

## Investigation of a Gel Effect on Dental Disease by Generalized Linearized Model

### Introduction:

In this project, the investigators were interested in a new gel treatment for gum disease on 130 participants. The hypothesis of interest was whether the gel treatment would lower the pocket depth or attachment loss at one year. There were 4 categorical variables, treatment group, race, sex and smoker; and 6 numerical variables, age, sites, attachbase, attach1year, pdbase and pd1year in the dataset. Since there are very high correlations between “the baseline score” and “the score at one year”, hence we generated two new variables, “attachchange” and “dpchange”, in order to measure the changes of the pocket depth and attachment loss adjusting other factors. The diagnostic plots showed the data has missing data, but it is hard to impute the accurate data since there was very limited data points. GLM and T-test were applied for comparison to test the treatment significance among the different groups and between two different groups. GLMselect by stepwise selection was used to generate the regression linear model, which confirmed the result of GLM, and indicated the treatments were not significant.

### Methods:

Since there were missing data which were noted by “NA” in the dataset, the data were difficult to properly be imported in the SAS Studio (although it can be easily read and changed in R Studio). I had to change the NA into “.” in the Excel file before reading the data into SAS Studio.

The study interest was **whether treatment results in lower average pocket depth and attachment loss at one year**. The score difference/change was the interested

outcome for this research. Hence two new variables, attachchange and pdchange, were generated for data cleaning. In order to select the variables in the model, the data was investigated for correlations between the variables by “Proc Corr”.

Then the missing data pattern was showed that age had one missing data. Some of the participants were difficult to be followed up after 1 year by various reasons. There were 27 data points missing in attach loss at one year and pocket depth at one year.

The Gplot was to check the relationship between the outcomes and the variables. The numerical variables including age, sites, attachbase, attach1year, pdbase and pd1year are normally distributed or spread evenly, so we don't have to transform the data.

The mean and frequency were checked separately for the numerical variables and categorical variables in order to get the basic statistics for the data. The treatment groups were random selected and balanced, and there are 26 participants in each treatment group; sex and smoker were balanced too. There is 87% white in the race, although highly weighted than other three populations, randomly balanced dispersed in the treatment groups.

Since there were categorical variables and numerical variables in the dataset, the Generalized Linear Model (GLM) was used to check if the different treatment groups had the different effects on the two outcomes, pdchange and attachchange without the adjustment at the beginning. Fourthly, the Structured Query Language (SQL) was used to generated into different groups in pair and t-test was applied to test which two treatment groups could have different effects for two outcomes.

GLMSELECT was used to select the models and detect the potential significant interactions among the variables. The final regression models were obtained by using the selected variables from the GLMSELECT results. This step with interaction might complicate the study since the sample size was small in each treatment group. Without interaction, the final model was selected and treatment groups were excluded.

### **Results:**

Correlation was checked in the Form 1. There were very high correlations between the variables, attachbase and attach1year ( $r=0.946$ ), pdbase and pd1year ( $r=0.843$ ). But there were relative low correlations between the variables, attachchange and attachbase ( $r=-0.414$ ), also pdchange and pdbase ( $r=-0.203$ ). Hence the two variables, attachchange and pdchange, were selected as the new outcomes for the models.

### **From 1: Correlations between all the variables**

| Pearson Correlation Coefficients<br>Prob >  r  under H0: Rho=0<br>Number of Observations |                           |                           |                           |                           |                           |                           |                           |                           |
|--|---------------------------|---------------------------|---------------------------|---------------------------|---------------------------|---------------------------|---------------------------|---------------------------|
|  | pdchange                  | attachchange              | age                       | sites                     | pdbase                    | pd1year                   | attachbase                | attach1year               |
| pdchange   | 1.00000<br>103            | 0.53546<br><.0001<br>103  | -0.07422<br>0.4585<br>102 | -0.04812<br>0.6293<br>103 | -0.20267<br>0.0401<br>103 | 0.35543<br>0.0002<br>103  | -0.03676<br>0.7124<br>103 | 0.15128<br>0.1272<br>103  |
| attachchange   | 0.53546<br><.0001<br>103  | 1.00000<br>103            | -0.17458<br>0.0793<br>102 | 0.16209<br>0.1019<br>103  | -0.13472<br>0.1749<br>103 | 0.16531<br>0.0952<br>103  | -0.41437<br><.0001<br>103 | -0.09561<br>0.3367<br>103 |
| age  | -0.07422<br>0.4585<br>102 | -0.17458<br>0.0793<br>102 | 1.00000<br>129            | -0.11266<br>0.2037<br>129 | -0.11097<br>0.2106<br>129 | -0.12526<br>0.2097<br>102 | 0.12162<br>0.1698<br>129  | 0.08605<br>0.3898<br>102  |
| sites  | -0.04812<br>0.6293<br>103 | 0.16209<br>0.1019<br>103  | -0.11266<br>0.2037<br>129 | 1.00000<br>130            | -0.16634<br>0.0586<br>130 | -0.18852<br>0.0565<br>103 | -0.39708<br><.0001<br>130 | -0.36885<br>0.0001<br>103 |
| pdbase   | -0.20267<br>0.0401<br>103 | -0.13472<br>0.1749<br>103 | -0.11097<br>0.2106<br>129 | -0.16634<br>0.0586<br>130 | 1.00000<br>130            | 0.84327<br><.0001<br>103  | 0.58869<br><.0001<br>130  | 0.60498<br><.0001<br>103  |
| pd1year  | 0.35543<br>0.0002<br>103  | 0.16531<br>0.0952<br>103  | -0.12526<br>0.2097<br>102 | -0.18852<br>0.0565<br>103 | 0.84327<br><.0001<br>103  | 1.00000<br>103            | 0.54983<br><.0001<br>103  | 0.66049<br><.0001<br>103  |
| attachbase   | -0.03676<br>0.7124<br>103 | -0.41437<br><.0001<br>103 | 0.12162<br>0.1698<br>129  | -0.39708<br><.0001<br>130 | 0.58869<br><.0001<br>130  | 0.54983<br><.0001<br>103  | 1.00000<br>130            | 0.94556<br><.0001<br>103  |
| attach1year  | 0.15128<br>0.1272<br>103  | -0.09561<br>0.3367<br>103 | 0.08605<br>0.3898<br>102  | -0.36885<br>0.0001<br>103 | 0.60498<br><.0001<br>103  | 0.66049<br><.0001<br>103  | 0.94556<br><.0001<br>103  | 1.00000<br>103            |

Missing data analysis was shown in Form 2. There was one missing (0.77%) in age and 27 missing (20.77%) in pd1year and attach1year, which caused the same missing in the pdchange and attachchange. The reason for the missing was difficult for following up at the one year when the study ended. Since the percentage of the missing data was not

high and could be tolerate, and there were only two time points which was difficult to predict the trend at the one year, we analyzed the data without imputation for the missing data.

## Form 2: Missing data Descriptive for Continuous Variables

| The MEANS Procedure |     |        | The MI Procedure      |          |              |     |       |        |         |            |             |      |         |
|---------------------|-----|--------|-----------------------|----------|--------------|-----|-------|--------|---------|------------|-------------|------|---------|
| Variable            | N   | N Miss | Missing Data Patterns |          |              |     |       |        |         |            |             |      |         |
| pdchange            | 103 | 27     | Group                 | pdchange | attachchange | age | sites | pdbase | pd1year | attachbase | attach1year | Freq | Percent |
| attachchange        | 103 | 27     | 1                     | X        | X            | X   | X     | X      | X       | X          | X           | 102  | 78.46   |
| age                 | 129 | 1      | 2                     | X        | X            | .   | X     | X      | X       | X          | X           | 1    | 0.77    |
| sites               | 130 | 0      | 3                     | .        | .            | X   | X     | X      | .       | X          | .           | 27   | 20.77   |
| pdbase              | 130 | 0      |                       |          |              |     |       |        |         |            |             |      |         |
| pd1year             | 103 | 27     |                       |          |              |     |       |        |         |            |             |      |         |
| attachbase          | 130 | 0      |                       |          |              |     |       |        |         |            |             |      |         |
| attach1year         | 103 | 27     |                       |          |              |     |       |        |         |            |             |      |         |

Overall statistical description for variables was shown in Form 3-1 and 3-2. The two outcomes, pdchange and attachchange, had small mean values and small errors (standard deviation). The ages of the participants were ranged from 28 years old to 74 years old. The checked sites had broader range and collected enough numbers for statistical analysis. From 3-2 showed the smoker, age, treatment groups were well balanced, however, 87% white was in the race, and other three populations only had 13% total. The small sample number 26 in each treatment group indicated the study population was small.

## Form 3-1: The Means of the continuous variables

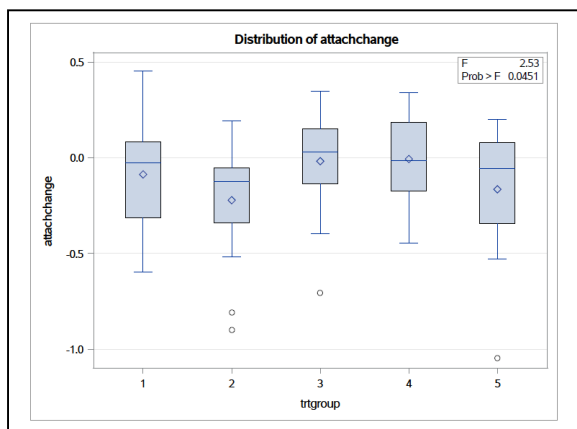
| The MEANS Procedure |     |             |            |             |             |
|---------------------|-----|-------------|------------|-------------|-------------|
| Variable            | N   | Mean        | Std Dev    | Minimum     | Maximum     |
| pdchange            | 103 | -0.2943543  | 0.2676105  | -0.8580247  | 0.4551282   |
| attachchange        | 103 | -0.0994510  | 0.2760304  | -1.0476190  | 0.4523810   |
| age                 | 129 | 49.9434708  | 10.0323275 | 28.5722108  | 74.5325120  |
| sites               | 130 | 157.5076923 | 11.3412509 | 114.0000000 | 168.0000000 |
| pdbase              | 130 | 3.1383709   | 0.4367214  | 2.2628205   | 5.2173913   |
| pd1year             | 103 | 2.8751627   | 0.4875549  | 1.9642857   | 4.8913043   |
| attachbase          | 130 | 2.1460753   | 0.7970523  | 0.8950617   | 5.0892857   |
| attach1year         | 103 | 2.1013885   | 0.7718840  | 0.8653846   | 5.3043478   |

### Form 3-2: The Frequency of the categorical variables

| Table of trtgroup by smoker |                               |                              |               | Table of trtgroup by race |                            |                             |                            |                               |               | Table of trtgroup by sex |                              |                               |               |
|-----------------------------|-------------------------------|------------------------------|---------------|---------------------------|----------------------------|-----------------------------|----------------------------|-------------------------------|---------------|--------------------------|------------------------------|-------------------------------|---------------|
| trtgroup                    | smoker                        |                              |               | trtgroup                  | race                       |                             |                            |                               |               | trtgroup                 | sex                          |                               |               |
|                             | 0                             | 1                            | Total         |                           | 1                          | 2                           | 4                          | 5                             | Total         |                          | 1                            | 2                             | Total         |
| 1                           | 15<br>11.63<br>57.69<br>18.52 | 11<br>8.53<br>42.31<br>22.92 | 26<br>20.16   | 1                         | 0<br>0.00<br>0.00<br>0.00  | 2<br>1.54<br>7.69<br>22.22  | 1<br>0.77<br>3.85<br>33.33 | 23<br>17.69<br>88.46<br>20.18 | 26<br>20.00   | 1                        | 11<br>8.46<br>42.31<br>20.37 | 15<br>11.54<br>57.69<br>19.74 | 26<br>20.00   |
| 2                           | 17<br>13.18<br>65.38<br>20.99 | 9<br>6.98<br>34.62<br>18.75  | 26<br>20.16   | 2                         | 1<br>0.77<br>3.85<br>25.00 | 1<br>0.77<br>3.85<br>11.11  | 1<br>0.77<br>3.85<br>33.33 | 23<br>17.69<br>88.46<br>20.18 | 26<br>20.00   | 2                        | 10<br>7.69<br>38.46<br>18.52 | 16<br>12.31<br>61.54<br>21.05 | 26<br>20.00   |
| 3                           | 18<br>13.95<br>69.23<br>22.22 | 8<br>6.20<br>30.77<br>16.67  | 26<br>20.16   | 3                         | 1<br>0.77<br>3.85<br>25.00 | 5<br>3.85<br>19.23<br>55.56 | 0<br>0.00<br>0.00<br>0.00  | 20<br>15.38<br>76.92<br>17.54 | 26<br>20.00   | 3                        | 11<br>8.46<br>42.31<br>20.37 | 15<br>11.54<br>57.69<br>19.74 | 26<br>20.00   |
| 4                           | 14<br>10.85<br>56.00<br>17.28 | 11<br>8.53<br>44.00<br>22.92 | 25<br>19.38   | 4                         | 0<br>0.00<br>0.00<br>0.00  | 0<br>0.00<br>0.00<br>0.00   | 1<br>0.77<br>3.85<br>33.33 | 25<br>19.23<br>96.15<br>21.93 | 26<br>20.00   | 4                        | 11<br>8.46<br>42.31<br>20.37 | 15<br>11.54<br>57.69<br>19.74 | 26<br>20.00   |
| 5                           | 17<br>13.18<br>65.38<br>20.99 | 9<br>6.98<br>34.62<br>18.75  | 26<br>20.16   | 5                         | 2<br>1.54<br>7.69<br>50.00 | 1<br>0.77<br>3.85<br>11.11  | 0<br>0.00<br>0.00<br>0.00  | 23<br>17.69<br>88.46<br>20.18 | 26<br>20.00   | 5                        | 11<br>8.46<br>42.31<br>20.37 | 15<br>11.54<br>57.69<br>19.74 | 26<br>20.00   |
| Total                       | 81<br>62.79                   | 48<br>37.21                  | 129<br>100.00 | Total                     | 4<br>3.08                  | 9<br>6.92                   | 3<br>2.31                  | 114<br>87.69                  | 130<br>100.00 | Total                    | 54<br>41.54                  | 76<br>58.46                   | 130<br>100.00 |
| Frequency Missing = 1       |                               |                              |               |                           |                            |                             |                            |                               |               |                          |                              |                               |               |

Generalized Linear Model (GLM) were applied to compare if the five different treatment groups would lead to significant changes in attach loss or pocket depth. Figure 1 showed there was significant difference for the five treatment groups for attach loss change ( $p=0.0451$ ); but there was not significant difference for the five treatment groups for pocket depth change ( $p=0.0899$  or  $p=0.1208$ ).

Figure 1



Form4: t-test of treatment for attachchange

| treatment | P-value | treatment | P-value |
|-----------|---------|-----------|---------|
| Trt1&2    | 0.0883  | Trt2&3    | 0.0174  |
| Trt1&3    | 0.3706  | Trt2&4    | 0.0096  |
| Trt1&4    | 0.2733  | Trt2&5    | 0.5735  |
| Trt1&5    | 0.3981  |           |         |

Form5: GLM for attachchange and pdchange

| Source     | DF | Type III SS | Mean Square | F Value | Pr > F | Source     | DF | Type III SS | Mean Square | F Value | Pr > F |
|------------|----|-------------|-------------|---------|--------|------------|----|-------------|-------------|---------|--------|
| trtgroup   | 4  | 0.47831707  | 0.11957927  | 1.97    | 0.1063 | trtgroup   | 4  | 0.52588496  | 0.13147124  | 1.88    | 0.1208 |
| attachbase | 1  | 0.84801112  | 0.84801112  | 13.95   | 0.0003 | attachbase | 1  | 0.00059475  | 0.00059475  | 0.01    | 0.9267 |
| race       | 1  | 0.10457025  | 0.10457025  | 1.72    | 0.1930 | sex        | 1  | 0.27805833  | 0.27805833  | 3.98    | 0.0492 |
| sex        | 1  | 0.02876959  | 0.02876959  | 0.47    | 0.4933 | age        | 1  | 0.07995641  | 0.07995641  | 1.14    | 0.2878 |
| age        | 1  | 0.08468370  | 0.08468370  | 1.39    | 0.2410 | race       | 1  | 0.06127884  | 0.06127884  | 0.88    | 0.3517 |
| smoker     | 1  | 0.14215054  | 0.14215054  | 2.34    | 0.1297 | smoker     | 1  | 0.00395873  | 0.00395873  | 0.06    | 0.8125 |
| sites      | 1  | 0.01509983  | 0.01509983  | 0.25    | 0.6194 | sites      | 1  | 0.03928912  | 0.03928912  | 0.56    | 0.4555 |

T-test was further to investigate which treatment groups were significant different for the outcome, attachment loss change. Without adjustment, the average change of attach

loss was statistical significance ( $P=0.0451$ ) among the five treatment groups, specifically between control group and low concentrations (Trt2&3:  $P=0.0174$ ), or medium concentrations group (Trt2&4:  $P=0.0096$ ). However, the Placebo group had not significant difference from all other groups. Adjusted by other factors, baseline, race, smoker, sex and age, by GLM shown in Form 5, the treatments were not significant anymore ( $p=0.1063$ ). GLMselect also excluded the treatment groups in the model.

### **Conclusions:**

The result showed that brushing the study gel on the gum with active ingredient, had no significant difference from the gel without any active ingredient. The new gel might not have clinical significance. After adjusted by baseline, race, smoker, sex and age, the treatments were not significant ( $p=0.1063$ ) and lost statistical significance. The new gel did not lower the pocket depth since there was no significant difference for the five treatment groups with/without adjusted by other variables ( $p=0.1208$  or  $p=0.0899$ ).

Hence, adjusted by demographic factors, the new gel had no significant effect on the treatment population, and was not promising to test for clinical trial. But baseline may affect attach loss ( $p<.0001$ ), and sex may affect pocket depth ( $p=0.049$ ).

**Limitation:** The number of the precipitants was small, 26 samples in each treatment group. If the population increased, the model might be more accurate and the clinical significance might be much better addressed. Since the small group in the data, and it as hard to track the reason and conditions for the persons who failed to show up at one year, it was difficult to impute the accurate values for the missing data.

Reproducible research information: See below (**Reproducible Research**).

```
/*read in the raw dental data*/
FILENAME Dental '/home/lingdizhang68/6624/Project0/Data/Project0_dental_data.csv';

PROC IMPORT DATAFILE=Dental
  DBMS=CSV replace
  OUT=WORK.IMPORT;
  GETNAMES=YES;
RUN;

/* generate new variables for the change from baseline to one year*/
DATA dentalclean;
set WORK.IMPORT;
attachchange=attach1year-attachbase;
pdchange=pd1year-pdbase;
RUN;

***Export the clean dataset for use in other analysis programs***;
PROC EXPORT DATA= WORK.dentalclean
  OUTFILE= '/home/lingdizhang68/6624/Project0/dentalclean.csv'
  DBMS=CSV REPLACE;
  PUTNAMES=YES;
RUN;

PROC PRINT data=Dentalclean;
run;

/****open ods (output delivery system) to write all my figures      ***
/****and tables to a directory and file of my choosing for editing ***
/****later                                                         ***

ods listing close;
ods html path='/home/lingdizhang68/6624/Project0/Reports' file = 'DDescriptives.html';

/*import the new clean dataset*/
PROC IMPORT OUT= WORK.dentalclean
  DATAFILE= '/home/lingdizhang68/6624/Project0/dentalclean.csv'
  DBMS=CSV REPLACE;
  GETNAMES=YES;
  DATAROW=2;
RUN;

/*check the missing data and mean for numerical variables*/
PROC MEANS data=Dentalclean N NMISS;
  var pdchange attachchange age sites pdbase pd1year attachbase attach1year;
  title 'Missing data description for Continuous Variables in Dental-Project0';
RUN;title;

ods select MissPattern;
PROC MI data=Dentalclean nimpute=0;
var pdchange attachchange age sites pdbase pd1year attachbase attach1year;
run;

/*Descriptive Statistics for the dataset*/
PROC MEANS data=Dentalclean;
  var pdchange attachchange age sites pdbase pd1year attachbase attach1year;
  title 'Descriptive Statistics for Continuous Variables in Dental-Project0';
RUN;title;

PROC FREQ data=Dentalclean;
  tables smoker sex trtgroup race;
  title 'Descriptive Statistics for the Categorical Variables in Dental-Project0';
RUN;title;

PROC FREQ data=Dentalclean;
  tables trtgroup*race;
```



```
title 'Descriptive Statistics for treatment and race';
RUN;title;

PROC CORR data=Dentalclean;
var pdchange attachchange age sites pdbase pd1year attachbase attach1year;
title 'Correlations between all variables: After data cleaning';
RUN;title;

PROC GPLOT data=Dentalclean;
plot attachchange*(trtgroup race age sites smoker sex);
symbol I=r1 value=dot color=black;
RUN;

PROC GPLOT data=Dentalclean;
plot pdchange*(trtgroup race age sites smoker sex);
symbol I=r1 value=dot color=red;
RUN;

QUIT

ods graphics on;

/*check if the treatment group has important effect on the outcomes*/

PROC GLM data=Dentalclean;
class trtgroup;
model attachchange=trtgroup;
run;
/*0.045*/
/* below t-test still find there is no clinical meaning for the treatment group*/

PROC GLM data=Dentalclean;
class trtgroup;
model attachchange=trtgroup attachbase race sex age smoker sites;
run;
/*0.1063*/

PROC GLM data=Dentalclean;
class trtgroup sex race;
model attachchange=trtgroup attachbase race sex age smoker sites;
run;
/*0.0907*/

quit

/*t-test to check if two treatment groups are significantly different */

ods graphics on;
proc sql;
create table trt12 as
select *
from Dentalclean
where trtgroup=1 or trtgroup=2;
quit;

proc sql;
create table trt13 as
select *
from Dentalclean
where trtgroup=1 or trtgroup=3;
quit;

proc sql;
create table trt14 as
select *
from Dentalclean
```

```
where trtgroup=1 or trtgroup=4;
quit;
```

```
proc sql;
create table trt15 as
select *
from Dentalclean
where trtgroup=1 or trtgroup=5;
quit;
```

```
proc sql;
create table trt23 as
select *
from Dentalclean
where trtgroup=2 or trtgroup=3;
quit;
```

```
proc sql;
create table trt24 as
select *
from Dentalclean
where trtgroup=2 or trtgroup=4;
quit;
```

```
proc sql;
create table trt25 as
select *
from Dentalclean
where trtgroup=2 or trtgroup=5;
quit;
```

```
proc ttest data=trt12;
class trtgroup;
var attachchange;
run;
```

```
/*0.0883*/
```

```
proc ttest data=trt13;
class trtgroup;
var attachchange;
run;
/*0.3706*/
```

```
proc ttest data=trt14;
class trtgroup;
var attachchange;
run;
/*0.2733*/
```

```
proc ttest data=trt15;
class trtgroup;
var attachchange;
run;
```

```
/0.3081*/
```

```
proc ttest data=trt23;
class trtgroup;
var attachchange;
run;
/*0.0176*/
```

```
proc ttest data=trt24;
class trtgroup;
var attachchange;
```

```
run;
/*0.0096*/
```

```
proc ttest data=trt25;
class trtgroup;
var attachchange;
run;
/*0.5635*/
```

```
proc ttest data=trt12;
class trtgroup;
var pdchange;
run;

/*0.8792*/
```

```
proc ttest data=trt23;
class trtgroup;
var pdchange;
run;
/*0.0944*/
```

```
proc ttest data=trt24;
class trtgroup;
var pdchange;
run;
/*0.0852*/
```

```
proc ttest data=trt25;
class trtgroup;
var pdchange;
run;
/*0.5717*/
```

```
PROC GLM data=Dentalclean;
class trtgroup;
model pdchange=trtgroup;
run;
/*0.089*/
```

```
PROC GLM data=Dentalclean;
class trtgroup;
model pdchange=trtgroup attachbase sex age race smoker sites;
run;
/0.1208*/
```

```
PROC GLM data=Dentalclean;
class trtgroup sex race;
model pdchange=trtgroup attachbase race sex age smoker sites;
run;
/*0.1276*/
```

```
/*model selection for the regression linear model*/
proc glmselect data=Dentalclean;
  class trtgroup sex race smoker;
  model attachchange=attachbase trtgroup age sex race smoker sites/selection=stepwise(select=SL SLS=0.05);
run;

/*attachbase is significant p<0.001*/
```

```
proc glmselect data=Dentalclean;
  class trtgroup sex race smoker;
  model pdchange=pbbase trtgroup age sex race smoker sites/selection=stepwise(select=SL SLS=0.05);
run;

/*sex is significant p=0.0330*/
```

```
/*please forget the interaction term and imputation*/

/*linear regression for model selection*/
ods graphics on;
proc glmselect data=Dentalclean;
  class trtgroup sex race smoker;
  model attachchange=attachbase|trtgroup|age|sex|race|smoker|sites/selection=stepwise(select=SL SLS=0.05);
run;

proc glmselect data=Dentalclean;
  class trtgroup sex race smoker;
  model attachchange=attachbase trtgroup age sex race smoker sites/selection=stepwise(select=SL SLS=0.05);
run;

/*attachbase is significant p<0.001*/

proc glmselect data=Dentalclean;
  class trtgroup sex race smoker;
  model pdchange=pdbase trtgroup age sex race smoker sites/selection=stepwise(select=SL SLS=0.05);
run;

/*sex is significant p=0.0330*/

proc glm data=Dentalclean PLOTS=(DIAGNOSTICS RESIDUALS);
  class trtgroup sex race smoker;
  model attachchange=attachbase trtgroup smoker attachbase*smoker/solution;
  output out=d_attach COOKD=COOKD STUDENT=STUDENT;
/*R=0.3492*/

proc glm data=Dentalclean PLOTS=(DIAGNOSTICS RESIDUALS);
  class trtgroup sex race smoker;
  model attachchange=attachbase trtgroup/solution;
  output out=d_attach COOKD=COOKD STUDENT=STUDENT;
/*R=0.237*/

quit;

ods html close;
ods listing;

/*impute the missing data*/

PROC MI DATA = Dentalclean;
EM OUT = Dentalimpute;
VAR attach1year pd1year age;
RUN;

PROC PRINT data=Dentalimpute;
run;

/*clean data by generating the change scores*/
DATA dental1;
set Dentalimpute;
dattach=attach1year-attachbase;
```

```
dchange=pd1year-pdbase;  
RUN;
```

---

```
PROC PRINT data=dental1;  
run;
```

```
/*select the interaction and run linear regression model*/
```

---

```
proc glmselect data=dental1;  
  class trtgroup sex race smoker;  
  model dattach=attachbase|trtgroup|age|sex|race|smoker|sites/selection=stepwise(select=SL SLS=0.05);  
run;
```

---

```
proc glmselect data=dental1;  
  class trtgroup sex race smoker;  
  model dchange=pdbase|trtgroup|age|sex|race|smoker|sites/selection=stepwise(select=SL SLS=0.05);  
run;
```

---

```
proc glm data=dental1 PLOTS=(DIAGNOSTICS RESIDUALS);  
  class trtgroup sex race smoker;  
  model dattach=attachbase trtgroup sex smoker attachbase*smoker/solution;  
  output out=d_attach COOKD=COOKD STUDENT=STUDENT;  
quit;
```

---

```
proc glm data=dental1 PLOTS=(DIAGNOSTICS RESIDUALS);  
  class trtgroup sex race smoker;  
  model dchange=pdbase trtgroup pdbase*sites*sex/solution;  
  output out=attach1year COOKD=COOKD STUDENT=STUDENT;
```