P8123 Final

Farizah Rob, Emily Potts, Harry Wang, William Anderson

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```
library(tidyverse)
library(survey)
library(ggsurvey)
library(tableone)
library(gtsummary)
library(sjPlot)

knitr::opts_chunk$set(
    message = FALSE,
    warning = FALSE
)

theme_set(theme_minimal() + theme(legend.position = "bottom"))

load("./data/nhanes_data.rda") #put files into data folder so everyone can run this
load("./data/nhanes_key.rda")
load("./data/key_guide.rda")
```

Problem statement: Controlling blood pressure (BP) reduces the risk for cardiovascular disease. However, the prevalence of BP control (i.e., systolic BP < 140 and diastolic BP < 90) among US adults with hypertension has decreased since 2013. We invite teams to analyze publicly available data from US adults to help identify potential causes or correlates of worsening BP control among US adults with hypertension over the past decade, as this may allow for development of effective interventions to help control BP and prevent cardiovascular disease.

Svydesign and subset for hypertension patients 2013 onwards

```
dstrat <- svydesign(ids=~svy_psu, strata=~svy_strata, weights=~svy_weight_mec, data=nhanes_data, nest=The late to subset(dstrat, svy_subpop_htn==1) #limited to hypertension patients

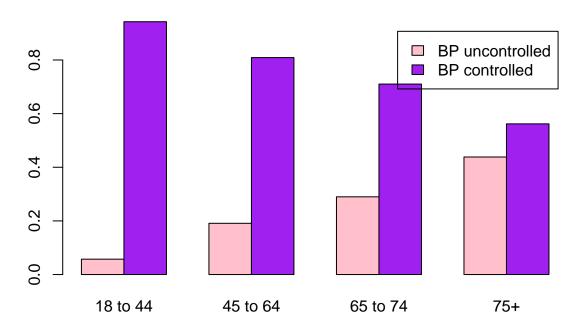
htsub2 <- subset(htsub, svy_year=="2013-2014" | svy_year=="2015-2016" | svy_year=="2017-2020") #limited
```

Some EDA

The outcome of interest -> bp_control_140_90 OR bp_uncontrolled_140_90

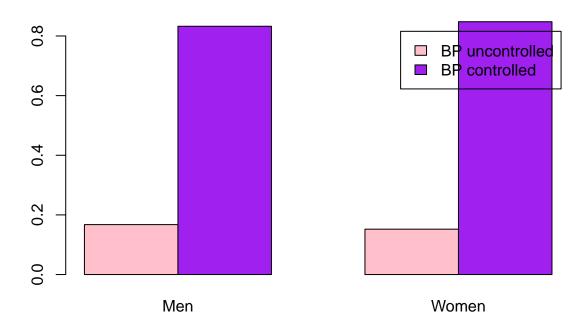
```
#age -> so for older people, BP control is definitely lower
age <- svyby(~factor(bp_control_140_90), ~factor(demo_age_cat), design=htsub2, svymean, na.rm=TRUE)
barplot(age, legend.text=c("BP uncontrolled", "BP controlled"), col=c("pink", "purple"), main="Proporti</pre>
```

Proportion of BP control by age category

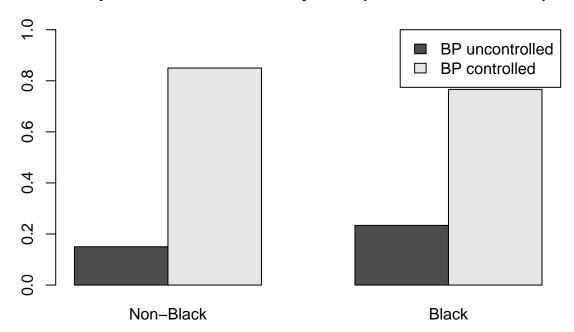


```
#gender -> not really any difference
gender <- svyby(~factor(bp_control_140_90), ~factor(demo_gender), design=htsub2, svymean, na.rm=TRUE)
barplot(gender, legend.text=c("BP uncontrolled", "BP controlled"), col=c("pink", "purple"), main="Proposition")</pre>
```

Proportion of BP control by gender

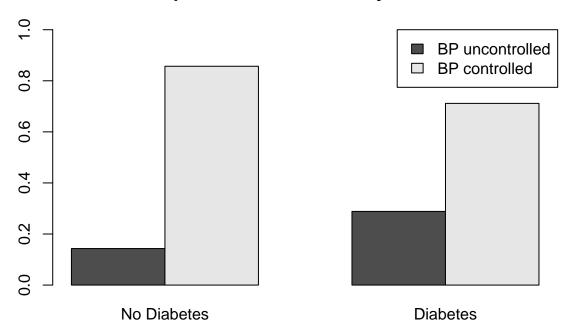


Proportion of BP control by race (non-Black vs Black)

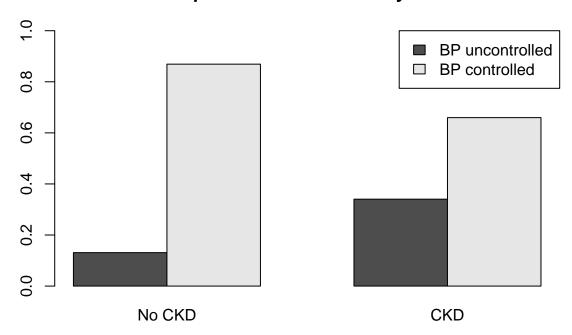


```
#diabetes
diabetes <- svyby(~factor(bp_control_140_90), ~factor(cc_diabetes), design=htsub2, svymean, na.rm=TRUE)
barplot(diabetes,
    legend.text=c("BP uncontrolled", "BP controlled"),
    names.arg=c("No Diabetes", "Diabetes"),
    ylim=c(0, 1.0),
    args.legend = list(x = "topright"),
    main="Proportion of BP control by diabetes")</pre>
```

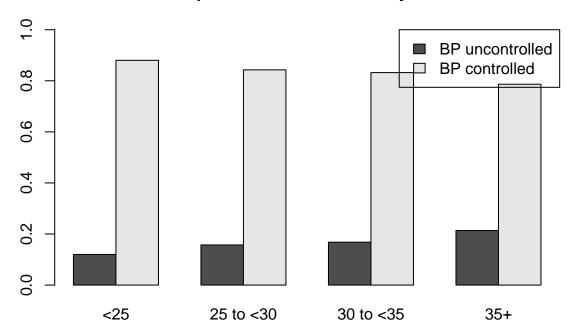
Proportion of BP control by diabetes



Proportion of BP control by CKD



Proportion of BP control by BMI



```
#meds recommended -> 100% for those with uncontrolled BP
# look at med recommended and meds taken ratio by different subgroups?
table(nhanes_data$bp_med_recommended_jnc7, nhanes_data$bp_med_use) #48% of people recommended medicatio
##
##
            No
                 Yes
##
    No 37755
     Yes 7062 14690
##
svyCreateCatTable(vars= c("bp_med_recommended_jnc7" ,"bp_med_use"),
                  strata="demo_race",
                  data=htsub2) #across all races, the % of med use is lower than % recommended -? this
##
                                      Stratified by demo_race
##
                                       Non-Hispanic White Non-Hispanic Black
##
                                       443841370.8
                                                           78256419.3
```

Stratified by demo_race

Non-Hispanic Asian Hispanic

111473163.2 (25.1) 23255215.8 (29.7)

6822774.8 (18.0) 15828214.5 (15.0)

105372525.4

37820092.9

bp_med_recommended_jnc7 = Yes (%) 156173388.2 (35.2) 33534720.2 (42.9)

bp_med_recommended_jnc7 = Yes (%) 11080831.4 (29.3) 26424308.7 (25.1)

##

##

##

##

##

bp_med_use = Yes (%)

bp_med_use = Yes (%)

```
##
                                       Stratified by demo_race
##
                                        Other
                                                          p
                                                                 test
                                        24220134.3
##
##
     bp_med_recommended_jnc7 = Yes (%) 8029041.0 (33.2) <0.001</pre>
     bp_med_use = Yes (%)
                                        5443707.4 (22.5) < 0.001
## table 1
reset_gtsummary_theme()
theme_gtsummary_compact()
tab1 <- tbl_svysummary(data = htsub2, by = "bp_control_140_90", include = c("demo_age_cat", "demo_gende
                       label = list(demo_age_cat ~ "Age",
                                     demo_gender ~ "Gender",
                                     demo_race ~ "Race",
                                     cc_bmi ~ "BMI",
                                     cc_ckd ~ "Chronic Kidney Disease",
                                     cc_diabetes ~ "Diabetes",
                                     cc_cvd_any ~ "Cardiovascular Disease",
                                     bp_med_use ~ "BP Medication Use",
                                     chol_total ~ "Total Cholesterol",
                                     chol_hdl ~ "HDL",
                                     chol_ldl ~ "LDL",
                                     chol_trig ~ "Triglycerides",
                                     chol_med_use ~ "Cholesterol Medication Use")) %>% add_stat_label()
  italicize_levels()
tab1
```

Table 1: Subject Characteristics (Survey Weighted)

Characteristic	No, N = 110,019,543	Yes, N = 579,490,999
Age, n (%)		
18 to 44	$18,218,707 \ (17\%)$	299,580,713 (52%)
45 to 64	45,525,230 (41%)	192,746,908 (33%)
65 to 74	23,816,079 (22%)	58,369,736 (10%)
75+	22,459,527 (20%)	28,793,643 (5.0%)
Gender, n (%)	,	,
Men	55,991,881 (51%)	278,426,528 (48%)
Women	54,027,662 (49%)	301,064,472 (52%)
Race, n (%)	,	,
Non-Hispanic White	$68,928,072 \ (63\%)$	374,913,299 (65%)
Non-Hispanic Black	18,291,583 (17%)	59,964,836 (10%)
Non-Hispanic Asian	$5,715,421 \ (5.2\%)$	$32,104,671 \ (5.5\%)$
Hispanic	13,404,152 (12%)	91,968,373 (16%)
Other	3,680,315 (3.3%)	$20,539,820 \ (3.5\%)$
BMI, n (%)	,	,
<25	$23,597,988 \ (22\%)$	173,248,879 (30%)
25 to <30	33,526,572 (31%)	179,795,985 (32%)
$30 \ to < 35$	23,810,864 (22%)	117,994,322 (21%)
35+	26,600,214 (25%)	97,987,747 (17%)
Unknown	2,483,906	10,464,066
Chronic Kidney Disease, n (%)	32,045,575 (29%)	62,111,324 (11%)
Diabetes, n (%)	22,552,554 (20%)	55,629,143 (9.6%)

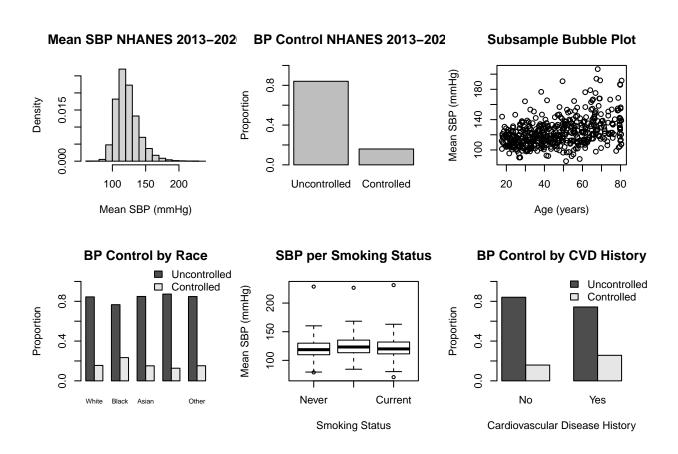
Characteristic	No, N = 110,019,543	Yes, N = 579,490,999
Cardiovascular Disease, n (%)	15,856,616 (14%)	40,856,909 (7.1%)
BP Medication Use, n (%)	51,549,103 (47%)	111,273,973 (19%)
Total Cholesterol, Median (IQR)	192 (163, 222)	183 (158, 211)
Unknown	63,100,407	$325,\!591,\!825$
HDL, Median (IQR)	52 (42, 65)	52(43, 63)
Unknown	63,100,407	325,591,825
LDL, Median (IQR)	112 (88, 139)	109 (86, 133)
Unknown	63,221,679	325,731,647
Triglycerides, Median (IQR)	105 (72, 155)	89 (61, 135)
Unknown	$63,\!221,\!679$	325,731,647
Cholesterol Medication Use, n (%)	16,391,227 (34%)	47,895,434 (19%)
Unknown	62,505,218	323,610,642

Possible Figure 1:

I made a six-panel figure that could serve as our single Figure 1, feel free to change these plots to different things or remove ones we might not need. - Will

```
par(mfrow = c(2, 3))
svyhist(~bp_sys_mean, htsub2, main = "Mean SBP NHANES 2013-2020", xlab = "Mean SBP (mmHg)")
prop <- svymean(~bp_uncontrolled_140_90, htsub2, na.rm=TRUE)</pre>
barplot(prop,
        ylim=c(0, 1.0),
        main="BP Control NHANES 2013-2020", names.arg = c("Uncontrolled", "Controlled"),
        ylab = "Proportion")
 svyplot(bp sys mean~demo age years, htsub2, style = "subsample", xlab = "Age (years)", ylab = "Mean SB
race <- svyby(~bp_uncontrolled_140_90, ~factor(demo_race), htsub2, svymean, na.rm=TRUE)
barplot(race,
        legend.text=c("Uncontrolled", "Controlled"),
        ylim=c(0, 1.0),
        args.legend = list(x = "topright", bty = "n", inset = c(-0.20, -0.2)),
        main="BP Control by Race",
        ylab = "Proportion",
        names.arg = c("White", "Black", "Asian", "Hispanic", "Other"),
        cex.names = 0.6)
svyboxplot(bp_sys_mean~factor(cc_smoke), htsub2, na.rm=TRUE, xlab = "Smoking Status", ylab = "Mean SBP
cvd <- svyby(~bp_uncontrolled_140_90, ~factor(cc_cvd_chd), htsub2, na.rm = T, svymean)</pre>
barplot(cvd,
        legend.text=c("Uncontrolled", "Controlled"),
        vlim=c(0, 1.0),
       args.legend = list(x = "topright", bty = "n", inset = c(-0.20, -0.1)),
        main="BP Control by CVD History",
```





Regression models - really don't know which variables to select

using the bp_uncontrolled_140_90 variable instead of bp_control for regression

```
m1 <- svyglm(bp_uncontrolled_140_90 ~ demo_age_cat + demo_race + demo_gender + cc_smoke + cc_bmi+ cc_di
#need to set BMI reference value to normal level rather than underweight
summary(m1)
```

```
##
## Call:
  svyglm(formula = bp_uncontrolled_140_90 ~ demo_age_cat + demo_race +
##
##
       demo_gender + cc_smoke + cc_bmi + cc_diabetes + cc_ckd +
       cc_cvd_any + chol_trig + chol_ldl + chol_hdl, design = htsub2,
##
##
       family = quasibinomial())
##
## Survey design:
  subset(htsub, svy_year == "2013-2014" | svy_year == "2015-2016" |
##
##
       svy year == "2017-2020")
##
```

```
## Coefficients:
##
                                 Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                               -4.2784289 0.2847878 -15.023 < 2e-16 ***
## demo_age_cat45 to 64
                                1.1528233 0.1252890
                                                       9.201 5.47e-11 ***
## demo_age_cat65 to 74
                                1.7627753
                                           0.1598648
                                                      11.027 4.28e-13 ***
## demo age cat75+
                                2.4495017 0.1681808
                                                     14.565
                                                             < 2e-16 ***
## demo raceNon-Hispanic Black 0.8670093
                                           0.1173396
                                                       7.389 1.02e-08 ***
## demo raceNon-Hispanic Asian 0.3999404
                                           0.1137010
                                                       3.517 0.001199 **
## demo raceHispanic
                                0.1559047
                                           0.1091321
                                                       1.429 0.161743
## demo_raceOther
                                0.0207119
                                           0.2431672
                                                       0.085 0.932594
## demo_genderWomen
                               -0.3087733
                                           0.1070130
                                                      -2.885 0.006566 **
## cc_smokeFormer
                                0.0729305
                                           0.1250743
                                                       0.583 0.563463
## cc_smokeCurrent
                                0.2004411
                                          0.1685633
                                                       1.189 0.242179
## cc_bmi25 to <30
                                          0.1116762
                               -0.0215815
                                                     -0.193 0.847849
## cc_bmi30 to <35
                                0.2261152 0.1130356
                                                       2.000 0.053045 .
## cc_bmi35+
                                0.7023231
                                           0.1345164
                                                       5.221 7.63e-06 ***
## cc_diabetesYes
                                0.3421597
                                           0.1262108
                                                       2.711 0.010214 *
## cc ckdYes
                                0.6520772 0.1190915
                                                       5.475 3.49e-06 ***
## cc_cvd_anyYes
                               -0.0273710
                                           0.1208875
                                                      -0.226 0.822158
## chol trig
                                0.0017308
                                           0.0003819
                                                       4.533 6.21e-05 ***
## chol_ldl
                                0.0052070
                                           0.0012622
                                                       4.125 0.000209 ***
## chol hdl
                                0.0061692
                                           0.0032180
                                                       1.917 0.063190 .
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for quasibinomial family taken to be 0.9486429)
##
## Number of Fisher Scoring iterations: 5
```

#need to look into subset of high cholesterol and hypertension?

Modeling medication use with bp_uncontrolled_140_90 outcome

Model is limited to survey design subset of hypertension patients from 2013-2020. We selected medication variables concerning self-reported blood pressure medication use (yes/no), the number of antihypertensive medication pills taken per day, the use of vasodilators (yes/no), alpha and beta blockers (yes/no), the use of angiotensin receptor blockers, and the use of statins or other cholesterol medication for cholesterol control (yes/no). These variables were selected based on their prior association with blood pressure levels in existing clinical trials and research studies.

```
meds_binary <- svyglm(bp_uncontrolled_140_90 ~ bp_med_use + bp_med_n_pills + bp_med_vasod + bp_med_alph</pre>
```

Use of blood pressure medications, number of antihypertensive pills (1-4+), and angiotensin receptor blockers, were significantly associated with having uncontrolled blood pressure. The use of vasodilators, statins or other cholesterol medication, alpha and beta blockers, had non-statistically significant associations with having uncontrolled blood pressure.

Modeling medication use with bp_sys_mean outcome

We now model the continuous outcome of the average systolic blood pressure with the same medication covariates among those with hypertension from 2013-2020.

```
meds_continuous <- svyglm(bp_sys_mean ~ bp_med_use + bp_med_n_pills + bp_med_vasod + bp_med_alpha + bp_med_state + bp_med_stat
```

The use of blood pressure medications, number of antihypertensive pills (1), and angiotensin receptor blockers were significantly associated with mean systolic blood pressure levels in those with hypertension from 2013-2020. The use of (2-4+) antihypertensive pills per day, vasodilators, alpha and beta blockers, and statins and other cholesterol medications, had non-statistically significant associations with mean systolic blood pressure levels in those with hypertension from 2013-2020.

Nested Models

Level One Demographic Characteristics.

```
model_1 <- svyglm(bp_uncontrolled_140_90 ~ demo_age_cat + demo_race + demo_gender, #demographics
                 design = htsub2, family=quasibinomial())
summary(model_1)
##
## Call:
## svyglm(formula = bp_uncontrolled_140_90 ~ demo_age_cat + demo_race +
      demo_gender, design = htsub2, family = quasibinomial())
##
## Survey design:
## subset(htsub, svy_year == "2013-2014" | svy_year == "2015-2016" |
      svy_year == "2017-2020")
##
##
## Coefficients:
                              Estimate Std. Error t value Pr(>|t|)
##
## (Intercept)
                              -2.92319 0.09320 -31.364 < 2e-16 ***
## demo_age_cat45 to 64
                               1.41571
                                          0.07074 20.014 < 2e-16 ***
## demo_age_cat65 to 74
                               2.01652
                                          0.07848 25.696 < 2e-16 ***
## demo_age_cat75+
                               2.70181
                                          0.10722 25.200 < 2e-16 ***
## demo_raceNon-Hispanic Black 0.86309
                                          0.07301 11.822 1.1e-15 ***
## demo_raceNon-Hispanic Asian 0.26243
                                          0.07708
                                                   3.405 0.00136 **
## demo_raceHispanic
                                                    3.075 0.00350 **
                               0.23525
                                          0.07650
## demo_raceOther
                               0.27350
                                          0.17971
                                                    1.522 0.13474
## demo_genderWomen
                                          0.06224 -3.436 0.00124 **
                              -0.21386
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for quasibinomial family taken to be 0.9818051)
## Number of Fisher Scoring iterations: 5
```

Level Two Health-related Characteristics

```
##
## Call:
## svyglm(formula = bp_uncontrolled_140_90 ~ demo_age_cat + demo_race +
      demo_gender + cc_smoke + cc_bmi + cc_diabetes + cc_ckd +
##
      cc_cvd_any + chol_trig + chol_ldl + chol_hdl, design = htsub2,
      family = quasibinomial())
##
##
## Survey design:
## subset(htsub, svy_year == "2013-2014" | svy_year == "2015-2016" |
      svy_year == "2017-2020")
##
##
## Coefficients:
                             Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                           -4.2784289 0.2847878 -15.023 < 2e-16 ***
                           1.1528233 0.1252890
                                                9.201 5.47e-11 ***
## demo_age_cat45 to 64
## demo_age_cat65 to 74
                            1.7627753  0.1598648  11.027  4.28e-13 ***
                            ## demo_age_cat75+
## demo_raceNon-Hispanic Black 0.8670093 0.1173396
                                               7.389 1.02e-08 ***
## demo_raceNon-Hispanic Asian 0.3999404 0.1137010 3.517 0.001199 **
                       0.1559047 0.1091321 1.429 0.161743
## demo_raceHispanic
## demo_raceOther
                          0.0207119 0.2431672 0.085 0.932594
                         ## demo_genderWomen
                          0.0729305 0.1250743 0.583 0.563463
## cc smokeFormer
                           0.2004411 0.1685633 1.189 0.242179
## cc smokeCurrent
## cc_bmi25 to <30
                          -0.0215815 0.1116762 -0.193 0.847849
## cc_bmi30 to <35
                          0.2261152 0.1130356 2.000 0.053045 .
## cc_bmi35+
                          0.7023231 0.1345164
                                               5.221 7.63e-06 ***
                        0.3421597 0.1262108
## cc_diabetesYes
                                               2.711 0.010214 *
## cc_ckdYes
                          ## cc_cvd_anyYes
                          -0.0273710 0.1208875 -0.226 0.822158
## chol_trig
                            0.0017308 0.0003819
                                                4.533 6.21e-05 ***
## chol_ldl
                            0.0052070 0.0012622 4.125 0.000209 ***
## chol_hdl
                            0.0061692 0.0032180 1.917 0.063190 .
## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for quasibinomial family taken to be 0.9486429)
## Number of Fisher Scoring iterations: 5
```

Level Three Medication Use

svyglm(formula = bp_uncontrolled_140_90 ~ demo_age_cat + demo_race +

```
##
      demo_gender + cc_smoke + cc_bmi + cc_diabetes + cc_ckd +
##
      cc_cvd_any + chol_trig + chol_ldl + chol_hdl + bp_med_use +
##
      bp_med_n_pills + bp_med_vasod + bp_med_alpha + bp_med_beta +
      bp_med_angioten + chol_med_statin + chol_med_use, design = htsub2,
##
##
      family = quasibinomial())
##
## Survey design:
  subset(htsub, svy_year == "2013-2014" | svy_year == "2015-2016" |
##
      svy year == "2017-2020")
##
  Coefficients:
##
                               Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                             -4.2354978 0.2697846 -15.700 1.86e-14 ***
## demo_age_cat45 to 64
                             1.0690468 0.1288047
                                                   8.300 1.19e-08 ***
## demo_age_cat65 to 74
                             1.6111371   0.1594032   10.107   2.58e-10 ***
## demo_age_cat75+
                             2.2844522 0.1762558
                                                  12.961 1.35e-12 ***
## demo_raceNon-Hispanic Black   0.8434587   0.1182782
                                                  7.131 1.79e-07 ***
## demo_raceNon-Hispanic Asian 0.3873648 0.1161263
                                                  3.336 0.002659 **
## demo_raceHispanic
                             0.1973600 0.1073986
                                                   1.838 0.078030
## demo raceOther
                             0.0066420 0.2505540
                                                   0.027 0.979062
## demo_genderWomen
                            ## cc smokeFormer
                             0.0752089 0.1281457
                                                  0.587 0.562533
## cc_smokeCurrent
                            0.2019008 0.1687557
                                                   1.196 0.242761
## cc bmi25 to <30
                            -0.0631372 0.1149384 -0.549 0.587664
## cc bmi30 to <35
                            0.1786980 0.1134249
                                                  1.575 0.127718
## cc bmi35+
                            0.5917701 0.1431533
                                                   4.134 0.000351 ***
## cc_diabetesYes
                                                   2.434 0.022397 *
                             0.3255830 0.1337452
## cc_ckdYes
                             0.6201295 0.1265869
                                                   4.899 4.84e-05 ***
## cc_cvd_anyYes
                            ## chol_trig
                             0.0016698 0.0003928
                                                  4.252 0.000259 ***
## chol_ldl
                             0.0050071 0.0012360
                                                   4.051 0.000434 ***
## chol_hdl
                             0.0063284 0.0032239
                                                   1.963 0.060870
## bp_med_useYes
                            0.5344699 0.2079053
                                                  2.571 0.016488 *
## bp_med_n_pillsOne
                            -0.1874577 0.2794390 -0.671 0.508476
## bp_med_n_pillsTwo
                            -0.2084591 0.2981528
                                                  -0.699 0.490901
## bp_med_n_pillsThree
                            -0.4789832 0.3351850 -1.429 0.165378
## bp_med_n_pillsFour or more -0.4355646 0.3967429 -1.098 0.282731
## bp_med_vasodYes
                             0.0736384 0.3720228
                                                  0.198 0.844692
## bp_med_alphaYes
                             0.1738527 0.3837901
                                                   0.453 0.654464
## bp_med_betaYes
                             0.2530225 0.1310938
                                                   1.930 0.065016
## bp med angiotenYes
                             0.3140413 0.1492004
                                                   2.105 0.045525 *
## chol med statinYes
                             0.0018895 0.3706140
                                                   0.005 0.995973
## chol med useYes
                             ## ---
## Signif. codes: 0 '*** 0.001 '** 0.01 '* 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for quasibinomial family taken to be 0.941334)
##
## Number of Fisher Scoring iterations: 5
```

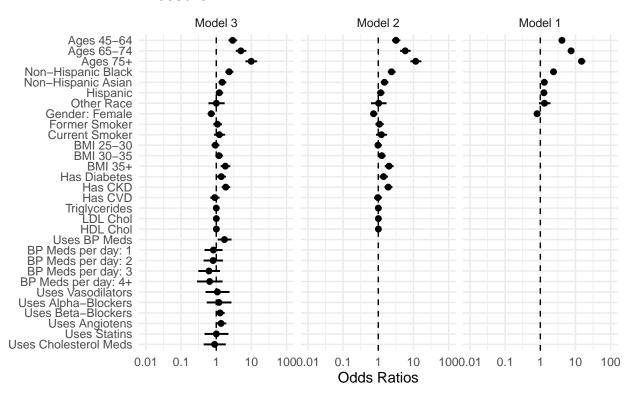
I added the nested models based on three levels. Should we also make figures based on the significant variables from these models? If these models look good, I will proceed making the table two - Harry

Figure 2: Forest plot of estimates from models 1-3

This plots estimates and 95% CI for odds ratios from models 1-3 and can be our figure 2 - Will

```
plot_models(model_1, model_2, model_3, transform = "exp", legend.title = "", show.p = T, grid = T, show
```

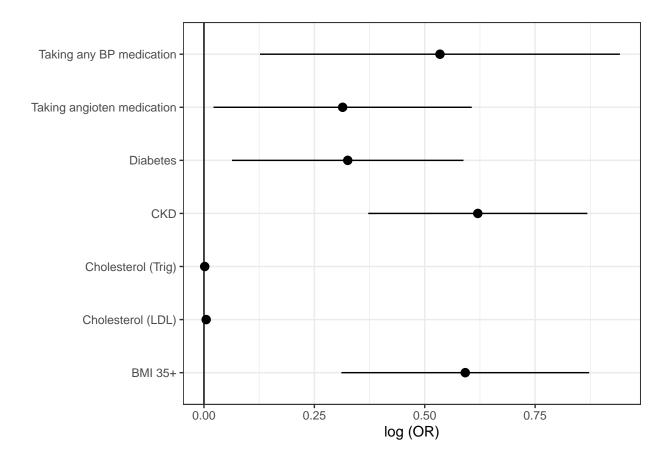
Odds Ratio Estimates for Uncontrolled Blood Pressure



Farizah's code for plotting forest plot of model 3, I don't think it's outputting correctly:

```
#age
reg_age <- matrix(c(1.0690468, 1.6111371, 2.2844522, 0.1288047, 0.1594032, 0.1762558), byrow=FALSE, nco
colnames(reg_age) <- c("log_or", "se")

reg_age <- reg_age %>% as_tibble()
reg_age$age <- c("45 to 64", "65 to 74", "75+" )
reg_age$195 <- reg_age$log_or-1.96*reg_age$se
reg_age$u95 <- reg_age$log_or+1.96*reg_age$se
age_plot <- ggplot(reg_age, aes(x = log_or, y = age)) + geom_point() + geom_pointrange(aes(xmin = 195, section = 195, section
```



```
library(ggpubr)
ggarrange(age_plot, race_plot, health_plot, ncol =3, nrow=1, labels = c("Age (ref = 18-44)", "Race (ref
```

