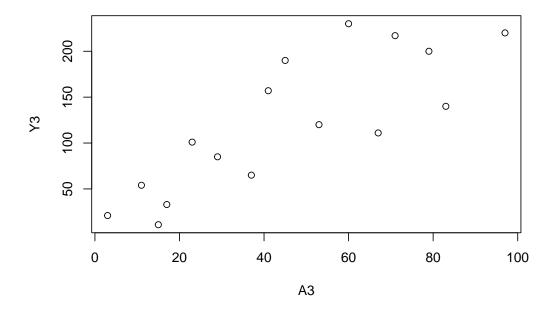
```
summary(Y2[A2 == 1])
     Min. 1st Qu. Median
##
                             Mean 3rd Qu.
                                             Max.
     40.0 47.5
##
                     65.0
                             70.0
                                     87.5
                                            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.
      50.0
             95.0
                   120.0
                            117.5 142.5
                                            180.0
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
- $\bullet$  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:
##
       Min
                 1Q
                      Median
                                   ЗQ
                                           Max
## -61.930 -30.564
                      -5.741
                               30.653
                                        77.225
##
## Coefficients:
##
              Estimate Std. Error t value Pr(>|t|)
## (Intercept) 24.5464
                           21.3300
                                     1.151 0.269094
## A3
                 2.1372
                            0.3997
                                     5.347 0.000103 ***
## 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))
##
       1
## 216.89
summary(glm(Y ~ A))
##
## Call:
## glm(formula = Y ~ A)
## Deviance Residuals:
      Min
                10
                    Median
                                  3Q
## -96.250 -40.000
                    3.125 35.938 102.500
##
## Coefficients:
              Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                 67.50
                            19.72
                                    3.424 0.00412 **
                 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
```

- 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
## 197.1269
```

# 12. IP Weighting and Marginal Structural Models

### Program 12.1

• Descriptive statistics from NHEFS data (Table 12.1)

```
library(here)
# install.packages("readxl") # install package if required
library("readxl")
nhefs <- read_excel(here("data", "NHEFS.xls"))</pre>
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)</pre>
# 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)
##
## 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))
##
## 4.525079
# No smoking cessation
predict(lm(wt82_71 ~ qsmk, data = nhefs.nmv), data.frame(qsmk = 0))
## 1.984498
```

```
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.
##
             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.
##
      1.00
             15.00
                     23.00
                             24.09
                                     32.00
                                              64.00
summary(nhefs.nmv[which(nhefs.nmv$qsmk == 1),]$age)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
##
     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.
##
     39.58
             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
                                              60.00
                                      35.00
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
                           1
     0 0.85382631 0.14617369
##
     1 0.91066998 0.08933002
##
table(nhefs.nmv$qsmk, nhefs.nmv$education)
##
##
         1
             2
##
     0 210 266 480 92 115
     1 81 74 157 29 62
prop.table(table(nhefs.nmv$qsmk, nhefs.nmv$education), 1)
##
##
                1
                           2
                                       3
     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
                                    2
                         1
##
     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 \sim 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)
##
## 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:
##
      Min
                1Q
                     Median
                                  3Q
                                         Max
  -1.5127 -0.7907 -0.6387
                              0.9832
                                       2.3729
##
## Coefficients:
                          Estimate Std. Error z value Pr(>|z|)
##
                        -2.2425191 1.3808360 -1.624 0.104369
## (Intercept)
                        -0.5274782  0.1540496  -3.424  0.000617 ***
## sex
                        ## race
                         0.1212052 0.0512663
                                              2.364 0.018068 *
## age
## I(age^2)
                        -0.0008246 0.0005361 -1.538 0.124039
## 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
                        -0.0735966  0.0277775  -2.650  0.008061 **
## smokeyrs
                         0.0008441 0.0004632 1.822 0.068398 .
## I(smokeyrs^2)
## as.factor(exercise)1
                         0.3548405 0.1801351 1.970 0.048855 *
                         0.3957040 0.1872400 2.113 0.034571 *
## as.factor(exercise)2
## as.factor(active)1
                         0.0319445 0.1329372
                                               0.240 0.810100
## as.factor(active)2
                         0.1767840 0.2149720
                                               0.822 0.410873
## wt71
                        -0.0152357 0.0263161 -0.579 0.562625
## 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</pre>
summary(nhefs.nmv$w)
##
      Min. 1st Qu. Median
                             Mean 3rd Qu.
                                              Max.
##
           1.230
                   1.373 1.996 1.990 16.700
sd(nhefs.nmv$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 ***
## qsmk
                 3.4405 0.5255 42.87 5.86e-11 ***
## ---
## 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)
SE <- coef(summary(msm.w))[, 2]</pre>
lcl \leftarrow beta - qnorm(0.975) * SE
ucl \leftarrow beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
                       lcl ucl
##
                beta
## (Intercept) 1.780 1.340 2.22
## qsmk
               3.441 2.411 4.47
# 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
    54 17 9
##
    55 9 4
##
    56 8 7
##
##
    57 9 2
    58 8 4
##
##
    59 5 4
##
    60 5 4
##
    61 5 2
    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|)
## (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
                                                -0.15 0.88465
                                     0.198351
## as.factor(education)3 0.086432
                                                 0.49 0.62744
                                     0.178085
## 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)
                          0.001045
                                     0.000287
                                                 3.65 0.00027 ***
## 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
                          0.395704
                                     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
                         -0.015236
## wt71
                                                -0.58 0.56262
                                     0.026316
## 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
                               30
                                      Max
## -0.771 -0.771 -0.771
                            1.648
                                    1.648
##
## Coefficients:
##
               Estimate Std. Error z value Pr(>|z|)
## (Intercept) -1.0598
                            0.0578
                                     -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: 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|)
## (Intercept) 1.780
                         0.225 62.7 2.3e-15 ***
                 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]
lcl <- beta - qnorm(0.975) * SE</pre>
ucl \leftarrow beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
               beta lcl ucl
## (Intercept) 1.78 1.34 2.22
               3.44 2.41 4.47
## qsmk
# 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 ciq/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</pre>
```

```
summary(nhefs.nmv.s$sw.a)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
##
      0.19
              0.89
                      0.97
                               1.00
                                               5.10
                                       1.05
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|)
## (Intercept)
                                              2.00452 0.29512 46.13 1.1e-11 ***
## 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 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Correlation structure = independence
## Estimated Scale Parameters:
##
##
               Estimate Std.err
## (Intercept)
                   60.5
                            4.5
## Number of clusters:
                         1162 Maximum cluster size: 1
beta <- coef(msm.sw.cont)</pre>
SE <- coef(summary(msm.sw.cont))[, 2]
lcl \leftarrow beta - qnorm(0.975) * SE
ucl \leftarrow beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
##
                                                            lcl
                                                                     ucl
                                                 beta
## (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(</pre>
  death ~ qsmk,
 data = nhefs.nmv,
  weights = sw,
 id = seqn,
 family = binomial(),
  corstr = "independence"
)
## 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
                       1 0.0678
## (Intercept)
## Number of clusters:
                          1566 Maximum cluster size: 1
beta <- coef(msm.logistic)</pre>
SE <- coef(summary(msm.logistic))[, 2]</pre>
lcl <- beta - qnorm(0.975) * SE</pre>
ucl \leftarrow beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)
##
                  beta
                           lcl
                                  ucl
## (Intercept) -1.4905 -1.645 -1.336
## qsmk
                0.0301 -0.278 0.338
```

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

## as.factor(exercise)2

```
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:
##
     Min
               1Q Median
                                      Max
  -1.513 -0.791 -0.639
                            0.983
                                    2.373
##
## Coefficients:
                          Estimate Std. Error z value Pr(>|z|)
##
## (Intercept)
                                     1.380836
                                              -1.62 0.10437
                         -2.242519
## 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 ***
                                                 2.36 0.01807 *
## age
                          0.121205
                                     0.051266
## I(age^2)
                         -0.000825
                                     0.000536
                                                -1.54 0.12404
## as.factor(education)2 -0.028776
                                                -0.15 0.88465
                                     0.198351
## as.factor(education)3 0.086432
                                     0.178085
                                                 0.49 0.62744
## as.factor(education)4  0.063601
                                                 0.23 0.81592
                                     0.273211
## as.factor(education)5 0.475961
                                     0.226224
                                                 2.10 0.03538 *
                                                -5.07 4.0e-07 ***
## smokeintensity
                         -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
                                                 1.82 0.06840 .
                                     0.000463
## as.factor(exercise)1
                          0.354841
                                     0.180135
                                                 1.97 0.04885 *
```

2.11 0.03457 \*

0.187240

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|)
## (Intercept)
                 -0.9016
                               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|)
##
## (Intercept)
                                     1.78445 0.30984 33.17 8.5e-09 ***
## 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
                                                                 0.88
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Correlation structure = independence
## Estimated Scale Parameters:
##
##
               Estimate Std.err
## (Intercept)
                   60.8
                           3.71
## 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 \leftarrow 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(gsmk)1:as.factor(sex)1 -0.15948 -2.210 1.891
```

- 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
              25
summary(nhefs[which(nhefs$cens == 0),]$wt71)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
##
      39.6
              59.5
                      69.2
                              70.8
                                       79.8
                                              151.7
summary(nhefs[which(nhefs$cens == 1),]$wt71)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
      36.2
              63.1
                      72.1
                              76.6
                                      87.9
##
                                              169.2
# 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:
               1Q Median
                               3Q
                                      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)
                                                3.07 0.00216 **
                         0.000846
                                    0.000276
## 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
                                                1.97 0.04846 *
                         0.355052
                                    0.179929
## as.factor(active)1
                         0.010875
                                    0.129832
                                                0.08 0.93324
## 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
                              3Q
                                     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
## ---
## 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:
##
               1Q Median
                               30
                                      Max
## -1.097 -0.287 -0.207 -0.157
                                     2.996
## Coefficients:
##
                          Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                          4.014466
                                     2.576106
                                                  1.56
                                                         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
## age
                         -0.269729
                                     0.117465
                                                 -2.30
                                                         0.0217 *
## I(age^2)
                          0.002884
                                     0.001114
                                                  2.59
                                                         0.0096 **
## as.factor(education)2 -0.440788
                                     0.419399
                                                -1.05
                                                         0.2933
## as.factor(education)3 -0.164688
                                     0.370547
                                                 -0.44
                                                         0.6567
## as.factor(education)4 0.138447
                                     0.569797
                                                  0.24
                                                         0.8080
## as.factor(education)5 -0.382382
                                     0.560181
                                                 -0.68
                                                         0.4949
## smokeintensity
                          0.015712
                                     0.034732
                                                  0.45
                                                         0.6510
## I(smokeintensity^2)
                         -0.000113
                                     0.000606
                                                 -0.19
                                                         0.8517
## smokeyrs
                          0.078597
                                     0.074958
                                                  1.05
                                                         0.2944
## 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
                                                 -1.57
                                     0.372313
                                                         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
                                               -2.20
                        -0.087887
                                    0.040012
                                                     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
                              3Q
                                     Max
## -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 ***
## as.factor(qsmk)1
                      0.641
                                 0.264
                                          2.43
                                                  0.015 *
## 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(nhefsqsmk == 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</pre>
summary(nhefs$sw.a)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
##
      0.33
              0.86
                      0.95
                              1.00
                                               4.21
                                       1.08
sd(nhefs$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(</pre>
 wt82_71 \sim qsmk,
 data = nhefs,
 weights = sw,
 id = seqn,
  corstr = "independence"
summary(msm.sw)
##
## Call:
## geeglm(formula = wt82_71 \sim qsmk, data = nhefs, weights = sw,
       id = seqn, corstr = "independence")
##
## Coefficients:
               Estimate Std.err Wald Pr(>|W|)
##
## (Intercept) 1.662 0.233 51.0 9.3e-13 ***
                          0.526 44.2 2.9e-11 ***
## qsmk
                  3.496
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## Correlation structure = independence
## Estimated Scale Parameters:
##
```

```
## Estimate Std.err
## (Intercept) 61.8 3.83
## Number of clusters: 1566 Maximum cluster size: 1
beta <- coef(msm.sw)
SE <- coef(summary(msm.sw))[, 2]
lcl <- beta - qnorm(0.975) * SE
ucl <- beta + qnorm(0.975) * SE
cbind(beta, lcl, ucl)

## beta lcl ucl
## (Intercept) 1.66 1.21 2.12
## qsmk 3.50 2.47 4.53</pre>
```

## 13. Standardization and the parametric G-formula

## Program 13.1

- 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"))</pre>
# some preprocessing of the data
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)</pre>
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)
##
## 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) + 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 **
                                   ## sex
## race
                                    0.5601096 0.5818888 0.963 0.335913
                                    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
                                    0.0491365 0.0517254
## smokeintensity
                                                        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
                                   ## 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)
## ATC: 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
                                          age education smokeintensity smokeyrs
     predicted.meanY qsmk
                             sex race
##
               <dbl> <dbl> <dbl> <dbl> <dbl> <
                                                  <dbl>
                                                                  <dbl>
                                                                           <dbl>
                         0
                                                                              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
                                      4.511
                                              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
```

## Program 13.2

- 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 <- rep(-1, N)
observed <- cbind(L, A, Y, interv)
```

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

## Program 13.3

- 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$interv <- 1
interv1$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 \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 = onesample
)
summary(std)
##
## 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 = onesample)
##
## 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
                                     -1.4302717   0.4689576   -3.050   0.002328 **
## sex
                                      0.5601096 0.5818888 0.963 0.335913
## race
                                      ## age
```

```
## I(age * age)
                                      -0.0061010 0.0017261 -3.534 0.000421 ***
## as.factor(education)2
                                       0.7904440 0.6070005
                                                            1.302 0.193038
## 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
                                      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
## wt71
                                      0.0455018 0.0833709
                                                              0.546 0.585299
## 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
```

## Program 13.4

- 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
  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$wt82_71 <- NA
  d.onesample <- rbind(d, d0, d1) # combining datasets</pre>
  # linear model to estimate mean outcome conditional on treatment and confounders
  # parameters are estimated using original observations only (interv= -1)
  # parameter estimates are used to predict mean outcome for observations with set
  # treatment (interv=0 and interv=1)
  fit <- glm(</pre>
   wt82_71 ~ qsmk + sex + race + age + I(age * age) +
      as.factor(education) + smokeintensity +
      I(smokeintensity * smokeintensity) + smokeyrs + I(smokeyrs *
                                                           smokeyrs) +
      as.factor(exercise) + as.factor(active) + wt71 + I(wt71 *
                                                            wt71),
   data = d.onesample
  )
  d.onesample$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]),
```

```
sd(results$t[, 2]),
        sd(results$t[, 3]),
        sd(results$t[, 4]))
mean <- results$t0</pre>
11 \leftarrow mean - qnorm(0.975) * se
ul \leftarrow mean + qnorm(0.975) * se
bootstrap <-
  data.frame(cbind(
    с(
      "Observed",
      "No Treatment",
      "Treatment",
      "Treatment - No Treatment"
    ),
    mean,
    se,
    11,
    ul
  ))
bootstrap
```

## 14. G-estimation of Structural Nested Models

## Program 14.1

- 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"))</pre>
# some processing of the data
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)</pre>
# ranking of extreme observations
#install.packages("Hmisc")
library(Hmisc)
## 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
                                                                .05
                                                      Gmd
                                                                         .10
##
       1566
                  63
                        1510
                                           2.638
                                                    8.337
                                                            -9.752
                                                                     -6.292
       . 25
                .50
                        .75
                                    .90
                                             .95
    -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)
                    + 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 ~ 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
                     Median
                1Q
                                 3Q
                                         Max
## -2.9959
            0.1571
                     0.2069
                             0.2868
                                      1.0967
##
## Coefficients:
##
                          Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                        -4.0144661 2.5761058 -1.558 0.11915
## qsmk
                        -0.5168674 0.2877162 -1.796 0.07242
## sex
                        -0.0573131 0.3302775
                                             -0.174 0.86223
## race
                        0.0122715 0.4524887
                                               0.027 0.97836
## age
                         0.2697293 0.1174647
                                               2.296 0.02166 *
## I(age^2)
                        1.051 0.29326
## as.factor(education)2 0.4407884 0.4193993
## 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
## smokeintensity
                       -0.0157119 0.0347319 -0.452 0.65100
## I(smokeintensity^2)
                        0.0001133 0.0006058
                                              0.187 0.85171
## smokeyrs
                        -0.0785973 0.0749576 -1.049 0.29438
## 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$pd.c <- predict(cw.denom, nhefs, type="response")
nhefs$wc <- ifelse(nhefs$cens==0, 1/nhefs$pd.c, NA)
# observations with cens=1 only contribute to censoring models</pre>
```

### 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

```
#install.packages("geepack")
library("geepack")
nhefs$psi <- 3.446
nhefs$Hpsi <- nhefs$wt82_71 - nhefs$psi*nhefs$qsmk</pre>
fit <- geeglm(qsmk ~ sex + race + age + I(age*age) + as.factor(education)
           + smokeintensity + I(smokeintensity*smokeintensity) + smokeyrs
           + I(smokeyrs*smokeyrs) + as.factor(exercise) + as.factor(active)
           + wt71 + I(wt71*wt71) + Hpsi, family=binomial, data=nhefs,
           weights=wc, id=seqn, corstr="independence")
## Warning in eval(family$initialize): non-integer #successes in a binomial glm!
summary(fit)
##
## Call:
## 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 .
                                      -5.137e-01 1.536e-01 11.193 0.000821 ***
## sex
                                      -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
## smokeintensity
                                     -7.834e-02 1.464e-02 28.635 8.74e-08 ***
## 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 .
                                      3.800e-01 1.889e-01 4.049 0.044187 *
## as.factor(exercise)2
## as.factor(active)1
                                      3.412e-02 1.339e-01 0.065 0.798778
                                      2.135e-01 2.121e-01 1.012 0.314308
## as.factor(active)2
## wt71
                                    -7.661e-03 2.562e-02 0.089 0.764963
## I(wt71 * wt71)
                                     8.655e-05 1.582e-04 0.299 0.584233
                                     -1.903e-06 8.839e-03 0.000 0.999828
## Hpsi
## ---
## 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 \leftarrow seq(from = 2,to = 5, by = 0.1)
j = 0
Hpsi.coefs <- cbind(rep(NA,length(grid)), rep(NA, length(grid)))</pre>
colnames(Hpsi.coefs) <- c("Estimate", "p-value")</pre>
for (i in grid){
 psi = i
 j = j+1
  nhefs$Hpsi <- nhefs$wt82_71 - psi * nhefs$qsmk</pre>
  gest.fit <- geeglm(qsmk ~ sex + race + age + I(age*age) + as.factor(education)</pre>
                   + smokeintensity + I(smokeintensity*smokeintensity) + smokeyrs
                   + I(smokeyrs*smokeyrs) + as.factor(exercise) + as.factor(active)
                   + wt71 + I(wt71*wt71) + Hpsi, family=binomial, data=nhefs,
                   weights=wc, id=seqn, corstr="independence")
  Hpsi.coefs[j,1] <- summary(gest.fit)$coefficients["Hpsi", "Estimate"]</pre>
  Hpsi.coefs[j,2] <- summary(gest.fit)$coefficients["Hpsi", "Pr(>|W|)"]
}
```

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

```
logit.est <- glm(qsmk ~ sex + race + age + I(age^2) + as.factor(education)</pre>
                + smokeintensity + I(smokeintensity^2) + smokeyrs
                + I(smokeyrs^2) + as.factor(exercise) + as.factor(active)
                + wt71 + I(wt71^2), data = nhefs, weight = wc,
                family = binomial())
## 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
                                    1.31e+00 -1.83 0.06743 .
## (Intercept)
                                               -3.42 0.00062 ***
## sex
                        -5.14e-01 1.50e-01
## race
                        -8.61e-01 2.06e-01
                                               -4.18 2.9e-05 ***
                         1.15e-01 4.95e-02
                                               2.33 0.01992 *
## age
## 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
                                                2.13 0.03314 *
## as.factor(education)5 4.71e-01
                                    2.21e-01
## smokeintensity
                        -7.83e-02
                                    1.49e-02
                                             -5.27 1.4e-07 ***
## I(smokeintensity^2)
                         1.07e-03
                                    2.78e-04
                                                3.85 0.00012 ***
## smokeyrs
                        -7.11e-02
                                   2.71e-02
                                               -2.63 0.00862 **
## 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
                                               -0.31 0.75530
                                    2.46e-02
## 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$pqsmk
##
                                                                .05
                                                                         .10
                                  Info
                                            Mean
                                                      Gmd
          n missing distinct
##
                         1629
                                     1
                                          0.2622
                                                   0.1302
                                                            0.1015
                                                                     0.1261
       1629
                   0
##
                 .50
                          .75
                                    .90
                                             .95
        .25
##
     0.1780
              0.2426
                       0.3251
                                0.4221
                                          0.4965
##
## lowest : 0.05145 0.05157 0.05438 0.05583 0.05931
## highest: 0.67208 0.68643 0.71391 0.73330 0.78914
summary(nhefs$pqsmk)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
  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)),]</pre>
with(nhefs.c, sum(wc*wt82_71*(qsmk-pqsmk)) / sum(wc*qsmk*(qsmk - pqsmk)))
## [1] 3.446
```

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

## [1,] 2.859 0.03004

```
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]
```

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

```
library("readxl")
nhefs <- read_excel(here("data", "NHEFS.xls"))</pre>
nhefs$cens <- ifelse(is.na(nhefs$wt82), 1, 0)</pre>
# regression on covariates, allowing for some effect modification
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) + I(qsmk*smokeintensity), data=nhefs)
summary(fit)
##
## 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:
##
       Min
                 1Q Median
                                   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
```

```
## qsmk
                                       2.5595941 0.8091486
                                                              3.163 0.001590 **
## 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 *
## I(age * age)
                                      -0.0061010 0.0017261 -3.534 0.000421 ***
## as.factor(education)2
                                      0.7904440 0.6070005
                                                            1.302 0.193038
## 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
                                     -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
                                      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
## wt71
                                      0.0455018 0.0833709
                                                              0.546 0.585299
## 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
     (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
  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)'] \leftarrow c(5, 40)
# (step 3) check the contrast matrix
##
                                                        (Intercept) qsmk sex race
## Effect of Quitting Smoking at Smokeintensity of 5
## 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
## Effect of Quitting Smoking at Smokeintensity of 40
                                                        as.factor(education)4
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
##
                                                       as.factor(education)5
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
##
                                                       smokeintensity
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                       I(smokeintensity * smokeintensity)
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                                         0
                                                       smokeyrs
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                       I(smokeyrs * smokeyrs)
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                       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
## Effect of Quitting Smoking at Smokeintensity of 40
                                                       as.factor(active)2 wt71
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                                             0
                                                       I(wt71 * wt71)
## Effect of Quitting Smoking at Smokeintensity of 5
## Effect of Quitting Smoking at Smokeintensity of 40
                                                       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)</pre>
  summary(estimates1)
##
##
     Simultaneous Tests for General Linear Hypotheses
##
## Fit: 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)
##
## Linear Hypotheses:
##
                                                            Estimate Std. Error
## Effect of Quitting Smoking at Smokeintensity of 5 == 0
                                                              2.7929
                                                                         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 \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), data = nhefs)
##
##
## Deviance Residuals:
##
      Min
               10 Median
                               30
                                       Max
                  -0.318
## -42.332
          -4.216
                            3.807
                                    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
                                   0.5864117 0.5816949 1.008 0.313560
                                   ## 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
## smokeintensity
                                   0.0651533 0.0503115
                                                       1.295 0.195514
## I(smokeintensity * smokeintensity) -0.0010468 0.0009373 -1.117 0.264261
## smokeyrs
                                   0.1333931 0.0917319
                                                       1.454 0.146104
## 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
## 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
```

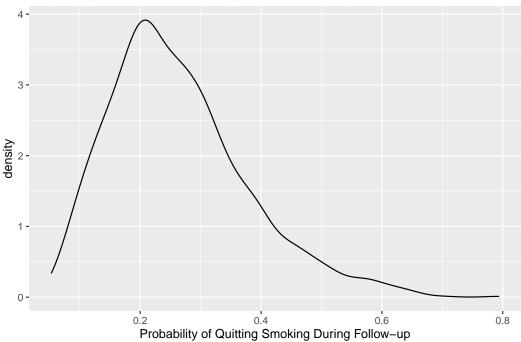
#### Program 15.2

- Estimating and plotting the propensity score
- Data from NHEFS

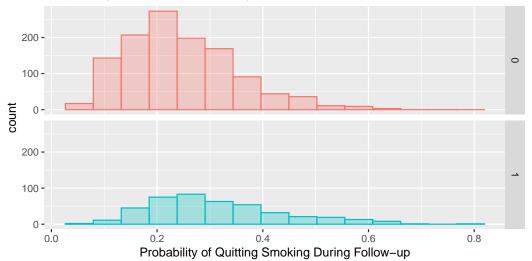
```
##
## Call:
  glm(formula = qsmk ~ sex + race + age + I(age * age) + as.factor(education) +
      smokeintensity + I(smokeintensity * smokeintensity) + smokeyrs +
##
##
      I(smokeyrs * smokeyrs) + as.factor(exercise) + as.factor(active) +
      wt71 + I(wt71 * wt71), family = binomial(), data = nhefs)
##
##
## Deviance Residuals:
                   Median
                                3Q
##
               1Q
                                       Max
  -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 *
                                  -0.0006052 0.0005074 -1.193 0.232973
## I(age * age)
## 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
                                  -0.0733708 0.0269958
                                                       -2.718 0.006571 **
## I(smokeyrs * smokeyrs)
                                   0.0008384 0.0004435
                                                        1.891 0.058669 .
## as.factor(exercise)1
                                   0.2914117 0.1735543
                                                        1.679 0.093136 .
                                   0.3550517 0.1799293
## as.factor(exercise)2
                                                       1.973 0.048463 *
```

```
## as.factor(active)1
                                     0.0108754 0.1298320
                                                          0.084 0.933243
## as.factor(active)2
                                     ## wt71
## I(wt71 * wt71)
                                                          0.895 0.370957
                                     0.0001209 0.0001352
## ---
## 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$qsmk==0])
     Min. 1st Qu. Median
                            Mean 3rd Qu.
                                            Max.
## 0.05298 0.16949 0.22747 0.24504 0.30441 0.65788
summary(nhefs$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



#### 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())
```

## Program 15.3

- Stratification on the propensity score
- Data from NHEFS

```
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
## X1 1 151 0.1 0.02 0.11 0.1 0.02 0.05 0.13 0.08 -0.55 -0.53 0
## -----
## : 2
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## -----
## : 3
## : 0
  vars n mean sd median trimmed mad min max range skew kurtosis se
## -----
## : 4
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
    1 129 0.21 0.01 0.21 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
## -----
## : 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
##
```

```
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
## -----
## : 10
## : 0
## vars n mean sd median trimmed mad min max range skew kurtosis se
## -----
## : 1
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
## 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
## X1 1 27 0.15 0.01 0.15 0.15 0.01 0.13 0.17 0.03 -0.03 -1.34 0
## -----
## : 3
## : 1
## vars n mean sd median trimmed mad min max range skew kurtosis se
## X1 1 29 0.18 0.01 0.18 0.01 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
## X1 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
   1 45 0.26 0.01 0.26 0.26 0.01 0.25 0.27 0.03 0.2 -1.12 0
## -----
## : 7
## : 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
## -----
## : 1
   vars n mean sd median trimmed mad min max range skew kurtosis se
## -----
## : 10
## : 1
  vars n mean sd median trimmed mad min max range skew kurtosis
## X1 1 77 0.52 0.08 0.51
                            0.51 0.08 0.42 0.79 0.38 0.88 0.81 0.01
# 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 between group 0 and group 1 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
## alternative hypothesis: true difference in means between group 0 and group 1 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 between group 0 and group 1 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 between group 0 and group 1 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 between group 0 and group 1 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 between group 0 and group 1 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 between group 0 and group 1 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 between group 0 and group 1 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 between group 0 and group 1 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 between group 0 and group 1 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)</pre>
summary(fit.psdec)
##
## Call:
## glm(formula = wt82_71 ~ qsmk + as.factor(ps.dec), data = nhefs)
##
## Deviance Residuals:
##
      Min
                     Median
                                           Max
                 10
                                   3Q
##
  -43.543
            -3.932
                     -0.085
                                4.233
                                        46.773
##
## Coefficients:
                       Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                    0.6089
                                            6.159 9.29e-10 ***
                         3.7505
                         3.5005
                                    0.4571
                                            7.659 3.28e-14 ***
## qsmk
## 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

## Program 15.4

- 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)
                + 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
11 \leftarrow mean - qnorm(0.975)*se
ul \leftarrow mean + qnorm(0.975)*se
bootstrap <- data.frame(cbind(c("Observed", "No Treatment", "Treatment",
                                "Treatment - No Treatment"), mean, se, ll, ul))
bootstrap
##
                           V1
                                                                                 11
                                           mean
                                                               se
                     Observed 2.63384609228479 0.312266616839415 2.02181476970537
## 1
                 No Treatment 1.71983636149843 0.230246012511331 1.26856246939226
## 2
                    Treatment 5.35072300362993 0.806949126721575 3.7691317778996
## 3
## 4 Treatment - No Treatment 3.6308866421315 0.683375663777217 2.29149495321701
## 1 3.24587741486422
## 2 2.17111025360459
## 3 6.93231422936027
       4.970278331046
# regression on the propensity score (linear term)
model6 <- glm(wt82_71 ~ qsmk + ps, data = nhefs) # p.qsmk</pre>
summary(model6)
##
## Call:
## glm(formula = wt82_71 ~ qsmk + ps, data = nhefs)
##
## Deviance Residuals:
##
       Min
                 1Q
                      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 ***
                                     7.765 1.47e-14 ***
## qsmk
                 3.5506
                            0.4573
## 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
  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
mean0
## [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 ## 2.673886 , 4.438444

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$price82)
##
      Min. 1st Qu. Median
                              Mean 3rd Qu.
                                               Max.
                                                       NA's
##
             1.740
                    1.815
                             1.806
                                      1.868
                                              2.103
                                                          92
# for simplicity, ignore subjects with missing outcome or missing instrument
nhefs.iv <- nhefs[which(!is.na(nhefs$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
     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 between group 0 and group 1 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|)
##
## (Intercept) 2.068164
                           5.085098 0.40671 0.68428
                2.396270 19.840037 0.12078 0.90388
## qsmk
##
## 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
```

- 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
- $\bullet\,$  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")</pre>
```

```
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
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Correlation structure = independence
## Estimated Scale Parameters:
##
##
               Estimate Std.err
                      1 0.7607
## (Intercept)
## 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)
##
                    beta
                               lcl
                                       ucl
## (Intercept) 3.555e+00 3.23152 3.87917
               2.748e-07 -0.04456 0.04456
## Hpsi
```

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

```
summary(tsls(wt82_71 \sim qsmk, \sim 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|)
                  -7.89
                             42.25 -0.187
## (Intercept)
                                               0.852
                  41.28
                             164.95
                                      0.250
                                               0.802
## qsmk
##
## Residual standard error: 18.6055 on 1474 degrees of freedom
summary(tsls(wt82_71 \sim qsmk, \sim 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
                                       18.1
                                               75.3
##
               Estimate Std. Error t value Pr(>|t|)
##
                             48.08
## (Intercept)
                  13.16
                                    0.274
                                               0.784
                 -40.91
                            187.74 -0.218
## qsmk
                                               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 -8.31
                     -3.44
                              0.00
                                      7.27
                                              60.53
##
               Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                  8.086
                             7.288
                                      1.110
                                               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 \sim qsmk, \sim 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|)
                  5.963
                             6.067
                                     0.983
## (Intercept)
                                              0.326
                -12.811
                            23.667 -0.541
                                              0.588
## qsmk
##
## 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

```
model2 <- tsls(wt82_71 ~ qsmk + sex + race + age + smokeintensity + smokeyrs +
                     as.factor(exercise) + as.factor(active) + wt71,
            ~ highprice + sex + race + age + smokeintensity + smokeyrs + as.factor(exercise) +
              as.factor(active) + wt71, data = nhefs.iv)
summary(model2)
##
##
   2SLS Estimates
##
## Model Formula: wt82_71 ~ qsmk + sex + race + age + smokeintensity + smokeyrs +
      as.factor(exercise) + as.factor(active) + wt71
##
##
## 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|)
                       17.280330
                                             7.399 2.3e-13 ***
## (Intercept)
                                   2.335402
                       -1.042295 29.987369 -0.035
                                                      0.9723
## qsmk
## sex
                       -1.644393
                                   2.630831 -0.625
                                                      0.5320
                       -0.183255
                                   4.650386 -0.039
                                                      0.9686
## race
## age
                       -0.163640
                                   0.240548 -0.680
                                                      0.4964
## smokeintensity
                                             0.040
                                                      0.9684
                        0.005767
                                   0.145504
## smokeyrs
                        0.025836
                                   0.161421
                                             0.160
                                                      0.8729
## as.factor(exercise)1 0.498748
                                   2.171239
                                             0.230
                                                      0.8184
## as.factor(exercise)2 0.581834
                                             0.267
                                   2.183148
                                                      0.7899
## as.factor(active)1
                                   0.607466 -1.926
                       -1.170145
                                                      0.0543 .
## as.factor(active)2 -0.512284 1.308451 -0.392
                                                      0.6955
## wt71
                       -0.097949
                                   0.036271 -2.701
                                                      0.0070 **
```

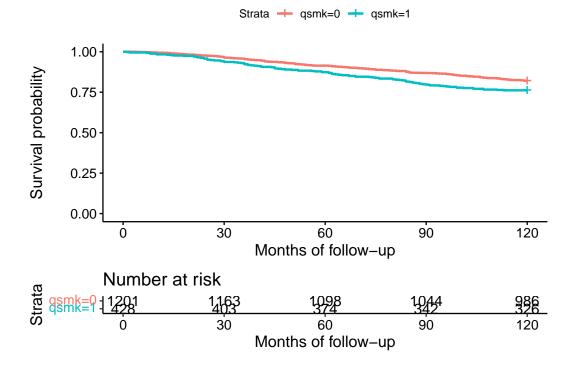
```
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 7.7162 on 1464 degrees of freedom
```

# 17. Causal survival analysis

- Nonparametric estimation of survival curves
- 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.
                                               Max.
      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")
## Loading required package: ggpubr
## Attaching package: 'survminer'
## The following object is masked from 'package:survival':
##
##
       myeloma
```

```
survdiff(Surv(survtime, death) ~ qsmk, data=nhefs)
## Call:
## survdiff(formula = Surv(survtime, death) ~ qsmk, data = nhefs)
##
             N Observed Expected (O-E)^2/E (O-E)^2/V
                           237.5
## qsmk=0 1201
                    216
                                       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)
```



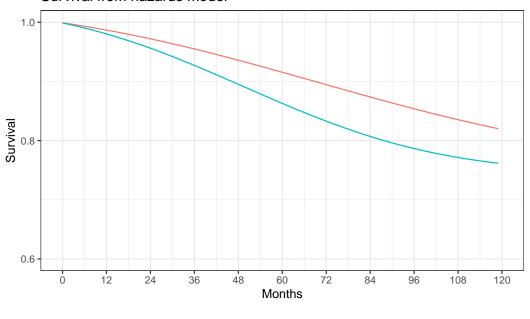
- Parametric estimation of survival curves via hazards model
- Data from NHEFS

```
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
                 10
                     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 ***
## qsmk
                    -3.355e-01 3.970e-01 -0.845
                                                    0.3981
## 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 *
## timesq
                    1.256e-04 6.686e-05 1.878
                                                    0.0604 .
## ---
## 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"))
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")</pre>
```

#### Survival from hazards model



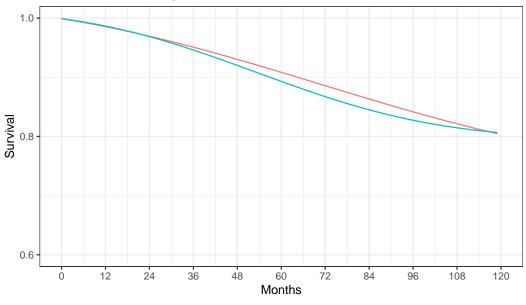
### A: — 0 — 1

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

```
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,</pre>
                     (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) +</pre>
                   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
                 10
                      Median
                                   30
                                           Max
                     0.0595
## -7.1859
             0.0528
                               0.0640
                                        0.1452
##
## Coefficients:
                     Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                     6.897e+00 2.208e-01 31.242
                                                     <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
## time
                   -1.889e-02 8.053e-03 -2.345
                                                     0.0190 *
                    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 \leftarrow data.frame(cbind(seq(0, 119), 0, (seq(0, 119))^2))
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



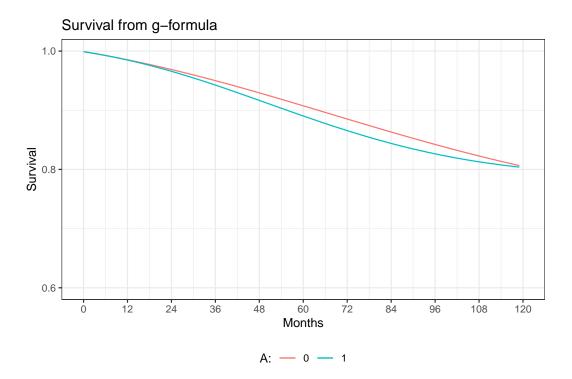
#### A: — 0 — 1

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

```
# fit of hazards model with covariates
gf.model <- glm(event==0 ~ qsmk + I(qsmk*time) + I(qsmk*timesq)</pre>
                + 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),
                data=nhefs.surv, family=binomial())
summary(gf.model)
##
## 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
                                    3Q
                 1Q
                      Median
                                            Max
## -4.3160
             0.0244
                      0.0395
                               0.0640
                                         0.3303
```

```
##
## Coefficients:
##
                                       Estimate Std. Error z value Pr(>|z|)
                                      9.272e+00 1.379e+00 6.724 1.76e-11 ***
## (Intercept)
                                      5.959e-02 4.154e-01 0.143 0.885924
## qsmk
                                     -1.485e-02 1.506e-02 -0.987 0.323824
## I(qsmk * time)
                                      1.702e-04 1.245e-04 1.367 0.171643
## I(qsmk * timesq)
                                     -2.270e-02 8.437e-03 -2.690 0.007142 **
## time
                                      1.174e-04 6.709e-05 1.751 0.080020 .
## timesq
                                      4.368e-01 1.409e-01 3.101 0.001930 **
## sex
                                     -5.240e-02 1.734e-01 -0.302 0.762572
## race
                                     -8.750e-02 5.907e-02 -1.481 0.138536
## age
                                      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
                                      3.750e-01 2.386e-01 1.571 0.116115
## as.factor(education)5
                                     -1.626e-03 1.430e-02 -0.114 0.909431
## smokeintensity
## 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
                                     -1.677e-02 3.065e-02 -0.547 0.584153
## smokeyrs
## 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
## wt.71
                                     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$time <- rep(seq(0, 119), nrow(nhefs))</pre>
gf.qsmk0$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 <- 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^{smk}=1 \& psi \le log(120/d^{survtime})),
  } 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</pre>
  testlow <- objfunc</pre>
  countlow <- 0
  while (testlow < 3.84 & countlow < 100){
    psilow <- psilow - increm
    testlow <- sumeef(psilow)</pre>
    countlow <- countlow + 1</pre>
  }
  # Upper bound of 95% CI
  psihigh <- psi1
  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
    } 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</pre>
        fleft <- fmiddle
      middle <- (left + right) / 2
      fmiddle <- for_conf(middle)</pre>
      diff <- right - left
      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 4.1.2 (2021-11-01)
## os
           macOS Big Sur 10.16
## system x86_64, darwin17.0
## ui
           X11
## language (EN)
## collate en_GB.UTF-8
## ctype
           en_GB.UTF-8
           Europe/London
## tz
##
   date
           2022-02-20
           2.17.1.1 @ /Applications/RStudio.app/Contents/MacOS/quarto/bin/ (via rmarkdown)
##
   pandoc
##
##
   package
              * version date (UTC) lib source
  bookdown
               0.24
                      2021-09-02 [1] CRAN (R 4.1.0)
## cli
               3.2.0
                      2022-02-14 [1] CRAN (R 4.1.2)
               0.6.29 2021-12-01 [1] CRAN (R 4.1.0)
## digest
## evaluate
               0.15
                      2022-02-18 [1] CRAN (R 4.1.2)
## fastmap
               1.1.0 2021-01-25 [1] CRAN (R 4.1.0)
## htmltools
               0.5.2
                      2021-08-25 [1] CRAN (R 4.1.0)
               1.37
## knitr
                      2021-12-16 [1] CRAN (R 4.1.0)
               2.0.2
                      2022-01-26 [1] CRAN (R 4.1.2)
## magrittr
                      2022-02-03 [1] CRAN (R 4.1.2)
## rlang
               1.0.1
               2.11
                      2021-09-14 [1] CRAN (R 4.1.0)
## rmarkdown
                      2020-11-12 [1] CRAN (R 4.1.0)
## rstudioapi
               0.13
## sessioninfo 1.2.2
                      2021-12-06 [1] CRAN (R 4.1.0)
## stringi
               1.7.6
                      2021-11-29 [1] CRAN (R 4.1.0)
## stringr
               1.4.0
                      2019-02-10 [1] CRAN (R 4.1.0)
               0.29
                      2021-12-14 [1] CRAN (R 4.1.0)
##
  xfun
##
  yaml
               2.3.4
                      2022-02-17 [1] CRAN (R 4.1.2)
##
   [1] /Library/Frameworks/R.framework/Versions/4.1/Resources/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
1 190
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
```

Y

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

-> A = 0

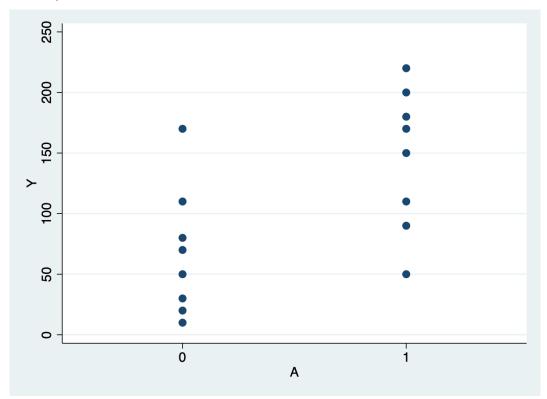
14. 0 50 15. 0 10 16. 0 20 17. end

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

-----

-> A = 1

Variable	Obs	Mean	Std. dev.	Min	Max
Y	   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
         Α
                 Y
 1. 1 110
 2. 1 80
 3. 1 50
 4. 1 40
 5. 2 170
 6. 2 30
 7. 2 70
 8. 2 50
 9. 3 110
10. 3 50
11. 3 180
12. 3 130
13. 4 200
14. 4 150
15. 4 220
16. 4 210
17. end
-> A = 1
  Variable | Obs Mean Std. dev. Min
                        70 31.62278
        ΥI
               4
                                         40
                                                 110
-> A = 2
  Variable | Obs
                       Mean Std. dev. Min
                                                 Max
______
               4
        Υ |
                     80 62.18253 30
                                                 170
```

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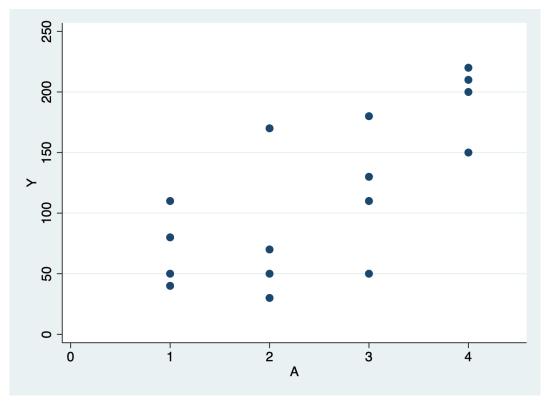
-> A = 3

Variable	1	0bs	Mean	Std. dev.	Min	Max
Υ	+ 	4	117.5	 53.77422	50	180

\_\_\_\_\_

-> A = 4

Variable	1	0bs	Mean	Std.	dev.	Min	Max
Y	+ 	4	195	31.09	9126	 150	220



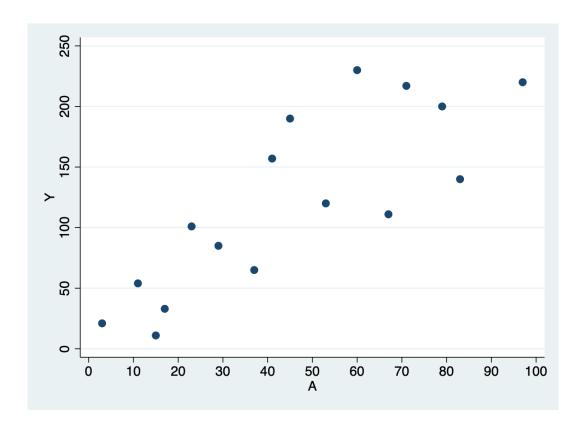
```
clear
**Figure 11.3**
input A Y
3
   21
11 54
17 33
23 101
29 85
37 65
41 157
53 120
67 111
79 200
83 140
97 220
```

```
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
```

```
Α
                       Y
 1. 3
       21
 2. 11
            54
 3. 17
            33
 4. 23
            101
 5. 29
            85
 6. 37
            65
 7. 41
            157
 8. 53
            120
 9.67
            111
10. 79
            200
11. 83
            140
12. 97
            220
13. 60
            230
14. 71
            217
15. 15
16. 45 190
17. end
```



- 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]

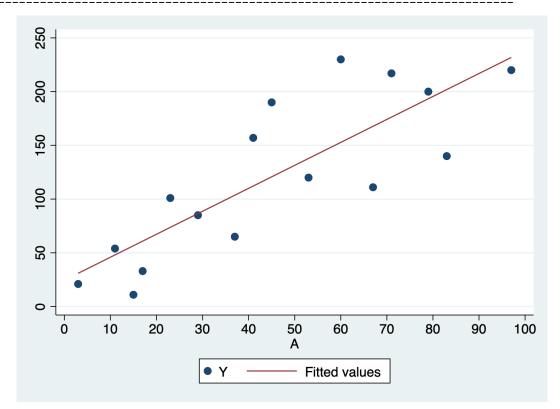
*Plot the data with the regression line: Fig 11.4*
scatter Y A, ylab(0(50)250) xlab(0(10)100) xscale(range(0 100)) || lfit Y A
qui gr export figs/stata-fig-11-4.png, replace
```

\_cons | 24.55 21.33 1.15 0.269 -21.20 70.29

(1) 90\*A + cons = 0

di 67.50 + 78.75

Coefficient		• • •	[95% conf.	interval]
216.89			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]*
```

Y | Coefficient 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

- 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)

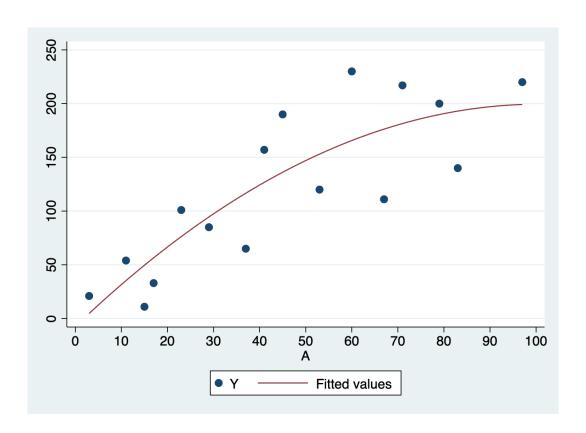
*Output the estimated mean Y value when A = 90*
lincom _b[_cons] + 90*_b[A] + 90*90*_b[Asq]

*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
```

			•	Coefficient				2 - 7	interval]
A   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	A Asq	 	A   Asq	4.11 -0.02	1.53 0.02	2.68 -1.33	0.019 0.206	0.80 -0.05	0.01

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

Coefficient		 [95% conf.	- · · · · -
 197.1269		142.7687	



# 12. IP Weighting and Marginal Structural Models: Stata

### Program 12.1

• 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)
}
 2. tabdisp qsmk, cell(`var') format(%3.1f)
 3. }
_____
quit smoking between |
baseline and 1982 | Age, years
----+----
No smoking cessation |
                       42.8
  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
  Smoking cessation |
_____
quit smoking between |
baseline and 1982 | University, %
-----
No smoking cessation |
                          9.9
                     15.4
  Smoking cessation |
```

quit smoking between baseline and 1982	   Years smoking
No smoking cessation Smoking cessation	70.3   72.4
quit smoking between baseline and 1982	
No smoking cessation	
Smoking cessation	18.6 
quit smoking between baseline and 1982	
No smoking cessation Smoking cessation	
quit smoking between baseline and 1982	
No smoking cessation Smoking cessation	
quit smoking between baseline and 1982	
No smoking cessation Smoking cessation	

- 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
                                                  Number of obs = 1,566
Logistic regression
                                                  LR chi2(18) = 109.16
                                                  Prob > chi2 = 0.0000
Log likelihood = -838.44842
                                                  Pseudo R2 = 0.0611
       qsmk | Coefficient 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
smokeintens~y | -.0772704
                          .0152499
                                     -5.07 0.000
                                                    -.1071596
                                                               -.0473812
          c. |
smokeintens~y#|
         c. |
```

smokeintens~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
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	0bs	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)

_	   Coefficient				[95% conf.	interval]
qsmk	3.440535 1.779978	.5258294	6.54	0.000	2.409131 1.338892	4.47194 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
                                                   Pseudo R2
                                                               = 0.0611
Log likelihood = -838.44842
        qsmk | Coefficient 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
I						
c.age#c.age	0008246	.0005361	-1.54	0.124	0018753	.0002262
, , ,						
education	455000				0.4.00=4.4	
1	4759606	.2262238	-2.10	0.035	9193511	0325701
2		.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
smokeintens~y	0772704	.0152499	-5.07	0.000	1071596	0473812
smokeIntens~y	0772704	.0152499	-5.07	0.000	1071596	04/3012
c. I						
smokeintens~y#						
c.						
smokeintens~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
ا						
smokeyrs	0735966	.0277775	-2.65	0.008	1280395	0191538
I						
c.smokeyrs#						
c.smokeyrs	.0008441	.0004632	1.82	0.068	0000637	.0017519
<u> </u>						
exercise						
0		.1872401	-2.11	0.035	7626878	0287201
1	0408635	.1382674	-0.30	0.768	3118627	.2301357
active	450504	0.4.000.4			5001015	0445505
0	176784	.2149721	-0.82	0.411	5981215	. 2445535
1	1448395	.2111472	-0.69	0.493	5586806	.2690015
ı wt71 ∣	0152357	.0263161	-0.58	0.563	0668144	.036343
W C / T	.0102307	.0203101	0.50	0.503	.000144	.030343
c.wt71#c.wt71	.0001352	.0001632	0.83	0.407	0001846	.000455
	1001002		3.00	0.101		. 300 100
_cons	-1.19407	1.398493	-0.85	0.393	-3.935066	1.546925
<u>-</u>						

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

-	Coefficient			interval]
			-1.173114	946529

(1,566 missing values generated)

(403 real changes made)

(1,163 real changes made)

Variable	l	Obs 1	Mean Std.	dev.	Min Max
	+				
sw_a	1,	566 .9988	3444 .2882	2233 .3312	2489 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)

Robust \_\_\_\_\_ qsmk | 3.440535 .5258294 6.54 0.000 2.409131 4.47194 \_cons | 1.779978 .2248742 7.92 0.000 1.338892 2.221065

| quit smoking between | baseline and 1982

age	1	No	smokin	Smoking	С	I	Total
	+-					+-	
25	1		24		3	I	27
26	1		14		5	1	19
27	1		18		2	1	20
28	1		20		5	1	25
29			15		4	1	19
30	1		14		5	1	19
31	1		11		5	1	16
32	1		14		7	1	21
33	1		12		3	1	15
34			22		5	1	27
35			16		5	1	21

36	13	3	16
37	14	1	l 15
38	6	2	8
39	19	4	23
40	10	4	l 14
41	13	3	l 16
42	16	3	l 19
43	14	3	17
44	9	4	13
45	12	5	17
46	19	4	23
47	19	4	23
48	19	4	23
49	11	3	14
50	18	4	22
51	9	3	1 12
52	11	3	14
53	11	4	15
54	17	9	1 26
55	9	4	13
56	8	7	15
57	9	2	11
58	8	4	12
59	5	4	l 9
60	5	4	1 9
61	5	2	7
62	6	5	11
63	3	3	1 6
64	7	1	8
65	3	2	J 5
66	4	0	l 4
67	2	0	1 2
69	6	2	8
70	2	1	3
71	0	1	1
72	2	2	4
74	0	1	1
Total	524	164	688
	quit smoki	ng between	
	baseline	and 1982	
	1 37 3 .	C1	1 70-4-3

age | No smokin Smoking c | Total 25 | 3 1 | 26 | 3 0 | 3 3 1 4 28 | 1 | 4 29 | 1 0 | 30 I 0 |

		•	
31	3	0	] 3
32	1 8	0	8
33	1 2	0	1 2
34	1 2	1	3
35	3	0	1 3
36	1 5	0	1 5
37	3	1	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	1 6
44	3	0	] 3
45	1	3	1 4
46	J 5	0	J 5
47	3	0	] 3
48	4	0	1 4
49	1	1	1 2
50	2	0	1 2
51	4	0	4
52	1	0	1
53	1 2	0	1 2
54	2	0	1 2
55	3	0	] 3
56	2	1	] 3
57	2	1	] 3
61	1	1	1 2
67	1	0	1
68	1	0	1
69		0	
70	1 0	1	1
	· ·+		+
Total	l 97	19	116

- Estimating the parameters of a marginal structural mean model with a continuous treatment Data from NHEFS
- Section 12.4

```
"" weights and calculate the mean expected smoking inte
"" which is 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)
                                                               1,162
     Source |
                            df
                                    MS
                                           Number of obs =
  ----- F(18, 1143)
                                                                5.39
                            18 553.164252 Prob > F
     Model | 9956.95654
                                                                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
smkintensi~71 | Coefficient Std. err.
                                    t P>|t|
                                                  [95% conf. interval]
        sex | 1.087021 .7425694 1.46 0.144
                                                   -.3699308
                                                               2.543973
        race | .2319789 .8434739 0.28 0.783 -1.422952
                                                               1.88691
        age | -.8099902 .2555388 -3.17 0.002 -1.311368 -.3086124
                .0066545 .0026849 2.48 0.013
 c.age#c.age |
                                                   .0013865
                                                               .0119224
   education |
         1 | 1.508097 1.184063 1.27
                                           0.203
                                                   -.8150843
                                                               3.831278
               2.02692 1.133772
                                     1.79 0.074
                                                   -.1975876
                                                               4.251428
         3 | 2.240314 1.022556
                                     2.19 0.029
                                                   .2340167
                                                               4.246611
```

1.75 0.081

-.3103458

5.36788

4 | 2.528767 1.44702

   smokeintens~y	3589684	. 2246653	-1.60	0.110	799771	.0818342
c. I						
smokeintens~y#  c.						
smokeintens~y		.0085753	0.23	0.819	0148668	.0187832
smokeyrs	.3857088	.1416765	2.72	0.007	.1077336	.6636841
c.smokeyrs#						
c.smokeyrs   	0054871	.0023837	-2.30	0.022	0101641	0008101
exercise						
	1.996904					
1	.988812	.6929239	1.43	0.154	3707334	2.348357
active						
	.8451341	1 098573	0.77	0 442	-1 310312	3 000581
	.800114					
i						
wt71	0656882	. 136955	-0.48	0.632	3343996	.2030232
c.wt71#c.wt71	.0005711	.000877	0.65	0.515	0011496	.0022918
_cons	16.86761	7.109189	2.37	0.018	2.91909	30.81614
Variable	Obs	Mean	Std. d	ev.	Min I	Max
sw_a	1,162	.9968057	.32229	37 .193	38336 5.1023	339
(sum of wgt is	1,158.2881828	36955)				
Linear regressi	on			Number o	of obs =	1 162
Linear regressi	.01					12.75
				-		0.0000
				R-square	ed =	0.0233
				Root MSE	=	7.7864
		(Std.	err. adju	sted for	1,162 cluster	rs in seqn)
I		Robust				
wt82_71	Coefficient				[95% conf	
smkintensi~71						
c.						
smkintensi~71#						

```
c. |
smkintensi~71 | .0026949 .0024203 1.11 0.266 -.0020537 .0074436
   _cons | 2.004525 .295502 6.78 0.000 1.424747 2.584302
(1) _cons = 0
______
  ______
    (1) | 2.004525 .295502 6.78 0.000 1.424747
 ._____
(1) 20*smkintensity82_71 + 400*c.smkintensity82_71#c.smkintensity82_71 +
   _{cons} = 0
  wt82_71 | Coefficient Std. err. t P>|t| [95% conf. interval]
_______
    (1) | .9027234 1.310533
                   0.69 0.491
                           -1.668554
                                 3.474001
```

- Estimating the parameters of a marginal structural logistic model
- Data from NHEFS
- Section 12.4

```
use ./data/nhefs, clear
/*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
```

```
/*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 Pseudo R2 = 0.0611

Log likelihood = -838.44842

qsmk	Coefficient	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
smokeintens~y	0772704	.0152499	-5.07	0.000	1071596	0473812
с.						
smokeintens~y#						
с.						
smokeintens~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
smokeyrs	0735966	.0277775	-2.65	0.008	1280395	0191538
c.smokeyrs#	l					
c.smokeyrs	.0008441	.0004632	1.82	0.068	0000637	.0017519

exercise	1						
0	1	395704	.1872401	-2.11	0.035	7626878	0287201
1	1	0408635	.1382674	-0.30	0.768	3118627	.2301357
	1						
active	1						
0	1	176784	.2149721	-0.82	0.411	5981215	.2445535
1	1	1448395	.2111472	-0.69	0.493	5586806	.2690015
	1						
wt71	1	0152357	.0263161	-0.58	0.563	0668144	.036343
	1						
c.wt71#c.wt71		.0001352	.0001632	0.83	0.407	0001846	.000455
_cons	1	-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 =.

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

qsmk | Coefficient Std. err. z P>|z| [95% conf. interval]

\_cons | -1.059822 .0578034 -18.33 0.000 -1.173114 -.946529

(1,566 missing values generated)

(403 real changes made)

(1,163 real changes made)

Variable	Obs	Mean	Std. dev.	Min	Max
sw_a	1,566	.9988444	.2882233	.3312489	4.297662

Logistic regression Number of obs = 1,566

Wald chi2(1) = 0.04

Prob > chi2 = 0.8482

Log pseudolikelihood = -749.11596 Pseudo R2 = 0.0000

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

Note: \_cons estimates baseline odds.

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

```
use ./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)
```

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

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 Pseudo R2 = 0.0584

Log likelihood = -883.34876

qsmk	Coefficient	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
١						
c.age#c.age	0006052	.0005074	-1.19	0.233	0015998	.0003893
I						
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 I	4221892	.2717235	-1.55	0.120	9547574	.110379
I						
smokeintens~y	0651561	.0147589	-4.41	0.000	0940831	0362292
<u> </u>						
c.						
smokeintens~y#						
c.						
smokeintens~y	.0008461	.0002758	3.07	0.002	.0003054	.0013867
	0722700	0000050	0.70	0 007	1000010	00046
smokeyrs	0733708	.0269958	-2.72	0.007	1262816	02046
c.smokeyrs#						
c.smokeyrs	.0008384	.0004435	1.89	0.059	0000307	.0017076
C.SMOKEYIS	.0000304	.0004455	1.03	0.039	.0000307	.0017070
exercise						
0 1		. 1799293	-1.97	0.048	7077067	0023967
1 1		.1351256	-0.47	0.638	3284812	.2012013
 I						
active						
0	0683123	.2087269	-0.33	0.743	4774095	.3407849
1	057437	. 2039967	-0.28	0.778	4572632	.3423892
ĺ						
wt71	0128478	.0222829	-0.58	0.564	0565214	.0308258
I						
c.wt71#c.wt71	.0001209	.0001352	0.89	0.371	000144	.0003859
I						

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 Pseudo R2 = 0.0050

Log likelihood = -933.49126

qsmk | Coefficient 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

<del>-</del>

(1,629 missing values generated)

(428 real changes made)

(1,201 real changes made)

Variable	Obs	Mean	Std. dev.	Min	Max
sw_a	1,629	.9991318	.2636164	.2901148	3.683352

(sum of wgt is 1,562.01032829285)

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)

- 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 |
                             Total
-----+----+-----
      0 | 1,163
                     403 |
                             1,566
           96.84 94.16 |
       96.13
     1 |
             38
                      25 I
       | 3.16 5.84 | 3.87
   Total | 1,201
                    428 |
                             1,629
       | 100.00 100.00 | 100.00
\rightarrow cens = 0
  Variable | Obs Mean Std. dev. Min
     wt71 | 1,566 70.83092
                               15.3149
                                         39.58
                                                151.73
-> cens = 1
  Variable | Obs Mean Std. dev. Min
     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 Pseudo R2 = 0.0584

Log likelihood = -883.34876

qsmk	Coefficient	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	-1.19	0.233	0015998	.0003893
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
smokeintens~y	0651561	.0147589	-4.41	0.000	0940831	0362292
c. l						
smokeintens~y#						
c.						
smokeintens~y	.0008461	.0002758	3.07	0.002	.0003054	.0013867
smokeyrs	0733708	.0269958	-2.72	0.007	1262816	02046
c.smokeyrs#						
c.smokeyrs	.0008384	.0004435	1.89	0.059	0000307	.0017076
exercise						
0	3550517	.1799293	-1.97	0.048	7077067	0023967
1	06364	.1351256	-0.47	0.638	3284812	.2012013
active	 					
0	0683123	.2087269	-0.33	0.743	4774095	.3407849
1 1	057437	.2039967	-0.28			.3423892
<u> </u>	.007407	. 2003301	0.20	0.770	.4072002	.0420032
wt71	0128478	.0222829	-0.58	0.564	0565214	.0308258
c.wt71#c.wt71	.0001209	.0001352	0.89	0.371	000144	.0003859
_cons	-1.185875	1.263142	-0.94	0.348	-3.661588	1.289838

Iteration 0: log likelihood = -938.14308
Iteration 1: log likelihood = -938.14308

Logistic regression Number of obs = 1,629

LR chi2(0) = 0.00

Prob > chi2 = .

Log likelihood = -938.14308 Pseudo R2 = 0.0000

qsmk | Coefficient Std. err. z P>|z| [95% conf. interval]

\_\_\_\_\_

Iteration 0: log likelihood = -266.67873
Iteration 1: log likelihood = -238.48654

Iteration 1: log likelihood = -238.48654 Iteration 2: log likelihood = -232.82848

Iteration 3: log likelihood = -232.68043

Iteration 4: log likelihood = -232.67999

Iteration 5:  $\log likelihood = -232.67999$ 

Logistic regression Number of obs = 1,629

LR chi2(19) = 68.00

Prob > chi2 = 0.0000

Log likelihood = -232.67999 Pseudo R2 = 0.1275

cens	Coefficient	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	4999559	0395027
	I					
c.age#c.age	.0028837	.0011135	2.59	0.010	.0007012	.0050661
	I					
education	I					
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
	I					
smokeintens~y	.0157119	.0347319	0.45	0.651	0523614	.0837851
_	I					
С.	I					

smokeintens~y#						
c.						
smokeintens~y	0001133	.0006058	-0.19	0.852	0013007	.0010742
1						
smokeyrs	.0785973	.0749576	1.05	0.294	0683169	.2255116
1						
c.smokeyrs#						
c.smokeyrs	0005569	.0010318	-0.54	0.589	0025791	.0014653
1						
exercise						
0	.583989	.3723133	1.57	0.117	1457317	1.31371
1	3874824	.3439133	-1.13	0.260	-1.06154	.2865754
1						
active						
0	7065829	.3964577	-1.78	0.075	-1.483626	.0704599
1	9540614	.3893181	-2.45	0.014	-1.717111	1910119
1						
wt71	0878871	.0400115	-2.20	0.028	1663082	0094659
1						
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

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

Log likelihood = -263.88009 Pseudo R2 = 0.0105

	Coefficient				[95% conf.	_
qsmk	.6411113	. 2639262	2.43	0.015	.1238255 -3.744273	1.158397

(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)

Linear regression Number of obs = 1,566 F(1, 1565) = 44.19 Prob > F = 0.0000 R-squared = 0.0363 Root MSE = 7.8652

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

wt82_71	Coefficient				[95% conf.	_
qsmk	3.496493 1.66199	.5259796	6.65 7.14	0.000	2.464794 1.205164	4.528192

# 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

```
/* 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
```

Iteration 0: log likelihood = -5328.5765

note: 1.qsmk omitted because of collinearity.

note: smokeintensity omitted because of collinearity.

Generalized lin				Residu	r of obs = mal df = parameter =	1,566 1,545 53.5683
Deviance	= 82763.02	862		(1/df)	Deviance =	53.5683
Pearson	= 82763.02	862		(1/df)	Pearson =	53.5683
Variance functi	ion: $V(u) = 1$			[Gauss		
Link function	: g(u) = u			[Ident	city]	
				AIC	=	6.832154
Log likelihood	= -5328.576	456		BIC	=	71397.58
	 	 OIM				
wt82_71	Coefficient	std. err.	z	P> z	[95% conf.	interval]
qsmk			3.16		.973692	
	-1.430272		-3.05	0.002	-2.349412	
race			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
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
smokeintens~y	   .0491365	.0517254	0.95	0.342	0522435	.1505165
c.						
smokeintens~y#						
c.   smokeintens~y		.000938	-1.06	0.291	0028292	.0008479
Smokerneens y	.0003301	.000500	1.00	0.201	.0020232	.0000473
smokeyrs	.1343686 	.0917122	1.47	0.143	045384	.3141212
c.smokeyrs#						
c.smokeyrs	0018664	.0015437	-1.21	0.227	0048921	.0011592
exercise						
0	3539128	.5588587	-0.63	0.527	-1.449256	.7414301
1	0579374	.4316468	-0.13	0.893	9039497	.7880749
active	 					
0	.2613779	.6845577	0.38	0.703	-1.08033	1.603086
1	6861916	.6739131	-1.02	0.309	-2.007037	.6346539

(option mu assumed; predicted mean wt82\_71)

Variable	Obs	Mean	Std. dev.	Min	Max
meanY	1,566	2.6383	3.034683	-10.87582	9.876489
+					
Variable	Obs	Mean	Std. dev.		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

```
"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
/* 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 original
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
                                    Α
                                               Y
             ID
                         L
 1. "Rheia"
                     0 0 0
  2. "Kronos"
                     0 0 1
  3. "Demeter"
                     0 0 0
  4. "Hades"
                     0 0 0
  5. "Hestia"
                     0 1 0
  6. "Poseidon"
                     0 1 0
 7. "Hera"
                     0 1 0
 8. "Zeus"
                     0 1 1
 9. "Artemis"
                     1 0 1
 10. "Apollo"
                     1 0 1
 11. "Leto"
                     1 0 0
 12. "Ares"
                     1 1 1
```

13. "Athena"

1 1 1

- 14. "Hephaestus" 1 1 1
- 15. "Aphrodite" 1 1 1
- 16. "Cyclope" 1 1 1
- 17. "Persephone" 1 1 1
- 18. "Hermes" 1 1 0
- 19. "Hebe" 1 1 0
- 20. "Dionysus" 1 1 0
- 21. end
- (20 observations created)
- (20 observations created)
- (20 real changes made)

Expanded observation			
type	Freq.	Percent	Cum.
	+		
-1	J 20	33.33	33.33
Original observation	1 20	33.33	66.67
Duplicated observation	1 20	33.33	100.00
Total	60	100.00	

(40 real changes made, 40 to missing)

(13 real changes made)

(7 real changes made)

-----

#### -> interv = -1

Variable	Ob	s Mean	Std. de	ev. Min	Max
Α	l 2	 0 .65	.48936	05 0	1

\_\_\_\_\_

#### -> interv = Original

Variable	Obs	Mean	Std. dev.	Min	Max
A	20	0	0	0	0

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

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

Variable	Obs	Mean	Std. de	V.	Min	Max	
·	20	1		0	1	1	
	SS	df			er of obs		20
·					16)		
	.833333333				> F		
	4.16666667			_			
	5			_	_		
	Coefficient		t	 P> t	[95% cor	of. int	terval]
·	1.05e-16		0.00	1.000	7649549		7649549
1.L	.4166667	.389756	1.07	0.301	4095791	1	. 242912
A#L   1 1	-5.83e-17	. 4959325	-0.00	1.000	-1.05133	3 :	1.05133
_cons	. 25	. 2551552	0.98	0.342	2909048	3 .7	7909048
-> interv = -1							
	Obs					Max	
	20					866667	
	riginal						
Variable	Obs			v.	Min	Max	
predY	20	.5	. 20942	7	.25 .66	366667	
	uplicat						
	0bs	Mean	Std. de	v.	Min	Max	
predY		.5	.20942	 7	.25 .66	66667	

#### observe[4,2]

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

use ./data/nhefs-formatted, clear

\*ii.Estimation in original sample\*

- Standardizing the mean outcome to the baseline confounders:
- Data from NHEFS
- Section 13.3

```
*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
```

```
*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 origi:
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)
 Expanded observation |
              type |
                      Freq. Percent
                                            Cum.
                                33.33
                -1 |
                       1,566
                                            33.33
                       1,566
                                 33.33
 Original observation |
                                           66.67
Duplicated observation | 1,566
                                 33.33
                                          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 | Obs Mean Std. dev. Min Max
      qsmk | 1,566 .2573436 .4373099
                                               0
-> interv = Original
   Variable | Obs Mean Std. dev. Min Max
                       0 0
      qsmk | 1,566
                                            0
```

-> interv = Duplicat

Variable	Obs	Mean	Std. dev	<b>J</b> .	Min M	ax
qsmk	1,566	1	(	)	1	1
Source	SS	df	MS		r of obs = 1545) =	
Model	14412.558	20	720 6279	-	> F =	
	82763.0286				ared =	
+-					-squared =	
Total	97175.5866	1,565	62.0930266	=	=	7.319
wt82_71	Coefficient	Std. err	t	P> t	[95% conf.	interval]
qsmk			3.16	0.002		
sex			-3.05		-2.350132	
	.5601096		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	.194977	.7413692	0.26	0.793	-1.259219	1.649173
2		.7012116	1.41	0.160	390006	2.360848
3			1.19	0.236	4921358	1.994715
4	1.686547	.8716593	1.93	0.053	0232138	3.396307
smokeintens~y	.0491365	.0517254	0.95	0.342	052323	.1505959
c.						
smokeintens~y#						
c.						
smokeintens~y	0009907	.000938	-1.06	0.291	0028306	.0008493
smokeyrs	.1343686	.0917122	1.47	0.143	045525	.3142621
c.smokeyrs#						
c.smokeyrs	0018664	.0015437	-1.21	0.227	0048944	.0011616
exercise						
0	3539128	.5588587	-0.63	0.527	-1.450114	.7422889
1	0579374	.4316468	-0.13	0.893	904613	.7887381
i		. 1313100	0.10		.001010	
active						
0		.6845577	0.38	0.703	-1.081382	1.604138
1	6861916	.6739131	-1.02	0.309	-2.008073	.6356894
wt71	.0455018	.0833709	0.55	0.585	1180303	.2090339

   c.wt71#c.wt71     qsmk#	0009653	.0005247	-1.84	0.066	001	19945	.0000639
c.							
smokeintens~y							
Smoking ce	.0466628	.0351448	1.33	0.184	022	22737	.1155993
_cons   	-1.690608	4.388883	-0.39	0.700	-10.	. 2994	6.918188
-> interv = -1							
Variable	0bs	Mean	Std. de	v.	Min	Ma	ax 
predY	1,566	2.6383	3.03468	3 -10.8	37582	9.87648	39
	ginal						
Variable	0bs	Mean	Std. de	v.	Min	Ma	ıx 
predY	1,566	1.756213	2.82627	1 -11.8	33737	6.73349	98
interv = Dup	licat						
Variable	0bs	Mean	Std. de	v.	Min	Ma	ıx 
predY	1,566	5.273587	2.92053	32 -9.09	91126	11.050	06

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)
```

12.

Command: bootstdz

EY\_a0: r(boot\_0)
EY\_a1: r(boot\_1)
difference: r(boot\_diff)

Simulations (10)

• • • • • • • • • •

(loading observe[4,2])

pe[1,3]

r2 r3 r4 c2 1.7562131 5.2735873 3.5173742

Bootstrap results

Number of obs = 1,629

Replications = 10

	Observed coefficient		z	P> z	Normal	
EY_a0   EY_a1   difference	1.756213 5.273587	.2157234 .4999001 .538932	8.14 10.55 6.53	0.000 0.000 0.000	1.333403 4.293801 2.461087	2.179023 6.253374 4.573662

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

## Program 14.1

- 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
```

```
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
                                        Number of obs = 1,629
Optimization : ML
                                        Residual df =
                                                          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 | Coefficient 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
```

/\*Analyses restricted to N=1566\*/

drop if wt82 == .

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
education   1	.3823818	.5601808	0.68	0.495	7155523	1.480316
- '			-0.10	0.495		
		.5749586			-1.185305	1.068491
3	.2176937	.5225008	0.42	0.677	8063891	1.241776
4	.5208288	.6678735	0.78	0.435	7881792	1.829837
smokeintens~y	.0157119	.0347319	0.45	0.651	0523614	.0837851
I						
c. l						
smokeintens~y#						
c.						
smokeintens~y	0001133	.0006058	-0.19	0.852	0013007	.0010742
smokeyrs	.0785973	.0749576	1.05	0.294	068317	.2255116
c.smokeyrs#						
c.smokeyrs		.0010318	-0.54	0.589	0025791	.0014653
١						
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
_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	<u> </u>	0bs	Mean	Std.	dev.	Min	Max
w_cens	   1,	,566 1.	039197	.05	5646 1.0	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
                                                     Number of obs = 1,566
Logistic regression
                                                     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 segn)
                            Robust
       qsmk | Coefficient 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 | -.5000231 .2208583
                              -2.26 0.024
                                             -.9328974 -.0671487
        3 | -.3833788 .195914 -1.96 0.050
                                             -.7673632 .0006056
        4 | -.4047116 .2836068
                                -1.43 0.154
                                             -.9605707
                                                        .1511476
smokeintens~y | -.0783425 .014645 -5.35 0.000
                                             -.1070461
                                                       -.0496389
        c. |
smokeintens~y#|
smokeintens~y | .0010722 .0002651 4.04 0.000 .0005526 .0015917
   smokeyrs | -.0711097 .026398 -2.69 0.007
                                             -.1228488 -.0193705
          c.smokeyrs#|
  c.smokeyrs | .0008153
                       .0004491
                               1.82 0.069
                                             -.000065
                                                        .0016955
   exercise |
        0 | -.3800465 .1889205 -2.01 0.044
                                             -.7503238 -.0097692
        1 | -.0437043
                                -0.32 0.750
                       .1372725
                                             -.3127534
                                                        .2253447
     active |
        0 | -.2134552 .2122025 -1.01
                                      0.314
                                             -.6293645
                                                        .2024541
        1 | -.1793327
                       .207151 -0.87 0.387
                                             -.5853412
                                                        .2266758
       wt71 | -.0076607 .0256319 -0.30 0.765
                                             -.0578983
                                                        .0425769
c.wt71#c.wt71 | .0000866 .0001582
                               0.55 0.584
                                             -.0002236
                                                        .0003967
      Hpsi | -1.90e-06 .0088414 -0.00 1.000
                                              -.0173307
                                                        .0173269
      -4.00316
                                                        1.326426
```

-1.905e-06

```
6. matrix p_mat = r(table)
7. matrix p_mat = p_mat["pvalue","qsmk:Hpsi"]
8. local p = p_mat[1,1]
9. local b = _b[Hpsi]
10. di "coeff", %6.3f `b', "is generated from psi", %4.1f `i'
11. matrix results[`j',1]= `i'
```

```
12.
            matrix results['j',2]= 'b'
13.
            matrix results[`j',3] = abs(`b')
14.
            matrix results['j',4]= 'p'
15. }
coeff 0.027 is generated from psi
coeff 0.025 is generated from psi 2.1
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
                                   2.6
coeff 0.014 is generated from psi
                                   2.7
coeff 0.012 is generated from psi
coeff 0.010 is generated from psi
coeff 0.008 is generated from psi
coeff 0.006 is generated from psi
coeff 0.005 is generated from psi
coeff 0.003 is generated from psi
                                   3.3
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 4.2
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
                                   4.6
coeff -0.024 is generated from psi
                                   4.7
coeff -0.026 is generated from psi
                                   4.8
coeff -0.028 is generated from psi 4.9
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
            3.7 -.00479666
r18
                               .00479666
                                           .59230593
            3.8 -.00669505
r19
                              .00669505
                                           .45731304
            3.9 -.00859969
                              .00859969
r20
                                            .3425138
r21
              4 -.01051072
                              .01051072
                                            .2488326
            4.1 -.01242824
                              .01242824
                                           .17537691
r22
r23
            4.2 -.01435235
                              .01435235
                                            .1199593
            4.3 -.01628313
                              .01628313
                                           .07967563
r24
r25
            4.4 -.01822063
                              .01822063
                                           .05142147
            4.5 -.02016492
r26
                              .02016492
                                           .03227271
r27
            4.6 -.02211603
                              .02211603
                                           .01971433
            4.7 -.02407401
                              .02407401
                                           .01173271
r28
r29
            4.8 -.02603888
                              .02603888
                                           .00680955
            4.9 -.02801063
r30
                              .02801063
                                           .00385828
r31
              5 -.02998926
                              .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]
> }
  3.4
: 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 **/
mata
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
            log pseudolikelihood = -877.82423
Iteration 4:
                                                 Number of obs = 1,566
Logistic regression
                                                 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
       qsmk | Coefficient std. err. z P>|z| [95% conf. interval]
  __________
        sex | -.5137295 .1533507 -3.35 0.001 -.8142913 -.2131677
       race | -.8608919 .2099555 -4.10 0.000 -1.272397 -.4493867
```

age	.1151581	.0503079	2.29	0.022	.0165564	.2137598
ا   c.age#c.age 	0007593	.00053	-1.43	0.152	0017981	.0002795
education						
1	4710854	.2247796	-2.10	0.036	9116454	0305255
2	5000247	.220776	-2.26	0.024	9327378	0673116
3	3833802	.1954991	-1.96	0.050	7665515	0002089
4	4047148	.2833093	-1.43	0.153	9599908	.1505613
I						
smokeintens~y	0783426	.0146634	-5.34	0.000	1070824	0496029
<u> </u>						
c.						
smokeintens~y#						
C.		0000655	4.04	0.000	0005510	0015005
smokeintens~y	.0010722	.0002655	4.04	0.000	.0005518	.0015925
smokeyrs	0711099	.0263523	-2.70	0.007	1227596	0194602
c.smokeyrs#						
c.smokeyrs	.0008153	.0004486	1.82	0.069	0000639	.0016945
I						
exercise						
0	3800461	.1890123	-2.01	0.044	7505034	0095887
1	0437044	.137269	-0.32	0.750	3127467	.225338
ا محدثمه ا						
active   0	2134564	.2121759	-1.01	0.314	6293135	.2024007
1	2134564 1793322	.2121759	-1.01 -0.87	0.314	6293135 5852109	.2024007
1 1	1793322	.2070040	-0.07	0.300	5052109	.2205400
wt71	0076609	.0255841	-0.30	0.765	0578048	.042483
c.wt71#c.wt71	.0000866	.0001572	0.55	0.582	0002216	.0003947
I						
_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)

```
----- mata (type end to exit) -----
: 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		of obs =	1,566
Model	14412.558	20	720.6279		1545) =	13.45 0.0000
·	82763.0286		53.5683033	R-squa		
residual	02/03.0200	1,545	55.5005055	-	-squared =	
Total	97175.5866	1 565	62.0930266	_	Squared =	0.1373 7.319
TOTAL	91113.3000	1,000	02.0930200	1,000 F	IDE -	7.519
wt82_71	Coefficient	Std. err	. t	P> t	[95% conf.	interval]
qsmk	2.559594	.8091486	3.16	0.002	.9724486	4.14674
qsmkintensity	.0466628	.0351448	1.33	0.184	0222737	.1155993
smokeintens~y	.0491365	.0517254	0.95	0.342	052323	.1505959
I						
c.						
smokeintens~y#						
c.						
smokeintens~y	0009907	.000938	-1.06	0.291	0028306	.0008493
I						
sex	-1.430272	.4689576	-3.05	0.002	-2.350132	5104111
race	.5601096	.5818888	0.96	0.336	5812656	1.701485
age	.3596353	.1633188	2.20	0.028	.0392854	.6799851
I						
c.age#c.age	006101	.0017261	-3.53	0.000	0094868	0027151
1						
education						
1		.7413692	0.26	0.793	-1.259219	1.649173
2		.7012116	1.41	0.160	390006	2.360848
3		.6339153	1.19	0.236	4921358	1.994715
4	1.686547	.8716593	1.93	0.053	0232138	3.396307
I						
smokeyrs	.1343686	.0917122	1.47	0.143	045525	.3142621
I						
c.smokeyrs#						
c.smokeyrs	0018664	.0015437	-1.21	0.227	0048944	.0011616
l						
exercise						
0		.5588587		0.527	-1.450114	.7422889
1	0579374	.4316468	-0.13	0.893	904613	.7887381
active						
0		.6845577		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
c.wt71#c.wt71	0009653	.0005247	-1.84	0.066	0019945	.0000639
_cons	-1.690608 	4.388883	-0.39	0.700	-10.2994	6.918188
( 1) qsmk +	5*qsmkintensit	ty = 0				
_	Coefficient				[95% conf.	interval]
•					1.482117	4.1037
	40*qsmkintens:  Coefficient		t	 P> t	 [95% conf.	 interval]
+						
(1)	4.426108 	.8477818 	5.22 	0.000 	2.763183 	6.089032
	SS 	df			er of obs = , 1546) =	•
	14318.1239				> F =	
Residual	82857.4627 	1,546 	53.5947365		uared = R-squared =	
Total	97175.5866	1,565	62.0930266	_	MSE =	
wt82_71	Coefficient	Std. err.	 t	P> t	[95% conf.	interval]
qsmk	3.462622	.4384543	7.90	0.000	2.602594	4.32265
smokeintens~y	.0651533	.0503115	1.29	0.196	0335327	.1638392
c. smokeintens~y# c.	Ì					
smokeintens~y	0010468	.0009373	-1.12	0.264	0028853	.0007918
sex	-1.46505	.468341	-3.13	0.002	-2.3837	5463989
race		.5816949	1.01	0.314	5545827	1.727406
age		.1633431	2.22	0.027	.0422649	.6830599
c.age#c.age	  0061377	.0017263	-3.56	0.000	0095239	0027515

1						
education						
1 l	.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
1						
c.smokeyrs#						
c.smokeyrs	001827	.0015438	-1.18	0.237	0048552	.0012012
1						
exercise						
0	3628786	.5589557	-0.65	0.516	-1.45927	.7335129
1 l	0421962	.4315904	-0.10	0.922	8887606	.8043683
1						
active						
0	.2580374	.6847219	0.38	0.706	-1.085044	1.601119
1 l	68492	.6740787	-1.02	0.310	-2.007125	.6372851
1						
wt71	.0373642	.0831658	0.45	0.653	1257655	.200494
1						
c.wt71#c.wt71	0009158	.0005235	-1.75	0.080	0019427	.0001111
ĺ						
cons	-1.724603	4.389891	-0.39	0.694	-10.33537	6.886166

# Prorgam 15.2

- $\bullet\,$  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
                                                  Pseudo R2
Log likelihood = -838.44842
                                                              = 0.0611
       qsmk | Coefficient 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
                                    -1.54 0.124
                          .0005361
                                                   -.0018753
                                                               .0002262
   education |
         1 | -.4759606
                                    -2.10
                                           0.035
                        .2262238
                                                   -.9193511
                                                              -.0325701
         2 | -.5047361
                                    -2.32 0.020
                          .217597
                                                   -.9312184 -.0782538
         3 | -.3895288
                                    -2.03 0.042
                                                   -.7647351
                                                              -.0143226
                          .1914353
         4 | -.4123596 .2772868
                                    -1.49 0.137
                                                   -.9558318
                                                             .1311126
smokeintens~y | -.0772704 .0152499
                                  -5.07 0.000
                                                   -.1071596 -.0473812
```

c.						
smokeintens~y#						
c. l						
smokeintens~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 I	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

\_\_\_\_\_

#### -> qsmk = No smoking cessation

Variable		Std. dev.	 Max
	. 2392928		.6814955

-----

### -> qsmk = Smoking cessation

Variable	0bs	Mean	Std. dev.	Min	Max
	403	.3094353	.1290642	.0598799	.7768887

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

#### -> qsmk = No smoking cessation

+		+
obs:	ps	seqn 
1		'
979.	.0510008	22941
945.	.0527126	1769
1023.	.0558418	21140
115.	.0558752	2522
478.	.0567372	12639
+		+

+-				-+
1	463.	.6337243	17096	١
1	812.	.6345721	17768	١
1	707.	.6440308	19147	١
1	623.	.6566707	21983	١
1	1033.	.6814955	22773	١
+-				-+

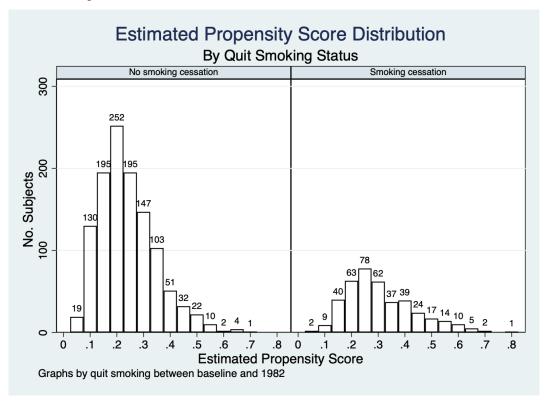
\_\_\_\_\_

#### -> qsmk = Smoking cessation

+		+
obs:	ps	seqn
1223.	.0598799	4289
1283.	.0600822	23550
1253.	.0806089	24306
1467.	.0821677	22904
1165.	.1021875	24584
+		+
+		+
1399.	.635695	17738

```
| 1173. .6659576 22272 |
| 1551. .7166381 14983 |
| 1494. .7200644 24817 |
| 1303. .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				Min	Max	
·			.0185215	.0510008	.1240482	
-> ps_dec = 2						
Variable		Mean		Min	Max	
ps I	157	.1430792	.0107751	.1241923	.1603558	
Variable	0bs	Mean	Std. dev.	Min	Max	
ps	156	. 1750423	.008773	.1606041	.1893271	
-> ps_dec = 4						
Variable	Obs	Mean	Std. dev.	Min	Max	
ps	157	.2014066	.0062403	.189365	.2121815	
Variable	Obs	Mean	Std. dev.	Min	Max	
ps	156	. 2245376	.0073655	.2123068	. 237184	
		Mean	Std. dev.	Min	Max	
ps		. 2515298	.0078777	.2377578	.2655718	
Variable	Obs	Mean	Std. dev.	Min	Max	

ps	I	157	. 2827476	.0099	986	. 265572	4 .2994	968
-> ps_dec = 3	8							
Variable			Mean					
ps	I	156	.3204104	.0125	5102	. 299758	1 .3438	3773
			Mean					Max
			.375637					631
			Mean					Max
			.5026508					8887
	test w	_						
Group	Obs	Me	ean Std.	err.	Std.	dev. [	95% conf.	interval]
No smoki   Smoking	146 11	3.742 3.9497	236 .6533 703 2.333	1341 2995	7.891 7.737	.849 2 7668 -1	.451467	5.033253
Combined	157	3.7568		0464	7.856	8869		
diff		20734	131 2.464	4411		-5	.075509	4.660822
			mean(Smoki				t	= -0.0841 = 155
Pr(T < t) =	Ha: diff < 0							
-> ps_dec = :	2							

Two-sample t test with equal variances

		Mean				
		2.813019				
		7.726944				
Combined	157	3.532893	.5519826	6.916322		4.623217
•		-4.913925			-7.907613	
	nean(No s	moki) - mean	(Smoking)	Degrees	t of freedom	= -3.2425 = 155
Ha: diff	< 0		Ha: diff !=	0	Ha: d	iff > 0
		Pr(				
	test wi	th equal var				
-		Mean				
•		3.25684				
_		7.954974		7.504324	5.045101	10.86485
Combined	156	4.100095	.5245749	6.551938	3.063857	5.136334
		-4.698134			-7.301973	-2.094294
diff = m $HO: diff = 0$		moki) - mean	(Smoking)	Degrees	t of freedom	= -3.5644 = 154
Ha: diff	< 0		Ha: diff !=	0	Ha: d	iff > 0
Pr(T < t) =	0.0002	Pr(	T  >  t ) =	0.0005	Pr(T > t	a) = 0.9998
	4					
Two-sample t	test wi	th equal var	iances			
_		Mean				
•		3.393929				
Smoking	36	5.676072				
Combined		3.917223	.5412091	6.78133	2.848179	4.986266
diff		-2.282143	1.278494		-4.807663	.2433778

diff = mean(No smoki) - mean(Smoking) t = -1.7850H0: diff = 0Degrees of freedom = 155 Ha: diff < 0 Ha: diff != 0 Ha: diff > 0Pr(T < t) = 0.0381Pr(|T| > |t|) = 0.0762Pr(T > t) = 0.9619\_\_\_\_\_\_  $-> ps_dec = 5$ Two-sample t test with equal variances \_\_\_\_\_\_ Mean Std. err. Std. dev. [95% conf. interval] Obs Group | \_\_\_\_\_ No smoki | 119 .8042619 8.773461 -.2242199 1.368438 2.961095 Smoking | 37 5.195421 1.388723 8.44727 2.378961 8.011881 \_\_\_\_\_\_\_ .7063778 8.822656 Combined | 156 2.27612 .8807499 \_\_\_\_\_\_\_\_\_\_ diff | -3.826983 1.637279 -7.061407 \_\_\_\_\_ diff = mean(No smoki) - mean(Smoking) t = -2.3374H0: diff = 0Degrees of freedom = 154 Ha: diff < 0 Ha: diff != 0 Ha: diff > 0 Pr(T < t) = 0.0104 Pr(|T| > |t|) = 0.0207Pr(T > t) = 0.9896 $\rightarrow$  ps\_dec = 6 Two-sample t test with equal variances \_\_\_\_\_ Group | Obs Mean Std. err. Std. dev. [95% conf. interval] \_\_\_\_\_ No smoki | 2.25564 .6850004 7.249362 112 .8982664 3.613014 45 7.199088 1.724899 11.57097 Smoking | 3.722782 10.67539 Combined | 157 3.672552 .7146582 8.954642 2.260897 diff | -4.943447 1.535024 -7.975714 -1.911181 diff = mean(No smoki) - mean(Smoking) t = -3.2204H0: 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

 $<sup>\</sup>rightarrow$  ps\_dec = 7

Two-sample t test with equal variances

-		Mean				
No smoki   Smoking	116 41	.7948483 6.646091	.7916172 1.00182	8.525978 6.414778	773193 4.621337	2.36289 8.670844
Combined	157	2.32288	.6714693	8.413486	.9965349	3.649225
diff		-5.851242	1.45977		-8.734853	-2.967632
	mean(No s	smoki) - mean				= -4.0083
Ha: di: Pr(T < t)		Pr(	Ha: diff != T  >  t ) =			
-> ps_dec :	= 8					

Two-sample t test with equal variances

Group		Mean	Std. err.		[95% conf.	- · · · · -
No smoki Smoking	l 107 l 49	1.063848 3.116263	.5840159 1.113479	6.041107 7.794356	0940204 .8774626	2.221716 5.355063
Combined	l 156	1.708517		6.684666	.6512864	2.765747
diff	I	-2.052415	1.144914		-4.31418	.2093492
diff =		smoki) - mean	n(Smoking)			= -1.7926

HO: diff = 0Degrees of freedom = 154

 $\rightarrow$  ps\_dec = 9

#### Two-sample t test with equal variances

Group	Obs	Mean	Std. err.		[95% conf.	_
No smoki   Smoking	100 57	0292906 .9112647	.7637396 .9969309	7.637396 7.526663	-1.544716 -1.085828	1.486134 2.908357
Combined	157	.3121849	.6054898		8838316	1.508201

```
-.9405554 1.26092
                                   -3.43136 1.550249
  diff |
  diff = mean(No smoki) - mean(Smoking)
H0: diff = 0
                              Degrees of freedom =
  Ha: diff < 0
                    Ha: diff != 0
                                      Ha: diff > 0
Pr(T < t) = 0.2284
               Pr(|T| > |t|) = 0.4568
                                   Pr(T > t) = 0.7716
 ._____
-> ps_dec = 10
Two-sample t test with equal variances
______
 Group |
               Mean Std. err. Std. dev. [95% conf. interval]
        Obs
______
        80 -.768504 .9224756 8.250872 -2.604646 1.067638
No smoki |
        76 2.39532 1.053132 9.180992 .2973737 4.493267
Smoking |
__________
Combined | 156 .7728463 .7071067 8.831759 -.6239631
__________
            -3.163824 1.396178
                                   -5.921957
______
  diff = mean(No smoki) - mean(Smoking)
                                       t = -2.2661
H0: diff = 0
                              Degrees of freedom = 154
  Ha: diff < 0
                   Ha: diff != 0
                                      Ha: diff > 0
Pr(T < t) = 0.0124 Pr(|T| > |t|) = 0.0248
                                   Pr(T > t) = 0.9876
   Source | SS df MS
                              Number of obs =
                                            1,566
 ------ F(10, 1555) =
                                            9.87
    Model | 5799.7817
                   10 579.97817 Prob > F
                                        = 0.0000
  Residual | 91375.8049 1,555 58.7625755 R-squared
                                        = 0.0597
   ------ Adj R-squared =
                                            0.0536
    Total | 97175.5866
                 1,565 62.0930266 Root MSE
                                            7.6657
       ._____
   wt82_71 | Coefficient Std. err. t P>|t| [95% conf. interval]
   ______
    qsmk |
          3.356927 .4580399 7.33 0.000
                                  2.458486 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
          4.36015 .8793345
                        4.96 0.000
      3 l
                                  2.635343
                                           6.084956
      4 |
          4.010061
                .8745966
                        4.59 0.000
                                 2.294548
                                           5.725575
          2.342505
                .8754878
                        2.68 0.008
      5 I
                                   .6252438 4.059766
      6 l
         3.572955
                .8714389
                        4.10 0.000
                                  1.863636
                                           5.282275
      7 | 2.30881
                .8727462
                        2.65 0.008 .5969261 4.020693
```

# 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
qenerate 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 Pseudo R2 = 0.0611

Log likelihood = -838.44842

qsmk	Coefficient	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
smokeintens~y	  0772704 	.0152499	-5.07	0.000	1071596	0473812
С.						
smokeintens~y#						
С.	•					
smokeintens~y	.0010451	.0002866	3.65	0.000	.0004835	.0016068
smokeyrs	I   −.0735966 	.0277775	-2.65	0.008	1280395	0191538
c.smokeyrs#						
c.smokeyrs		.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
wt71		0152357	.0263161	-0.58	0.563	0668144	.036343
c.wt71#c.wt71		.0001352	.0001632	0.83	0.407	0001846	.000455
_cons	1	-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)

Expanded observation   type	   Freq.	Percent	Cum.
-1	1,566	33.33	33.33
Original observation	1,566	33.33	66.67
Duplicated observation	1,566	33.33	100.00
Total	+ l 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	Obs	Mean	Std. dev.	Min	Max
qsmk	1,566	.2573436	. 4373099	0	1

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

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

Variable	Obs	Mean	Std. d	lev.	Min	Ма	х
•	1,566	0		0	0		0
-> interv = Dup	plicat						
Variable	0bs	Mean	Std. d	lev.	Min	Ma	x -
qsmk	1,566	1		0	1		1
	SS	df			per of obs		
	5287.31428			- (0)	1562) > F		0.0000
	91888.2723				quared		
+-					R-squared		
Total	97175.5866	1,565	62.093026	_	=		7.6699
wt82_71	Coefficient		t		 [95%	conf.	interval]
Smoking ce		1.13904	3.54	0.000	1.80	225	6.270665
	-12.3319						
qsmk#c.ps							
Smoking ce	-2.038829 	3.649684	-0.56	0.576	-9.197	'625	5.119967
_cons	4.935432	.5570216	8.86	0.000	3.842	2843	6.028021
-> Interv = -1							
	0bs						x -
•	1,566						1
-> interv = Original							
Variable	0bs	Mean	Std. d	lev.	Min	Ma	x -
	1,566			164 -4.6	345079 4	.30649	6

#### -> interv = Duplicat

Variable	l Obs	Mean	Std. dev.	Min	Max
predY	   1,566	5.273676	1.670225	-2.192565	8.238971

#### observe[4,2]

	interv	value
observed	-1	2.6382998
E(Y(a=0))	0	1.7618979
E(Y(a=1))	1	5.2736757
difference		3.5117778

#### (3,132 observations deleted)

(1,566 missing values generated)

11. predict ps\_b, pr
12.

Command: bootstdz /
 EY\_a0: r(boot\_0)
 EY\_a1: r(boot\_1)
difference: r(boot\_diff)

# Simulations (500)

 50
 100
 150
 200
 250
 300
 350
 400
 450

			••••	50	0	
pe[1,3] E(Y(a=0 value 1.76189	))) E(Y(a=1 979 5.27367					
Bootstrap resul	lts					f obs = 1,62 ions = 50
	Observed coefficient					mal-based nf. interval
EY_aO	1.761898 5.273676	.2255637 .4695378	7.81 11.23	0.000	4.35339	9 6.19395
Bootstrap resul	lts				of obs tions	•
	Observed		Boots	 trap		

| coefficient Bias std. err. [95% conf. interval]

.49707894

2.186845

6.109205

4.424034

2.466025

(P)

(P)

(P)

EY\_a0 | 1.7618979 .0026735 .22556365 1.269908

EY\_a1 | 5.2736757 -.0049491 .46953779 4.34944

3.5117778 -.0076226

Key: P: Percentile

difference |

# 16. Instrumental variables estimation: Stata

- 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

/*Calculate P[Y/Z=z]*/
ttest wt82_71, by(highprice)</pre>
```

```
/*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
nıım
denom
st_numscalar("iv_est", iv_est)
di scalar(iv_est)
```

file ./data/nhefs-highprice.dta saved

+-			+
I	Key		I
-			-
1	fı	requency	١
1	row	percentage	1
+-			+

Total	and 1982	quit smoki baseline No smokin	ĺ
, local	Silloking C	NO SHOKIH	nignprice (
100.00	8 19.51	33 80.49	0
1,435			1
	378 25.61	1,098 74.39	Total

Two-sample t test with equal variances

-			Std. err.			interval]
0   1	41 1,435	2.535729 2.686018	1.461629 .2084888	9.358993 7.897848	4183336 2.277042	3.094994
Combined	1,476	2.681843	. 2066282	7.938395	2.276527	3.087159
diff		1502887			-2.617509	2.316932
	mean(0)					= -0.1195
	ff < 0 = 0.4525	Pr(	Ha: diff !=	-		iff > 0 ) = 0.5475

Numerator, E[Y|Z=1] - E[Y|Z=0] = .15

Denominator: denom = P[A=1|Z=1] - P[A=1|Z=0] = .063

IV estimator = 2.3809524

```
pa[1,2]
                   c2
         c1
r1 .19512195 .25783972
ey[1,2]
                   c2
         c1
    2.535729 2.6860178
r1
----- mata (type end to exit) -----
: 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
  .1502887173
: denom
  .06271777
: st_numscalar("iv_est", iv_est)
```

#### 2.3962701

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

```
/* 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

Instrumented: qsmk
Instruments: highprice

- 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
  highprice | Coefficient Std. err. z P>|z|
                                                        [95% conf. interval]
```

```
hspi | 2.75e-07 .0201749 0.00 1.000 -.0395419 .0395424

_cons | 3.555347 .1637931 21.71 0.000 3.234319 3.876376
```

- Estimating the average causal effect using the standard IV estimator based on alternative proposed instruments
- Data from NHEFS
- Section 16.5

```
use ./data/nhefs-highprice, clear
/*Instrument cut-point: 1.6*/
replace highprice = .
replace highprice = (price82 >1.6 & price82 < .)</pre>
ivregress 2sls wt82_71 (qsmk = highprice)
/*Instrument cut-point: 1.7*/
replace highprice = .
replace highprice = (price82 >1.7 & price82 < .)</pre>
ivregress 2sls wt82_71 (qsmk = highprice)
/*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
                                                                 =
                                                                 = 18.593
                                                  Root MSE
```

_	Coefficient				[95% conf.	interval]
qsmk	41.28124	164.8417	0.25	0.802		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 = .

Root MSE = 20.577

wt82\_71 | Coefficient 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

-----

_	Coefficient					interval]
	•				-76.78374	34.57691
_cons	8.086377	7.283314	1.11	0.267	-6.188657	22.36141

Instrumented: qsmk
Instruments: highprice

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

```
(1,476 real changes made)
```

Instrumental v	Instrumental variables 2SLS regression				er of obs chi2(1) > chi2 ared MSE	= = = =	1,476 0.29 0.5880
_	Coefficient				/ ·	conf.	interval]
qsmk     cons	-12.81141	23.65099	-0.54 0.98	0.588	-59.16		33.54368 17.84599

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 | Coefficient Std. err.
                                                P>|z|
                                                        [95% conf. interval]
        qsmk | -1.042295 29.86522 -0.03 0.972
                                                        -59.57705
                                                                     57.49246
```

3.490938

sex | -1.644393 2.620115 -0.63 0.530 -6.779724

race	I	1832546	4.631443	-0.04	0.968	-9.260716	8.894207
age		16364	.2395678	-0.68	0.495	6331844	.3059043
smokeintens~y	1	.0057669	.144911	0.04	0.968	2782534	.2897872
smokeyrs	 	.0258357	.1607639	0.16	0.872	2892558	.3409271
exercise	1						
1	1	.4987479	2.162395	0.23	0.818	-3.739469	4.736964
2	1	.5818337	2.174255	0.27	0.789	-3.679628	4.843296
	1						
active	1						
1	1	-1.170145	.6049921	-1.93	0.053	-2.355908	.0156176
2	1	5122842	1.303121	-0.39	0.694	-3.066355	2.041787
	1						
wt71	1	0979493	.036123	-2.71	0.007	168749	0271496
_cons		17.28033	2.32589	7.43	0.000	12.72167	21.83899

-----

Instrumented: qsmk

 ${\tt Instruments: sex \ race \ age \ smokeintensity \ smokeyrs \ 1.exercise \ 2.exercise}$ 

1.active 2.active wt71 highprice

## 17. Causal survival analysis: Stata

- 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 |
```

between	١	quit smoki	ng between		
1983 and		baseline	and 1982		
1992	١	No smokin	Smoking c	1	Total
	+			-+-	
0	1	963	312	1	1,275
1	1	200	91	1	291
	+			-+-	
Total	١	1,163	403	I	1,566

Survival-time data settings

Failure event: death==1
Observed 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

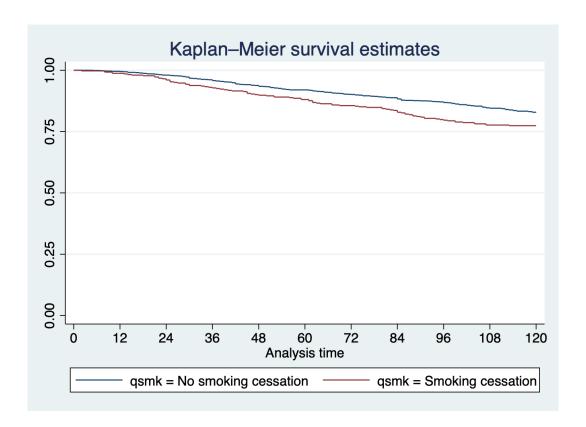
At risk from t = 0

0

Earliest observed entry t =

Last observed exit t = 120

Failure \_d: death==1
Analysis time \_t: survtime



- 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)
                Freq.
                          Percent
     event |
                                         Cum.
         0 | 170,785
                              99.83
                                         99.83
         1 |
                   291
                             0.17
                                         100.00
```

100.00

Total | 171,076

event	Odds ratio	Std. err.	z	P> z	[95% conf.	interval]
qsmk	1.402527	.6000025	0.79	0.429	.6064099	3.243815
qsmk#c.time						
Smoking ce	1.012318 	.0162153	0.76	0.445	.9810299	1.044603
qsmk#c.time#						
c.time						
Smoking ce	.9998342	.0001321	-1.25	0.210	.9995753	1.000093
time	1.022048	.0090651	2.46	0.014	1.004434	1.039971
c.time#c.time	.9998637	.0000699	-1.95	0.051	.9997266	1.000001
_cons	.0007992	.0001972	-28.90	0.000	.0004927	.0012963

Note: \_cons estimates baseline odds.

(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 = Original

Variable	Obs	Mean	Std. dev.	Min	Max
psurv	1,566	.8279829	0	.8279829	.8279829

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

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

Variable	Obs	Mean	Std. dev.	Min	Max
psurv	1,566	.774282	0	.774282	.774282

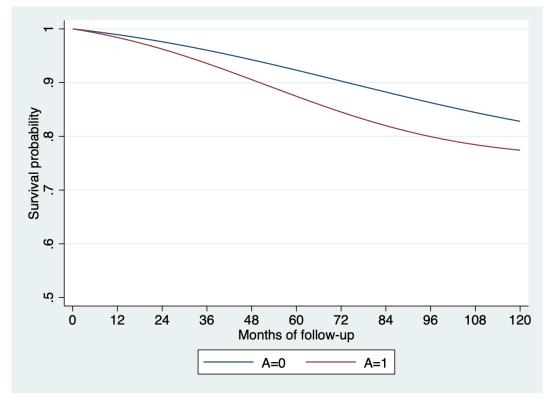
(3,132 observations created)

(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
psurv0	float	%9.0g		psurv, interv == Original observation
psurv1	float	%9.0g		<pre>psurv, interv == Duplicated   observation</pre>



## Program 17.3

 $\bullet\,$  Estimation of survival curves via IP weighted hazards model

- Data from NHEFS
- Section 17.4
- Generates Figure 17.6

```
use ./data/nhefs_surv, clear
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 newseqn 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

Logistic regression

Number of obs = 1,566 LR chi2(18) = 109.16 Prob > chi2 = 0.0000 Pseudo R2 = 0.0611

Log likelihood = -838.44842

qsmk	Coefficient	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     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
smokeintens~y	0772704	.0152499	-5.07	0.000	1071596	0473812
c.						
smokeintens~y#						
c.		000000	0.05	0 000	0004005	004000
smokeintens~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 1	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	.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 = 1,566

LR chi2(0) = -0.00

Prob > chi2 = .

Log likelihood = -893.02712 Pseudo R2 = -0.0000

\_\_\_\_\_

(128,481 missing values generated)

(128,481 real changes made)

Variable | Obs Mean Std. dev. Min Max -----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) = 22.74Prob > chi2 = 0.0004

Log pseudolikelihood = -2126.8554 Pseudo R2 = 0.0045

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

```
qsmk#c.time#|
     c.time |
Smoking ce.. | -.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 = Original
   Variable | Obs Mean Std. dev. Min
     psurv | 1,566 .8161003
                                   0 .8161003 .8161003
-> interv = Duplicat
   Variable | Obs Mean Std. dev. Min Max
```

psurv | 1,566 .8116784 0 .8116784 .8116784

```
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)

Variable name	Storage type	Display format	Value label	Variable label
psurv0	float	%9.0g		psurv, interv == Original observation
psurv1	float	%9.0g		psurv. interv == Duplicated

observation

#### (3,132 observations deleted)

```
5. predict p_qsmk, pr
6.
11.
23. drop if time != 119
24. bysort interv_b: egen meanS_b = mean(psurv)
25. keep newseqn qsmk meanS_b
26. drop if newseqn != 1 /* only need one pair */
27.

r; t=0.00 18:03:30

Command: bootipw_surv
    PrY_a0: r(boot_0)
    PrY_a1: r(boot_1)
```

difference: r(boot\_diff)

Simulations (10)

. . . . . . . . . .

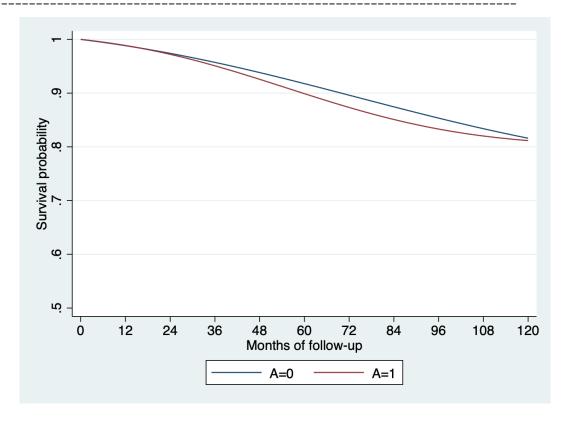
r; t=28.44 18:03:58

Bootstrap results

Number of obs = 1,629

Replications = 10

	Observed coefficient	std. err.			[95% conf.	-based interval]
PrY_a0	.8161003 .8116784	.0093124 .0237581 .0225007	87.64 34.16 -0.20	0.000 0.000 0.844	.7978484 .7651133 0485224	.8343522 .8582435 .0396786



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

- Section 17.5
- Generates Figure 17.7

```
use ./data/nhefs_surv, clear
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 = Original
               Obs Mean Std. dev. Min Max
   Variable |
______
      psurv | 1,566 .8160697 .2014345 .014127 .9903372
```

-> interv = Duplicat Variable | Obs Mean Std. dev. Min Max psurv | 1,566 .811763 .2044758 .0123403 .9900259 (372,708 missing values generated) -> interv = Original Variable | Obs Mean Std. dev. Min Max 0 .8160697 .8160697 1,566 .8160697 meanS | -> interv = Duplicat Variable | Obs Mean Std. dev. Min Max meanS | 1,566 .8117629 0 .8117629 .8117629 (3,132 observations created) (3,132 real changes made) (375,840 missing values generated) (375,840 real changes made) Variable Storage Display Value type format label Variable label name\_\_\_\_\_\_ meanS\_t0 float %9.0g meanS\_t, interv == Original

observation

file /Users/tom/Documents/GitHub/cibookex-r/figs/stata-fig-17-4.png saved as PNG format

(3,132 observations deleted)

5. drop if time != 0

6. /\*only predict on new version of data \*/

r; t=0.00 18:04:13

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

Simulations (10)

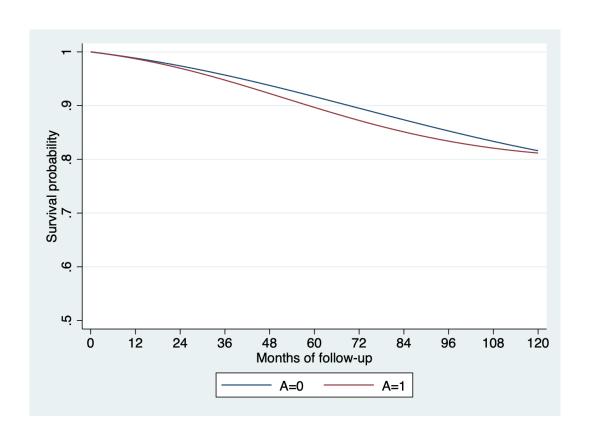
. . . . . . . . . .

r; t=31.74 18:04:45

Bootstrap results

Number of obs = 1,629 Replications = 10

| Observed Bootstrap | Normal-based | coefficient std. err. | z | P>|z| | [95% conf. interval] | PrY\_a0 | .8160697 .0087193 | 93.59 | 0.000 | .7989802 .8331593 | PrY\_a1 | .8117629 .0292177 | 27.78 | 0.000 | .7544973 | .8690286 | difference | -.0043068 | .0307674 | -0.14 | 0.889 | -.0646099 | .0559963



## Session information: Stata

```
library(Statamarkdown)
For reproducibility.
about
Revision 15 Feb 2022
Copyright 1985-2021 StataCorp LLC
Total physical memory: 8.01 GB
Stata license: Unlimited-user 2-core network, expiring 21 Jan 2023
Serial number: 501709302483
 Licensed to: Tom Palmer
              University of Bristol
# install.packages("sessioninfo")
sessioninfo::session_info()
- Session info -----
setting value
version R version 4.1.2 (2021-11-01)
         macOS Big Sur 10.16
system x86_64, darwin17.0
ui
         X11
language (EN)
collate en_GB.UTF-8
ctype en_GB.UTF-8
         Europe/London
tz
date
         2022-02-20
         2.17.1.1 @ /Applications/RStudio.app/Contents/MacOS/quarto/bin/ (via rmarkdown)
pandoc
- Packages -----
package
              * version date (UTC) lib source
bookdown
                0.24
                        2021-09-02 [1] CRAN (R 4.1.0)
cli
                3.2.0 2022-02-14 [1] CRAN (R 4.1.2)
                0.6.29 2021-12-01 [1] CRAN (R 4.1.0)
digest
evaluate
              0.15 2022-02-18 [1] CRAN (R 4.1.2)
fastmap
                1.1.0 2021-01-25 [1] CRAN (R 4.1.0)
htmltools
                0.5.2 2021-08-25 [1] CRAN (R 4.1.0)
```

```
1.37
                     2021-12-16 [1] CRAN (R 4.1.0)
knitr
magrittr
            2.0.2 2022-01-26 [1] CRAN (R 4.1.2)
             1.0.1 2022-02-03 [1] CRAN (R 4.1.2)
rlang
            2.11
                     2021-09-14 [1] CRAN (R 4.1.0)
rmarkdown
rstudioapi
              0.13
                     2020-11-12 [1] CRAN (R 4.1.0)
             1.2.2 2021-12-06 [1] CRAN (R 4.1.0)
sessioninfo
Statamarkdown * 0.7.0 2022-02-04 [1] Github (Hemken/Statamarkdown@a68a8b9)
             1.7.6 2021-11-29 [1] CRAN (R 4.1.0)
stringi
            1.4.0 2019-02-10 [1] CRAN (R 4.1.0)
stringr
xfun
            0.29
                     2021-12-14 [1] CRAN (R 4.1.0)
              2.3.4 2022-02-17 [1] CRAN (R 4.1.2)
yaml
```

[1] /Library/Frameworks/R.framework/Versions/4.1/Resources/library

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## Bibliography

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