Lab 5 BIS 505b

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- · Goal of Lab 5
- · Analysis Data Set
- Research Questions
- · Multiple Linear Regression
 - Binary Predictor Variable
 - Categorical Predictor Variable
 - The Unadjusted Model
 - The Adjusted Model
 - \circ Overall F-Test
 - \circ Partial F-Test
 - Interactions
- · Automated Variable Selection

Goal of Lab 5

In **Lab 5**, we will **(1)** work with categorical predictors, **(2)** create multiple linear regression models, **(3)** explore interactions between variables and **(4)** present automated variable selection methods.

Analysis Data Set

In this lab, we will analyze a subset of data from the **National Health and Nutrition Examination Survey** (NHANES) (n = 1430) nhanes.csv imported as the data frame nhanes in code chunk 3 above). The **Data Key** is provided below:

Variable Name	Definition
fastgluc	Fasting glucose level (mg/dL) (Our Response)
	88888 = Missing
age	Age at time of survey
sex	Sex
	0 = Male
	1 = Female
oralmed	Oral diabetes medication use
	0 = No
	1 = Yes
race	Race/Ethnicity

Variable Name	Definition
	1 = White
	2 = Black
	3 = Mexican-American
	4 = Other

After reviewing the Data Key, we see that missing values of fastgluc are coded as 88888. Begin by re-coding this numerical value of 88888 as NA in **R**.

```
# Re-code a `fastgluc` value of 88888 as NA
nhanes$fastgluc[nhanes$fastgluc == 88888] <- NA
# Checking missing values
summary(nhanes)</pre>
```

```
##
       fastgluc
                                                          oralmed
                          age
                                           sex
##
    Min.
           : 42.2
                    Min.
                            :30.00
                                             :0.0000
                                                       Min.
                                                               :0.0000
                                     Min.
    1st Qu.:108.1
                    1st Qu.:56.00
                                     1st Qu.:0.0000
                                                       1st Qu.:0.0000
##
    Median :149.2
##
                    Median :66.00
                                     Median :1.0000
                                                       Median :0.0000
##
   Mean
           :177.6
                            :64.41
                                             :0.5629
                                                              :0.4898
                    Mean
                                     Mean
                                                       Mean
    3rd Qu.:232.9
##
                    3rd Qu.:74.00
                                     3rd Qu.:1.0000
                                                       3rd Qu.:1.0000
           :594.2
                            :90.00
                                             :1.0000
                                                       Max.
                                                              :1.0000
##
    Max.
                    Max.
                                     Max.
                                                       NA's
    NA's
           :264
                                                              :5
##
##
         race
##
    Min.
           :1.000
##
    1st Qu.:1.000
    Median :2.000
##
##
    Mean
           :1.945
    3rd Qu.:3.000
##
##
           :4.000
   Max.
##
```

Missing Values:

Note: fastgluc contains 264 **missing values**. These observations will not be used to fit any of the models. In addition, individuals with a missing value for any of the explanatory variables included in a specific model will not be used to fit that model. This is an important consideration when thinking about which explanatory variables to include. For example, including a predictor that has a large proportion of missing values in the model will consequently exclude anyone who is missing a value for that variable from the regression model. This is called a **complete case analysis**. That is, only records with complete data on all variables included in a model, $y \sim x1 + x2 + x3$, (complete data for y, x1, x2 and x3) will be analyzed.

• Creating Factor Variables:

There are several **categorical variables** in this data set (sex , oralmed and race). Use the str() function to see how each variable is stored in **R** (numeric, integer, factor, character).

```
str(nhanes)
```

```
## 'data.frame': 1430 obs. of 5 variables:
## $ fastgluc: num 271.7 83.3 107.3 109.4 175.5 ...
## $ age : int 48 82 66 80 72 78 31 83 49 63 ...
## $ sex : int 0 1 1 1 0 1 0 1 1 1 ...
## $ oralmed : int 0 0 0 1 0 0 1 1 1 1 ...
## $ race : int 3 1 3 1 2 2 2 1 2 1 ...
```

Notice that the variables for race, oral diabetes medication use and sex are coded as integers, but should be coded as **factor variables**. We can use either the <code>mutate()</code> function in the <code>dplyr</code> package or the <code>factor()</code> function directly to redefine these variables as factors based on the Data Key.

```
## 'data.frame':
                  1430 obs. of 8 variables:
                  : num 271.7 83.3 107.3 109.4 175.5 ...
## $ fastgluc
                  : int 48 82 66 80 72 78 31 83 49 63 ...
## $ age
## $ sex
                  : int 0111010111...
## $ oralmed
                 : int 0001001111...
## $ race
                  : int 3 1 3 1 2 2 2 1 2 1 ...
##
  $ sex_factor : Factor w/ 2 levels "Male", "Female": 1 2 2 2 1 2 1 2 2 2 ...
   $ oralmed_factor: Factor w/ 2 levels "No", "Yes": 1 1 1 2 1 1 2 2 2 2 ...
##
## $ race factor
                 : Factor w/ 4 levels "White", "Black", ...: 3 1 3 1 2 2 2 1 2 1 ...
```

Note: When we include **factor variables** in our regression model, the **first level** specified in the factor() function is used as the **reference level**. For example, given the way sex_factor is defined above (levels = c(0, 1)), a linear regression model that includes sex_factor will assume *male* is the reference category. The effect of sex_factor then compares females (1) to males (0) (reference). Re-ordering the levels= argument of the factor() function allows us to specify the desired reference level by listing it first. For example, to report the average difference in fasting glucose level in males (0) vs. females (1) (reference), define sex_factor as sex_factor and sex_factor as sex_factor and sex_factor as sex_factor as sex_factor and sex_factor and s

Research Questions

We are interested in studying characteristics associated with **fasting glucose levels** fastgluc (our response variable, y). The **research questions** include:

- 1. Is there evidence of an association between age and fasting glucose levels, after controlling for sex, oral diabetes medication use and race/ethnicity?
- 2. In the multiple regression model, is race/ethnicity an important predictor of fasting glucose level?
- 3. Does the effect of age on fasting glucose levels differ by oral diabetes medication use?

Multiple Linear Regression

Multiple linear regression is used to describe the relationship between a quantitative response variable y and more than one explanatory variable x_1, \ldots, x_k . The *population regression model*,

 $\mu_{y|x}=lpha+eta_1x_1+\ldots+eta_kx_k$, is estimated using the method of **least squares**, giving a *fitted model*, $\hat{y}=a+b_1x_1+\ldots+b_kx_k$.

- The fitted model is used to **predict** or **estimate** the expected value of y for given values of x_1, \ldots, x_k by plugging these values of x into the fitted equation, $\hat{y} = a + b_1 x_1 + \ldots + b_k x_k$.
- The estimated slope b_j is used to **describe the association** between x_j and y, controlling for or holding all other explanatory variables constant.

A test of the slope, β_j (i.e., $H_0:\beta_j=0$ vs. $H_1:\beta_j\neq 0$) is used to determine if an association exists between x_j and y, holding all other explanatory variables constant. This test is performed using the t-statistic, $t=\frac{b_j}{s_{b_j}}$, which is compared to a t-distribution with n-p degrees of freedom. Note that p is equal to the number of regression parameters estimated in the model (k slopes + 1 intercept), so p=k+1.

The lm() function in **R** is used to estimate the regression coefficients (i.e., the intercept and slope parameter(s)) of the linear model. The result of the lm() function is usually saved as an object (e.g., regobject) and the summary() function is applied to that object (summary(regobject)) to output detailed results.

1m() Function Arguments	Option Definition
formula=	analysis_variable ~ predictor_variable1 + predictor_variable2
data=	Data frame containing sample data

A $100(1-\alpha)\%$ confidence interval for the parameter β_j is equal to $b_j \pm t_{1-\alpha/2;n-p} \ s_{b_j}$. The confint(regobject) function is used to return 95% confidence intervals for the model parameters. By default, 95% confidence intervals (level=0.95) are produced.

Finally, the fitted model can be used to estimate or predict a value of y for given values of x using the <code>predict()</code> function. A new data frame must be created and specified in the <code>newdata=</code> argument of the <code>predict()</code> function that contains the values of x used to predict values of y.

Binary Predictor Variable

So far, have only considered **quantitative predictor variables**. **Categorical variables** can be included as predictors in a regression model through the use of numeric 0/1 **dummy** or **indicator variables** z_j , where the reference level of the variable equals o.

When we include a **factor variable** in a regression model, **R** will automatically create the dummy variable(s) necessary to represent that categorical variable. The contrasts() function returns the dummy variable coding that **R** uses to represent a factor variable. For example, sex factor is a dummy variable (z_1) that equals 1 for

females and 0 for males (the reference category).

```
contrasts(nhanes$sex_factor)
```

```
## Female
## Female 1
```

• In an **unadjusted model** that contains sex_factor , $\hat{y}=a+b_1z_1$, the estimated slope of sex_factor b_1 equals the estimated *difference* in mean fasting glucose in females vs. males (ref) $(\bar{y}_f - \bar{y}_m)$. Including additional variables in the model will give *adjusted* differences in mean fasting glucose.

```
# SLR model including sex_factor
mod.sex <- lm(fastgluc ~ sex_factor, data = nhanes)
summary(mod.sex)</pre>
```

```
##
## Call:
## lm(formula = fastgluc ~ sex_factor, data = nhanes)
##
## Residuals:
      Min
##
               1Q Median
                               3Q
                                      Max
## -136.27 -69.36 -28.35
                            56.20 415.13
##
## Coefficients:
##
                   Estimate Std. Error t value
                                                          Pr(>|t|)
                                 3.934 44.672 <0.00000000000000000
## (Intercept)
                    175.724
## sex factorFemale
                      3.348
                                 5.260
                                         0.637
                                                             0.525
##
## Residual standard error: 89.18 on 1164 degrees of freedom
     (264 observations deleted due to missingness)
## Multiple R-squared: 0.0003479, Adjusted R-squared: -0.0005109
## F-statistic: 0.4052 on 1 and 1164 DF, p-value: 0.5246
```

```
## sex fastgluc.mean
## 1 Male 175.7237
## 2 Female 179.0721
```

• The **fitted model** is given by the equation, $\hat{y}=175.72+3.35$ Female. To make the interpretation of the fitted model easier, instead of using the variable name <code>sex_factor</code> in the written fitted model, I specified the level of the dummy variable that is being compared to the reference level (i.e., female vs. male).

- The **estimated intercept** a=175.72 is equal to the mean fastgluc when $z_1=0$ (i.e., the mean fasting glucose in the reference category (males)).
- The **estimated slope** of sex_{factor} $b_1 = 3.35$ is equal to the *difference* in mean fastgluc in females (179.07) minus males (175.72).
- A significance test of the slope $(H_0: \beta_1 = 0 \text{ vs. } \beta_1 \neq 0)$ reports a t-statistic t =0.64, which is compared to a t-distribution with 1164 degrees of freedom. This test does not support a significant difference in the mean fasting glucose in males vs. females (p-value = 0.525).

You can specify the reference category when creating a factor variable by listing that category as the **first level** in the levels= argument of the factor() function. For example, sex_factorv2 will set *female* as the reference category. Notice that the dummy variable for sex_factorv2 equals 1 for males and 0 for females.

```
## Male
## Female 0
## Male 1
```

The estimated slope of sex_factorv2 is equal to the estimated difference in mean fasting glucose in males vs. females (ref) $(\bar{y}_m - \bar{y}_f)$.

```
# SLR model including sex_factorv2
mod.sexv2 <- lm(fastgluc ~ sex_factorv2, data = nhanes)
summary(mod.sexv2)</pre>
```

```
##
## Call:
## lm(formula = fastgluc ~ sex factorv2, data = nhanes)
##
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -136.27 -69.36 -28.35
                            56.20 415.13
##
## Coefficients:
##
                   Estimate Std. Error t value
                                                          Pr(>|t|)
## (Intercept)
                    179.072
                                 3.493 51.271 <0.00000000000000000
## sex_factorv2Male -3.348
                                 5.260 -0.637
                                                             0.525
##
## Residual standard error: 89.18 on 1164 degrees of freedom
##
     (264 observations deleted due to missingness)
## Multiple R-squared: 0.0003479, Adjusted R-squared: -0.0005109
## F-statistic: 0.4052 on 1 and 1164 DF, p-value: 0.5246
```

We can also use the relevel() function to change the reference category (ref=) of an existing factor variable:

```
# Female will be the reference category (=0) of sex_factorv3
nhanes$sex_factorv3 <- relevel(nhanes$sex_factor, ref = "Female")
contrasts(nhanes$sex_factorv3)</pre>
```

```
## Male
## Female 0
## Male 1
```

Categorical Predictor Variable

Categorical variables with C levels are represented by a set of C-1 dummy variables. Again, when using factor versions of our categorical variables, $\mathbf R$ automatically creates the dummy variables needed to represent the categorical variable in a regression model. Be sure **not** to use ordered = TRUE when creating factor variables for inclusion in a regression model.

race_factor contains **4** levels (White, Black, Mexican-American, and Other) and must be represented by **3** dummy variables (z_1 , z_2 and z_3). When we created race_factor at the beginning of this Lab, White was specified as the first level (levels=c(1,2,3,4) corresponding to

labels=c("White", "Black", "Mexican-American", "Other")), thus "White" will be the reference category. All dummy variables will equal 0 for the reference level of the categorical variable. Below, we see the 3 dummy variables that describe race:

```
contrasts(nhanes$race_factor)
```

```
## Black Mexican-American Other

## White 0 0 0

## Black 1 0 0

## Mexican-American 0 1 0

## Other 0 0 1
```

- 1. z_1 equals 1 when race_factor == "Black" and equals 0 otherwise
- 2. z_2 equals 1 when race_factor == "Mexican-American" and equals 0 otherwise
- 3. z_3 equals 1 when race_factor == "Other" and equals 0 otherwise
- In an **unadjusted model**, $\hat{y}=a+b_1z_1+b_2z_2+b_3z_3$, the estimated slope of the first dummy variable b_1 equals the estimated difference in mean fasting glucose in Blacks vs. Whites (ref) $(\bar{y}_b-\bar{y}_w)$. The estimated slope of the second dummy variable b_2 equals the estimated difference in mean fasting glucose in Mexican-Americans vs. Whites (ref) $(\bar{y}_m-\bar{y}_w)$. The estimated slope of the third dummy variable b_3 equals the estimated difference in mean fasting glucose in Others vs. Whites (ref) $(\bar{y}_o-\bar{y}_w)$. Including additional variables in the model will give *adjusted* differences in mean fasting glucose.

```
# Model including race_factor
mod.race <- lm(fastgluc ~ race_factor, data = nhanes)
summary(mod.race)</pre>
```

```
##
## Call:
## lm(formula = fastgluc ~ race_factor, data = nhanes)
##
## Residuals:
##
      Min
                1Q Median
                                3Q
                                       Max
##
  -139.60 -67.91 -28.15
                             54.01 405.30
##
## Coefficients:
##
                               Estimate Std. Error t value
                                                                        Pr(>|t|)
                                             4.220 39.860 < 0.00000000000000002
## (Intercept)
                                168.222
## race factorBlack
                                 20.677
                                             6.547
                                                     3.158
                                                                         0.00163
## race factorMexican-American
                                 12.454
                                             6.233
                                                     1.998
                                                                         0.04595
## race factorOther
                                            16.029 -0.479
                                 -7.674
                                                                         0.63221
##
## Residual standard error: 88.83 on 1162 degrees of freedom
     (264 observations deleted due to missingness)
##
## Multiple R-squared: 0.009968,
                                    Adjusted R-squared: 0.007412
                  3.9 on 3 and 1162 DF, p-value: 0.008699
## F-statistic:
```

- The **fitted model** is given by the equation, $\hat{y}=$ 168.22 + 20.68 Black + 12.45 Mexican-American 7.67 Other.
- The average fasting glucose in Blacks is $b_1=20.68$ [95% CI (7.83, 33.52)] units higher than in Whites. A **significance test** of β_1 shows that there is a significant difference in the mean fasting glucose in Blacks vs. Whites (p-value = 0.002).
- The average fasting glucose in Mexican-Americans is $b_2 = 12.45$ [95% CI (0.22, 24.68)] units higher than in Whites. A **significance test** of β_2 shows that there is a significant difference in the mean fasting glucose in Mexican-Americans vs. Whites (p-value = 0.046).
- The average fasting glucose in Other races/ethnicities is $b_3 = -7.67$ [95% CI (-39.12, 23.77)] units different (lower) than in Whites. A **significance test** of β_3 shows that there is not a significant difference in the mean fasting glucose in Other races/ethnicities vs. Whites (p-value = 0.632).

The Unadjusted Model

The first research question asks if there is an association between age (age) and fasting glucose levels (fastgluc), after controlling for sex (sex_factor), oral diabetes medication use (oralmed_factor) and race/ethnicity (race_factor). Let's begin by estimating the unadjusted effect of age on fastgluc using a simple linear regression model. We call this the unadjusted effect because no other variables are being controlled for or adjusted for in the regression model.

• Unadjusted model: $\mu_{y|x} = lpha + eta_1$ Age

We fit the model using the lm() function and save the fitted model to the object mod.age. We output a summary of the results using summary(mod.age) and the 95% CIs of the model parameters using summary(mod.age).

```
# SLR model including age
mod.age <- lm(fastgluc ~ age, data = nhanes)
# Output results of fitted model
summary(mod.age)</pre>
```

```
##
## Call:
## lm(formula = fastgluc ~ age, data = nhanes)
##
## Residuals:
##
      Min
               1Q Median
                               3Q
                                      Max
## -162.13 -67.00 -26.35 52.64 414.70
##
## Coefficients:
                                                      Pr(>|t|)
##
              Estimate Std. Error t value
## (Intercept) 228.3632
                        12.7491 17.912 < 0.000000000000000002
               -0.8011
                           0.1970 -4.067
                                                     0.0000508
## age
##
## Residual standard error: 88.57 on 1164 degrees of freedom
    (264 observations deleted due to missingness)
## Multiple R-squared: 0.01401,
                                 Adjusted R-squared: 0.01316
## F-statistic: 16.54 on 1 and 1164 DF, p-value: 0.00005081
```

```
# Confidence intervals for model parameters (intercept and slope)
confint(mod.age)
```

```
## 2.5 % 97.5 %
## (Intercept) 203.349349 253.3769563
## age -1.187535 -0.4146312
```

- The **fitted model** is given by the equation, $\hat{y} = 228.36 0.8$ Age.
- The **estimated slope** of age $b_1 = -0.8$ [95% CI (-1.19, -0.41)] indicates that a 1-unit increase in age is associated with a -0.8-unit average change (a decrease) in fasting glucose.
- A significance test of the slope $(H_0: \beta_1 = 0 \text{ vs. } \beta_1 \neq 0)$ reports a t-statistic t =-4.07, which is compared to a t-distribution with 1164 degrees of freedom. This yields a highly significant p-value <.001.
- The **R-squared** of this model is low at 0.014, indicating that $_{\text{age}}$ only explains 1.4% of the total variability in fasting glucose. The variability about the regression line $\sigma_{y|x}$ is estimated by the "**residual standard error**" in the output above and is equal to $s_{y|x} = 88.57$.

The Adjusted Model

To **control** or **adjust for** sex, oral diabetes medication use and race/ethnicity when estimating the effect of age on fasting glucose, we will fit a **multiple linear regression model** that additionally includes the variables sex_factor, oralmed_factor and race_factor.

• Adjusted model: $\mu_{y|x}=\alpha+\beta_1$ Age $+\beta_2$ Female $+\beta_3$ MedUse $+\beta_4$ Black $+\beta_5$ Mexican-American $+\beta_6$ Other

```
# MLR model including age, sex_factor, oralmed_factor, race_factor
mod.mlr <- lm(fastgluc ~ age + sex_factor + oralmed_factor + race_factor, data = nhanes)
summary(mod.mlr)</pre>
```

```
##
## Call:
## lm(formula = fastgluc ~ age + sex_factor + oralmed_factor + race_factor,
##
       data = nhanes)
##
## Residuals:
##
       Min 1Q Median 3Q
                                        Max
## -153.65 -65.29 -27.40 49.84 399.79
##
## Coefficients:
##
                                Estimate Std. Error t value
                                                                          Pr(>|t|)
## (Intercept)
                                211.0088 15.1442 13.933 < 0.00000000000000002
## age
                                 -0.7065
                                             0.2061 -3.428
                                                                          0.000629
## sex_factorFemale
                                  2.0178
                                            5.2643 0.383
                                                                         0.701578
## oralmed_factorYes 9.6060 5.2393 1.833
## race_factorBlack 14.8757 6.8005 2.187
## race_factorMexican-American 6.0303 6.4286 0.938
                                                                          0.066994
                                                                         0.028910
                                                                         0.348418
                       -15.4705
## race factorOther
                                            16.1097 -0.960
                                                                          0.337094
##
## Residual standard error: 88.45 on 1156 degrees of freedom
    (267 observations deleted due to missingness)
## Multiple R-squared: 0.02231, Adjusted R-squared: 0.01723
## F-statistic: 4.396 on 6 and 1156 DF, p-value: 0.0002109
```

confint(mod.mlr)

```
## 2.5 % 97.5 %

## (Intercept) 181.2955887 240.7219343

## age -1.1108063 -0.3021315

## sex_factorFemale -8.3109948 12.3464959

## oralmed_factorYes -0.6736602 19.8856553

## race_factorBlack 1.5330274 28.2184219

## race_factorMexican-American -6.5827490 18.6433961

## race_factorOther -47.0779068 16.1369944
```

- The **fitted model** is given by the equation, $\hat{y}=211.01-0.71$ Age +2.02 Female +9.61 MedUse +14.88 Black +6.03 Mexican-American -15.47 Other.
- · Sex, medication use, and race-adjusted effect of age on fasting glucose:
 - The **estimated slope** of age $b_1 = -0.71$ [95% CI (-1.11, -0.3)] in the multiple linear regression model indicates that a 1-unit increase in age is associated with a -0.71-unit average change (a decrease) in fasting glucose, controlling for sex, oral diabetes medication use and race.
 - A significance test of the slope $(H_0: \beta_1 = 0 \text{ vs. } \beta_1 \neq 0)$ shows a highly significant association between age and fasting glucose when controlling for sex, oral diabetes medication use and race (p-value <.001).

- Age, medication use, and race-adjusted effect of sex on fasting glucose:
 - \circ The **estimated slope** of sex_factor $b_2=2.02$ [95% CI (-8.31, 12.35)] estimates the average difference fasting glucose in females vs. males (reference), controlling for age, oral diabetes medication use and race. The adjusted average fasting glucose level is 2.02-units higher in females than in males.
 - A significance test of the slope $(H_0: \beta_2 = 0 \text{ vs. } \beta_2 \neq 0)$ shows there is not a statistically significant difference in average fasting glucose in females and males when controlling for age, oral diabetes medication use and race (p-value = 0.702). Notice that the 95% confidence interval for β_2 supports this conclusion since it includes 0 (i.e., the value of β_2 hypothesized under H_0).
 - Since sex is not a significant predictor in the presence of age, oral diabetes medication use and race, we may want to remove this variable from the regression model. However, if **confounding** is a concern, we can retain the variable regardless of statistical significance.
- The **Adjusted R-squared** (R_a^2) of this model remains low at 0.017. The **residual standard error** is equal to $s_{y|x}$ = 88.45, and is only slightly smaller than the estimate from the unadjusted model.

Exercise: Interpret the effect of oral diabetes medication use in the model above.		
► Answer:		
Exercise: Interpret the effect of race/ethnicity in the model above.		
▶ Answer:		

Overall F-Test

The **Overall F-Test** tests whether the explanatory variables collectively have an effect on the response variable. Under H_0 , $\beta_1=\beta_2=\ldots=\beta_k=0$. Under H_1 , at least one $\beta_j\neq 0$ for $j=1,\ldots,k$. The **F-test statistic** is equal to the ratio of the variability explained by the *model* (MSM) to the mean squared *error* (MSE), giving $F=\frac{MSM}{MSE}$. The **F-test statistic** is compared to an F-distribution with *numerator degrees of freedom* k and denominator degrees of freedom k.

• Option 1: The overall F-test is presented in the last line of the summary() of model results.

```
# Overall F-test on last line of output
summary(mod.mlr)
```

```
##
## Call:
## lm(formula = fastgluc ~ age + sex_factor + oralmed_factor + race_factor,
##
       data = nhanes)
##
## Residuals:
                1Q Median
##
       Min
                                3Q
                                       Max
##
  -153.65 -65.29 -27.40
                             49.84 399.79
##
## Coefficients:
##
                               Estimate Std. Error t value
                                                                        Pr(>|t|)
## (Intercept)
                               211.0088
                                            15.1442 13.933 < 0.000000000000000002
## age
                                 -0.7065
                                             0.2061
                                                    -3.428
                                                                        0.000629
## sex_factorFemale
                                             5.2643
                                                                        0.701578
                                 2.0178
                                                      0.383
## oralmed factorYes
                                 9.6060
                                             5.2393
                                                     1.833
                                                                        0.066994
## race_factorBlack
                                14.8757
                                             6.8005
                                                     2.187
                                                                        0.028910
                                             6.4286
## race factorMexican-American
                                 6.0303
                                                      0.938
                                                                        0.348418
## race_factorOther
                               -15.4705
                                            16.1097 -0.960
                                                                        0.337094
##
## Residual standard error: 88.45 on 1156 degrees of freedom
     (267 observations deleted due to missingness)
## Multiple R-squared: 0.02231,
                                    Adjusted R-squared: 0.01723
## F-statistic: 4.396 on 6 and 1156 DF, p-value: 0.0002109
```

The **overall F-test** F-statistic, F = 4.396, is compared to an F-distribution with 6 and 1156 degrees of freedom, giving a p-value <.001. There is evidence to conclude that a significant association exists between fasting glucose level and at least one explanatory variable in the MLR model.

• Option 2: We can request the ANOVA table of a model using the <code>anova()</code> function on the model object (e.g., <code>mod.mlr</code>). Note that the ANOVA table has the contribution to the sum of squares broken down by predictor ("Sequential SS" or "Type I SS") and does **not** provide the overall F-statistic. To find the Model SS (SSM), add the sum of squares for all predictors. To find the MSM, divide SSM by the number of independent variables, k, which is the number of parameters tested under H_0 (i.e., 6 in this case; recall that race is represented by 3 dummy variables).

```
# ANOVA table anova(mod.mlr)
```

```
## Analysis of Variance Table
##
## Response: fastgluc
##
                    Df Sum Sq Mean Sq F value
                                                   Pr(>F)
## age
                       129225
                               129225 16.5177 0.00005146
                     1
## sex factor
                     1
                          1574
                                  1574 0.2012
                                                  0.65382
## oralmed_factor
                     1
                         22324
                                 22324 2.8535
                                                  0.09145
## race factor
                     3
                         53238
                                 17746
                                       2.2683
                                                  0.07898
## Residuals
                  1156 9043883
                                  7823
```

The **overall F-test** F-statistic, F = [(129225 + 1574 + 22324 + 53238)/(1 + 1 + 1 + 3)]/7823 = 4.396 agrees with the F-statistic reported in the summary() output.

- **Option 3:** Finally, we can perform the overall F-test using the anova() function to compare two **nested models** using the syntax anova(reducedmodel, fullmodel).
 - \circ Full model: $\mu_{y|x}=lpha+eta_1$ Age $+eta_2$ Female $+eta_3$ MedUse $+eta_4$ Black $+eta_5$ Mexican-American $+eta_6$ Other
 - \circ Reduced model (i.e., model under H_0): $\mu_{y|x}=lpha$

Under the reduced model, H_0 is assumed to be true (i.e., $\beta_1=\beta_2=\ldots=\beta_6=0$). When performing the overall F-test, the reduced model is also known as the **null model** since it contains only the intercept term and no explanatory variables. The **R** syntax for fitting a model that contains only the intercept is <code>yvariable ~ 1</code>, where <code>1</code> represents the intercept.

Note: Recall that each regression model only includes records with **complete data** on all variables included in that model (**complete case analysis**). Since there are some individuals with missing values for <code>oralmed_factor</code>, the sample size used to fit <code>mod.full</code> is <code>smaller</code> than the sample size used to fit <code>mod.null</code>. To compare the same subset of observations under both the full and reduced models, we must specify that the analysis data set used to fit <code>mod.full</code> should also be used to fit <code>mod.null</code>. The "complete case" data frame used to fit <code>mod.full</code> is <code>data = mod.full\$model</code> and is specified as the data source for <code>mod.null</code>:

```
# Full model
mod.full <- lm(fastgluc ~ age + sex_factor + oralmed_factor + race_factor, data = nhanes)

# Reduced model (null model) fit using the same observations used to fit full model
mod.null <- lm(fastgluc ~ 1, data = mod.full$model) # note **data=** here!

# F-test comparing full and reduced models
anova(mod.null, mod.full)</pre>
```

```
## Analysis of Variance Table
##
## Model 1: fastgluc ~ 1
## Model 2: fastgluc ~ age + sex_factor + oralmed_factor + race_factor
## Res.Df RSS Df Sum of Sq F Pr(>F)
## 1 1162 9250243
## 2 1156 9043883 6 206360 4.3962 0.0002109
```

The **overall F-test** F-statistic, F = [206360/6]/[9043883/1156] = 4.396 agrees with the F-statistic reported in the summary() output.

Partial F-Test

The **Partial F-Test** simultaneously tests the significance of a group or set of parameters. This test is commonly used to test the effect of categorical variables that are naturally made up of more than one dummy variable. For example, to test the significance of **race/ethnicity** in the adjusted model, we would test:

$$H_0:eta_4=eta_5=eta_6=0$$
 vs. $H_1:eta_4$, eta_5 , eta_6 not all 0.

Here, we are comparing two nested models

• Full model: $\mu_{y|x}=\alpha+\beta_1$ Age $+\beta_2$ Female $+\beta_3$ MedUse $+\beta_4$ Black $+\beta_5$ Mexican-American $+\beta_6$ Other

• Reduced model (i.e., model under H_0 , without race_factor): $\mu_{y|x}=\alpha+\beta_1$ Age $+\beta_2$ Female $+\beta_3$ MedUse

```
The partial F-test statistic is equal to F_0 = rac{\frac{SSM(F) - SSM(R)}{Number parameters tested under H_0}}{\frac{SSE(F)}{df_0(F)}}
```

The F-statistic is compared to an F-distribution with *numerator degrees of freedom* equal to the number of parameters tested under H_0 and *denominator degrees of freedom* n-p.

• **Option 1:** We can perform the partial F-test using the anova() function to compare two **nested models** using the syntax anova(reducedmodel, fullmodel).

As above, we need to be sure that we are fitting both the full and reduced models using the same data set. Thus, when fitting the *reduced model*, use the observations that were included in the full model by specifying data=mod.full\$model.

```
# Full model
mod.full <- lm(fastgluc ~ age + sex_factor + oralmed_factor + race_factor, data = nhanes)

# Reduced model (under H0, does not include race_factor)
# Fit using the same observations included in the full model
mod.red <- lm(fastgluc ~ age + sex_factor + oralmed_factor, data = mod.full$model)

# F-test comparing full and reduced models
anova(mod.red, mod.full)</pre>
```

```
## Analysis of Variance Table
##
## Model 1: fastgluc ~ age + sex_factor + oralmed_factor
## Model 2: fastgluc ~ age + sex_factor + oralmed_factor + race_factor
## Res.Df RSS Df Sum of Sq F Pr(>F)
## 1 1159 9097120
## 2 1156 9043883 3 53238 2.2683 0.07898
```

The **partial F-test** F-statistic, F = [53238/3]/[9043883/1156] = 2.268 is compared to an F-distribution with 3 and 1156 degrees of freedom. The effect of race is not statistically significant in the full model (p = 0.079), thus we cannot reject H_0 . Although we did see that β_4 (coefficient for dummy variable of Black vs. White) was significantly different from 0 in the individual t-tests of the slopes in mod.mlr, perhaps the effect was not strong enough to outweigh the lack of statistical significance seen in the other two dummy variables that make up $race_factor$.

• Option 2: Option 1 is a more flexible option for carrying out a partial F-test and be used to simultaneously test many different slope parameters involving different variables (e.g., simultaneously test the effect of age and sex_factor by testing $H_0: \beta_1=\beta_2=0$). However, if the goal of the partial F-test is to test C-1 dummy variables of a single C-level categorical variable, then we can use the Anova() function in the car package. The Anova() function applied to a model object (e.g., mod.full) returns individual F-tests for each variable in the model. A reduced model does not need to be explicitly specified in Option 2.

To test the effect of <code>race_factor</code> in a model containing age, sex and oral diabetes medication use, $\mu_{y|x}=\alpha+\beta_1$ Age $+\beta_2$ Female $+\beta_3$ MedUse $+\beta_4$ Black $+\beta_5$ Mexican-American $+\beta_6$ Other, we would test $H_0: \beta_4=\beta_5=\beta_6=0$ vs. $H_1: \beta_4, \beta_5, \beta_6$ not all 0.

```
mod.full <- lm(fastgluc ~ age + sex_factor + oralmed_factor + race_factor, data = nhanes)
# Anova() function in the "car" package
Anova(mod.full)</pre>
```

```
## Anova Table (Type II tests)
##
## Response: fastgluc
##
                  Sum Sq
                           Df F value
                                        Pr(>F)
## age
                   91939
                            1 11.7518 0.0006292
                            1 0.1469 0.7015782
## sex factor
                    1149
## oralmed_factor
                   26298
                            1 3.3615 0.0669941
## race factor
                            3 2.2683 0.0789751
                   53238
## Residuals
                 9043883 1156
```

Based on the output above, the **partial F-test** of all dummy variables that make up <code>race_factor</code> has an F-statistic = [53238/3]/[9043883/1156] = 2.268, which is compared to an F-distribution with 3 and 1156 degrees of freedom. As we saw above, the effect of race is not statistically significant in the presence of the other variables (p = 0.079), thus we cannot reject H_0 .

Interactions

Next, we would like to determine if the effect of age on fasting glucose level differs by oral diabetes medication use. Answering this question involves examining the **interaction** between age and oralmed_factor. The model that includes the interaction age*oralmed_factor also includes the main effects of age and oralmed_factor,

- Interaction model: $\mu_{y|x}=lpha+eta_1$ Age $+eta_2$ MedUse $+eta_3$ Age imes MedUse
- Model in those who use oral diabetes medication (MedUse = 1): $\mu_{y|x}=(lpha+eta_2)+(eta_1+eta_3)$ Age
- Model in those who do not use oral diabetes medication (reference) (MedUse = 0): $\mu_{y|x}=lpha+eta_1$ Age

Thus, a test of β_3 will determine if there is a significant interaction between age and oral diabetes medication use. If a significant interaction exists, then we can estimate the slope of age (i.e., the effect of age on fasting glucose level) separately in the two medication use categories. Note that for simplicity, I am examining this interaction in a model that does not control for sex or race. However, you could easily also include these variables in the model to examine the interaction while controlling for sex and race.

```
# Interaction model of age*oralmed_factor
mod.intx <- lm(fastgluc ~ age + oralmed_factor + age*oralmed_factor, data = nhanes)
summary(mod.intx)</pre>
```

```
##
## Call:
## lm(formula = fastgluc ~ age + oralmed_factor + age * oralmed_factor,
##
       data = nhanes)
##
## Residuals:
                1Q Median
##
      Min
                                3Q
                                       Max
##
  -146.12 -66.85 -26.22
                            51.04 409.13
##
## Coefficients:
##
                         Estimate Std. Error t value
                                                                Pr(>|t|)
                                     17.1431 11.763 <0.00000000000000000
## (Intercept)
                         201.6624
## age
                          -0.4448
                                     0.2655 -1.675
                                                                  0.0941
## oralmed_factorYes
                         59.5417
                                     25.6102 2.325
                                                                  0.0202
## age:oralmed factorYes -0.8033
                                      0.3957 -2.030
                                                                  0.0426
##
## Residual standard error: 88.45 on 1159 degrees of freedom
##
     (267 observations deleted due to missingness)
## Multiple R-squared: 0.01979,
                                   Adjusted R-squared: 0.01726
## F-statistic: 7.801 on 3 and 1159 DF, p-value: 0.00003703
```

```
confint(mod.intx)
```

```
## 2.5 % 97.5 %

## (Intercept) 168.0274868 235.29740684

## age -0.9658059 0.07613036

## oralmed_factorYes 9.2940930 109.78933801

## age:oralmed_factorYes -1.5797124 -0.02697211
```

- The **fitted model** is given by the equation, $\hat{y}=$ 201.66 0.44 Age + 59.54 MedUse 0.8 Age imes MedUse.
- The **estimated slope** of the interaction age*oralmed_factor $b_3 = -0.8$ [95% CI (-1.58, -0.03)] is equal to the *difference* in the slope of age in those who use oral diabetes medication vs. those who do not (ref).
- A significance test of the interaction term $(H_0: \beta_3 = 0 \text{ vs. } \beta_3 \neq 0)$ reports a t-statistic t =-2.03, which is compared to a t-distribution with 1159 degrees of freedom. This test supports a significant difference in the effect of age on fasting glucose level in those who use oral diabetes medication vs. those who do not (p-value = 0.043).

The effect (slope) of age on fasting glucose level in those who use oral diabetes medication is estimated by b_1+b_3 ; the effect (slope) of age in those who do not use oral diabetes medication is estimated by b_1 . We can use $\bf R$ to compute the slope in the medication use group and perform a hypothesis test to determine if age significantly affects fasting glucose level in those who use oral diabetes medication by testing $H_0: \beta_1+\beta_3=0$ vs. $H_1: \beta_1+\beta_3\neq 0$.

We can estimate this slope and test this **linear contrast** by using the glht() function in the multcomp package. We begin by specifying κ , which identifies the coefficients that are involved in the estimation (i.e., b_1+b_3), or c(0, 1, 0, 1). We then specify κ in the linfct= argument of the glht() function to specify the linear hypothesis to be tested. summary() returns the estimate of the effect and the hypothesis test results; confint() returns a confidence interval for the effect.

```
# b1 + b3: Effect of age in those with oralmed_factor = 1

# Vector that specifies linear combination of coefficients interested in
K <- rbind(c(0, 1, 0, 1))  # 1 = coefficients "on" when estimating slope in oralmed_factor = 1

# Label for comparison (printed in the output)
rownames(K) <- "b1+b3 (slope in oralmed_factor = 1)"

# Estimate of slope (b1 + b3) and hypothesis test, glht() function in the "multcomp" package
summary(glht(mod.intx, linfct = K))</pre>
```

```
# Confidence interval for beta1 + beta3
confint(glht(mod.intx, linfct = K))
```

```
##
     Simultaneous Confidence Intervals
##
##
## Fit: lm(formula = fastgluc ~ age + oralmed_factor + age * oralmed_factor,
##
       data = nhanes)
##
## Ouantile = 1.962
##
   95% family-wise confidence level
##
##
## Linear Hypotheses:
                                             Estimate lwr
## b1+b3 (slope in oralmed_factor = 1) == 0 -1.2482 -1.8238 -0.6726
```

- Effect of **age** on fasting glucose levels in those who *use oral diabetes medication* is estimated by $b_1+b_3=-1.25$ [95% CI (-1.82, -0.67)]. We have evidence to reject $H_0:\beta_1+\beta_3=0$ and conclude that there is a significant association between age and fasting blood glucose in those who use oral diabetes medication (p-value <.001).
- Effect of **age** on fasting glucose levels in those who *do not use oral diabetes medication* is estimated by $b_1 = -0.44$ [95% CI (-0.97, 0.08)]. We do not have evidence to reject $H_0: \beta_1 = 0$ and cannot conclude that there is a significant association between age and fasting blood glucose in those who do not use oral diabetes medication (p-value = 0.094).

• The **estimated slope** of the interaction $b_3 = -0.8$ is equal to the *difference* in the slope of age in those who use oral diabetes medication vs. those who do not (ref). The test of the interaction is telling us that there is a significant difference in the effect of age in these two groups.

Automated Variable Selection

Automated variable selection methods have been developed to choose the "best-fitting" model (i.e., the "best" subset of predictors). There are three automated variable selection procedures:

- Backward elimination begins with the full model and iteratively removes predictors that contribute least to
 the model until all variables remaining exceed a certain threshold. Once a variable is removed, it cannot reenter the model. Backward elimination tends to be helpful if you have a modest-sized model and would like
 to eliminate a few predictors.
- 2. **Forward selection** begins with the null model (no predictors) and iteratively adds the most important predictors, stopping when there the amount of improvement is below a certain threshold. Once a variable is entered into the model, it cannot be removed. Forward selection tends to be helpful if you have a large set of potential predictors and wish to identify a few important variables.
- 3. **Stepwise selection** is a combination of forward selection and backward elimination. This method begins with the null model and iteratively adds predictors. After each addition, there is the option of removing any of the variables already included if removing that variable improves the model fit.

Automated selection methods can be based variable p-value thresholds or other model fit statistics, such as R_a^2 . The **Akaike Information Criterion (AIC)** and the **Bayes Information Criterion (BIC)** are other commonly used criteria. The goal is to *minimize* AIC and BIC (i.e., smaller AIC and BIC is better). Just like R_a^2 , both of these statistics penalize larger models and will not automatically decrease when additional variables are added to the model.

The stepAIC() function in the MASS package can be used to conduct automated variable selection based on the AIC. The **null model** (intercept only model) and the **full model** (model that includes all candidate predictors) must be defined.

stepAIC() Function Arguments	Option Definition
object=	Model object (full model for backward elimination, null model for forward and stepwise selection)
scope=	Range of models =list(lower = nullmodel, upper = fullmodel)
direction=	=both (stepwise), =backward (backward), =forward (forward)

Again, to avoid error messages about missing observations resulting in different data sets used in the null model and the full model, we fit the null model using the observations included in the full model data=mod.full\$model.

```
# Full model (contains all predictors under consideration)
mod.full <- lm(fastgluc ~ age + sex_factor + oralmed_factor + race_factor, data = nhanes)
# Null model (intercept only, notice data= here)
mod.null <- lm(fastgluc ~ 1, data = mod.full$model)</pre>
```

Backward elimination stops when the AIC does not improve (i.e., does not decrease) after removing a
predictor.

```
## Start: AIC=10433.13
## fastgluc ~ age + sex_factor + oralmed_factor + race_factor
##
##
                   Df Sum of Sq
                                    RSS
                                          AIC
                           1149 9045032 10431
## - sex_factor
## <none>
                                9043883 10433
## - race_factor
                    3
1
                          53238 9097120 10434
## - oralmed factor 1
                          26298 9070181 10434
                    1
                          91939 9135822 10443
## - age
##
## Step: AIC=10431.28
## fastgluc ~ age + oralmed_factor + race_factor
##
##
                   Df Sum of Sq
                                    RSS
                                          AIC
## <none>
                                9045032 10431
## - race factor
                          54361 9099393 10432
                    3
## - oralmed_factor 1
                          25805 9070837 10433
## - age
                     1
                          92232 9137264 10441
```

```
##
## Call:
## lm(formula = fastgluc ~ age + oralmed_factor + race_factor, data = nhanes)
##
## Coefficients:
##
                    (Intercept)
                                                          age
##
                       212.1288
                                                      -0.7075
##
             oralmed_factorYes
                                            race_factorBlack
##
                         9.5030
                                                      15.1203
## race_factorMexican-American
                                            race_factorOther
                         6.1850
##
                                                     -15.1268
```

• Forward selection stops when the AIC does not improve after adding a predictor.

```
## Start: AIC=10447.37
## fastgluc ~ 1
##
##
                   Df Sum of Sq
                                   RSS
                                        AIC
## + age
                        129225 9121018 10433
## + race_factor
                    3
                         89134 9161108 10442
## + oralmed_factor 1
                         19536 9230707 10447
## <none>
                               9250243 10447
## + sex_factor
                       3123 9247120 10449
##
## Step: AIC=10433.01
## fastgluc ~ age
##
##
                   Df Sum of Sq
                                   RSS
                                         AIC
## + oralmed_factor 1
                          21625 9099393 10432
## + race_factor 3
                          50181 9070837 10433
## <none>
                               9121018 10433
## + sex_factor
                    1 1574 9119444 10435
##
## Step: AIC=10432.25
## fastgluc ~ age + oralmed_factor
##
##
                Df Sum of Sq
                                RSS
                                      AIC
                   54361 9045032 10431
## + race_factor 3
## <none>
                            9099393 10432
                 1 2273 9097120 10434
## + sex_factor
##
## Step: AIC=10431.28
## fastgluc ~ age + oralmed_factor + race_factor
##
##
               Df Sum of Sq
                               RSS
                                     AIC
                           9045032 10431
## <none>
## + sex_factor 1 1149.3 9043883 10433
##
## lm(formula = fastgluc ~ age + oralmed_factor + race_factor, data = mod.full$model)
```

```
##
## Coefficients:
##
                    (Intercept)
                                                           age
##
                       212.1288
                                                       -0.7075
##
             oralmed_factorYes
                                             race factorBlack
##
                         9.5030
                                                      15.1203
                                             race_factorOther
## race_factorMexican-American
##
                         6.1850
                                                      -15.1268
```

• Stepwise selection stops when the AIC does not improve after potentially adding or removing a predictor.

```
# Stepwise selection
stepAIC(mod.null, scope = list(lower = mod.null, upper = mod.full),
    data = nhanes, direction = 'both')
```

```
## Start: AIC=10447.37
## fastgluc ~ 1
##
                   Df Sum of Sq
##
                                     RSS AIC
## + age
                         129225 9121018 10433
## + race_factor
                    3
                          89134 9161108 10442
## + oralmed_factor 1
                          19536 9230707 10447
## <none>
                                9250243 10447
                       3123 9247120 10449
## + sex_factor 1
##
## Step: AIC=10433.01
## fastgluc ~ age
##
##
                    Df Sum of Sq
                                     RSS AIC
                          21625 9099393 10432
## + oralmed_factor 1
## + race_factor 3
                           50181 9070837 10433
## <none>
                                 9121018 10433
## + sex_factor 1 1574 9119444 10435
                         129225 9250243 10447
## - age
                    1
##
## Step: AIC=10432.25
## fastgluc ~ age + oralmed_factor
##
                   Df Sum of Sq
##
                                     RSS
                                          AIC
## + race_factor
                    3 54361 9045032 10431
                                9099393 10432
## <none>
## - oralmed_factor 1 21625 9121018 10433
## + sex_factor 1 2273 9097120 10434
## - age
                    1
                         131314 9230707 10447
##
## Step: AIC=10431.28
## fastgluc ~ age + oralmed_factor + race_factor
##
                   Df Sum of Sq
##
                                     RSS
                                          AIC
## <none>
                                 9045032 10431
## - race_factor
                    3
                           54361 9099393 10432
## - oralmed_factor 1
                          25805 9070837 10433
                    1 1149 9043883 10433
1 92232 9137264 10441
## + sex_factor
## - age
                           92232 9137264 10441
```

```
##
## lm(formula = fastgluc ~ age + oralmed_factor + race_factor, data = mod.full$model)
##
## Coefficients:
##
                   (Intercept)
                                                          age
##
                      212.1288
                                                      -0.7075
##
             oralmed factorYes
                                            race_factorBlack
##
                         9.5030
                                                      15.1203
## race_factorMexican-American
                                            race_factorOther
##
                         6.1850
                                                     -15.1268
```

All three selection procedures exclude sex from the selected model but retain age, oral diabetes medication use and race. We consistently saw that sex was not an important predictor of fasting glucose levels. The selected model is given by,

```
# Selected model
mod.selected <- lm(fastgluc ~ age + oralmed_factor + race_factor, data = nhanes)
summary(mod.selected)</pre>
```

```
##
## Call:
## lm(formula = fastgluc ~ age + oralmed_factor + race_factor, data = nhanes)
##
## Residuals:
##
      Min
                1Q Median
                                3Q
                                       Max
## -154.89 -65.19 -27.71
                             50.56 400.61
##
## Coefficients:
##
                               Estimate Std. Error t value
                                                                        Pr(>|t|)
                                           14.8541 14.281 < 0.00000000000000000
## (Intercept)
                               212.1288
## age
                                -0.7075
                                            0.2060
                                                    -3.435
                                                                       0.000614
## oralmed factorYes
                                 9.5030
                                            5.2305
                                                     1.817
                                                                       0.069501
## race_factorBlack
                                            6.7680
                                                     2.234
                                15.1203
                                                                       0.025668
## race_factorMexican-American
                                 6.1850
                                            6.4136
                                                     0.964
                                                                        0.335063
                                           16.0788 -0.941
## race_factorOther
                               -15.1268
                                                                        0.347008
##
## Residual standard error: 88.42 on 1157 degrees of freedom
     (267 observations deleted due to missingness)
## Multiple R-squared: 0.02218,
                                    Adjusted R-squared: 0.01796
## F-statistic: 5.25 on 5 and 1157 DF, p-value: 0.00008964
```

• The final **fitted model** is given by the equation, $\hat{y}=212.13-0.71$ Age +9.5 MedUse +15.12 Black +6.19 Mexican-American -15.13 Other.

Notice that the variable <code>oralmed_factor</code> is included in the final model despite not being "statistically significant" at the $\alpha=0.05$ -level. Not all variables must be statistically significant to contribute to the model. In addition, you can refine the model further by again exploring the interaction of age and oral medication use or other plausible interactions in this model. I recommend not relying solely on automated variable selection methods to choose a final model. Rather, use your subject-area knowledge to help you build and refine a final model that makes sense in your application.