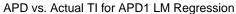
Infant Infection Timing Models

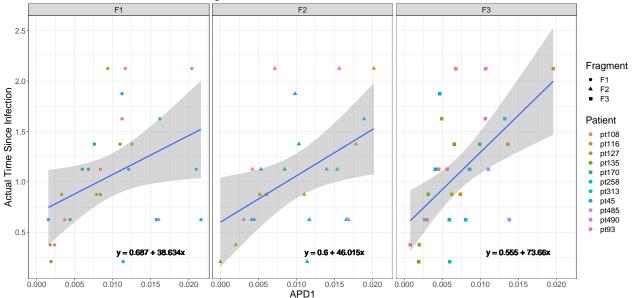
```
library(ggplot2)
library(devtools)
library(gridExtra)
library(ggpubr)
library(data.table)
library(Metrics)
library(tidyverse)
library(GGally)
source("Infant_functions.R")
source("Infant_predict.R")
source("Infant_plotting.R")
knitr::opts_chunk$set(fig.width=16, fig.height=8)
# Set working directory
setwd("/Users/magdalenarussell/Documents/Matsen_group/infection-timing/Infant_Model")
# Load existing data
training_data = read.csv("../_ignore/AllRunsAvg.csv")
testing_data = read.csv("../_ignore/AllRunsAvg.csv")
```

Note: for all of the following analyses, I have assigned the actual time since infection for all patients to be the midpoint of the third trimester (i.e. we are assigning birth to be time zero, so the actual infection time here is -0.125 years.)

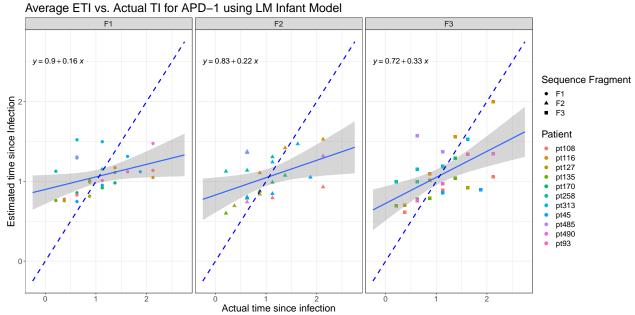
The following plots show the models created for the data using an APD cutoff of 0.01. First you will see the actual time since infection versus APD regression (using the indicated new modelling strategy). You will see the model statistics below these plots (i.e. model equation coefficients (also show on the plots in the form of an equation) and p values for each coefficient). Second you will see the relationship between the estimated time since infection and the actual time since infection. The perfect correlation (show as the blue spotted line) depicts the y = x line where actual time since infection = estimated time since infection. For these plots, I am showing a LM regression through the points with a confidence interval to show the trend of these estimated times. However, these estimated times were calculated using the indicated new modelling strategy (not always LM regression). Lastly, you will see, again, a plot comparing the estimated time since infection and the actual time since infection. Here, the plots show the trends per individual.

```
for (ty in c("LM", "LM_origin", "LAD", "LAD_origin", "LM_GEE")){
   together = compile_data(train_data = training_data, new_data = testing_data, apd = 1, type = ty)
   plot_APD_TI_regression(together, apd = 1, type = ty)
   show_model_statistics(apd = 1, type= ty)
   plot_ETI_TI_regression(together, apd = 1, type = ty)
}
```

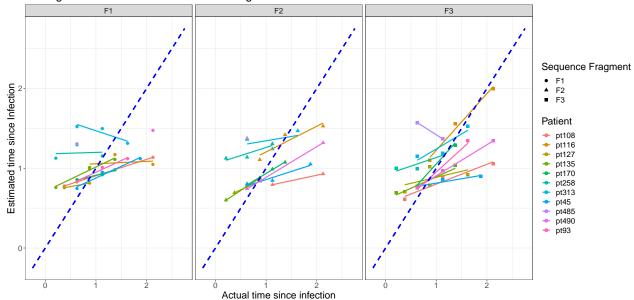




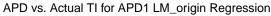
```
## [1] "LM Model for APD1 and Fragment F1 Summary Statistics:"
                 Estimate Std. Error t value
                           0.204124 3.364976 0.002473165
## (Intercept) 0.6868725
              38.6337771 17.886224 2.159974 0.040561325
  avg_apd1
## [1] "LM Model for APD1 and Fragment F2 Summary Statistics:"
                Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) 0.6001113 0.2144139 2.798844 0.01046298
               46.0145096 18.6146676 2.471949 0.02165105
## avg_apd1
## [1] "LM Model for APD1 and Fragment F3 Summary Statistics:"
##
                Estimate Std. Error t value
                                                 Pr(>|t|)
## (Intercept) 0.5554609 0.1612772 3.444138 0.0017649364
## avg_apd1
              73.6599533 19.6573417 3.747198 0.0007911118
```

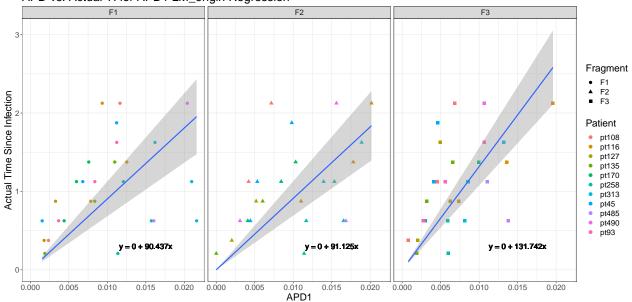


Average ETI vs. Actual TI for APD-1 using LM Infant Model

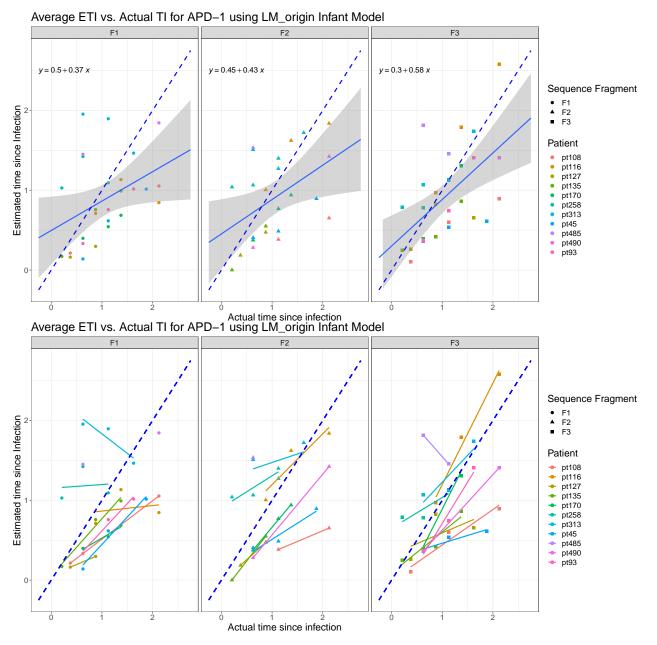


Saving 16 x 8 in image



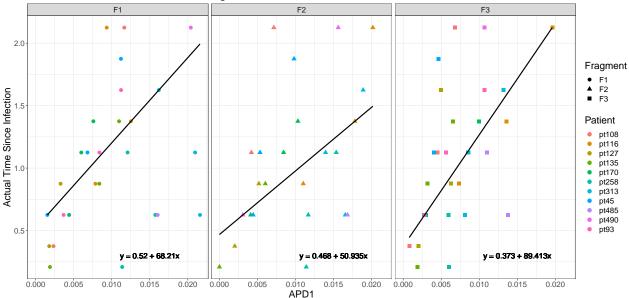


```
## [1] "LM_origin Model for APD1 and Fragment F1 Summary Statistics:"
## Estimate Std. Error t value Pr(>|t|)
## avg_apd1 90.43658 10.76303 8.402518 6.960345e-09
## [1] "LM_origin Model for APD1 and Fragment F2 Summary Statistics:"
## Estimate Std. Error t value Pr(>|t|)
## avg_apd1 91.12475 10.60674 8.591209 1.235489e-08
## [1] "LM_origin Model for APD1 and Fragment F3 Summary Statistics:"
## Estimate Std. Error t value Pr(>|t|)
## avg_apd1 131.7423 11.78763 11.17632 3.227911e-12
```



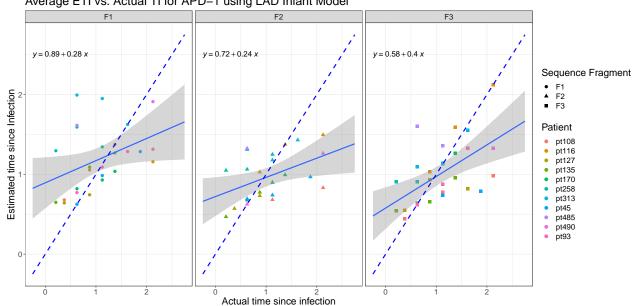
Saving 16 x 8 in image



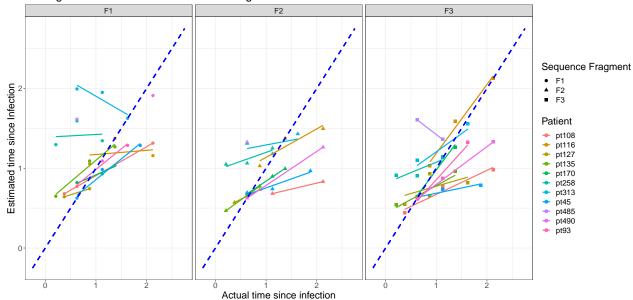


```
## [1] "LAD Model for APD1 and Fragment F1 Summary Statistics:"
                Estimate Std.Error Z value
                                                p-value
## (Intercept) 0.5201617 0.3100915 1.677446 0.09345531
               68.2096252 27.1715490 2.510333 0.01206175
  avg_apd1
## [1] "LAD Model for APD1 and Fragment F2 Summary Statistics:"
                Estimate Std.Error Z value
##
                                                p-value
               0.4681131 0.3108028 1.506142 0.13203072
## (Intercept)
              50.9345741 26.9828060 1.887668 0.05907056
## avg_apd1
## [1] "LAD Model for APD1 and Fragment F3 Summary Statistics:"
##
                Estimate Std.Error Z value
                                                  p-value
## (Intercept) 0.3732041 0.2183118 1.709500 0.0873582913
## avg_apd1
              89.4130783 26.6090256 3.360254 0.0007787091
```

Average ETI vs. Actual TI for APD-1 using LAD Infant Model

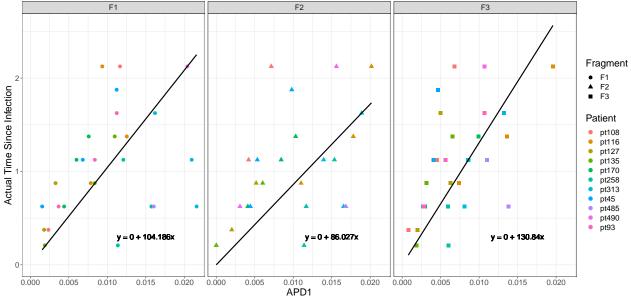


Average ETI vs. Actual TI for APD-1 using LAD Infant Model

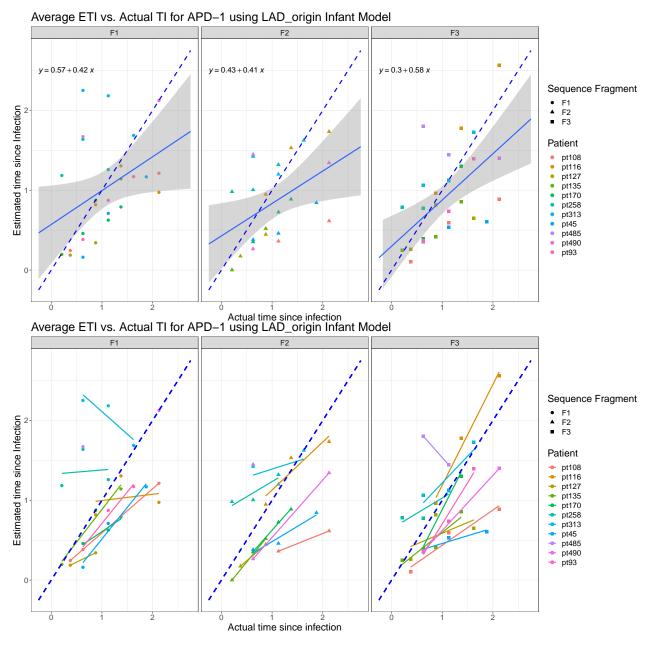


Saving 16 x 8 in image

APD vs. Actual TI for APD1 LAD_origin Regression

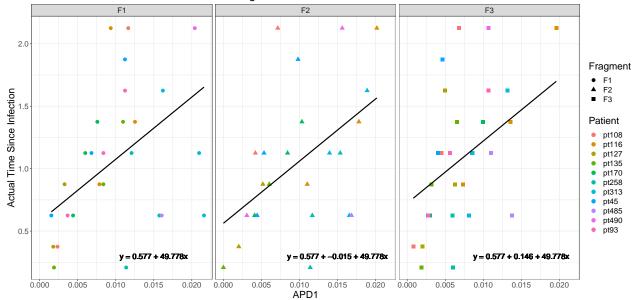


```
## [1] "LAD_origin Model for APD1 and Fragment F1 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd1 104.1856  16.23346 6.417952 1.3812e-10
## [1] "LAD_origin Model for APD1 and Fragment F2 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd1 86.02727  16.80154 5.120203 3.052072e-07
## [1] "LAD_origin Model for APD1 and Fragment F3 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd1 130.8403  17.24915 7.585319 3.316704e-14
## Saving 16 x 8 in image
```

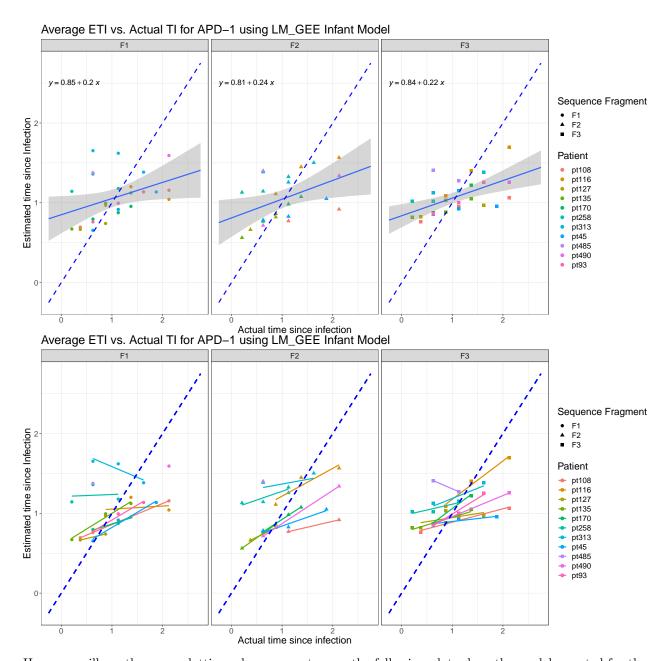


Saving 16 x 8 in image



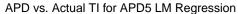


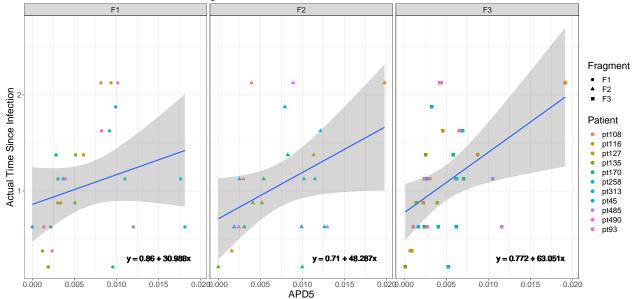
```
## [1] "LM_GEE Model for APD1 Summary Statistics:"
##
                         Estimate Naive S.E.
                                                Naive z Robust S.E.
## (Intercept)
                       0.57741090 0.1429107
                                              4.0403616
                                                         0.11171322
                      49.77758358 10.6470437
                                              4.6752493 15.16451484
## avg_apd1
## factor(fragment)F2 -0.01482986
                                  0.1419875 -0.1044448
                                                         0.05995904
## factor(fragment)F3 0.14614874
                                   0.1364798
                                             1.0708449
                                                         0.05297275
                        Robust z
## (Intercept)
                       5.1686893
## avg_apd1
                       3.2825042
## factor(fragment)F2 -0.2473333
## factor(fragment)F3
                      2.7589416
##
          (Intercept)
                                avg_apd1 factor(fragment)F2
                                               8.046503e-01
##
         2.357415e-07
                            1.028895e-03
## factor(fragment)F3
         5.798890e-03
## Saving 16 x 8 in image
```



Here you will see the same plotting scheme except, now, the following plots show the models created for the data using an APD cutoff of 0.05.

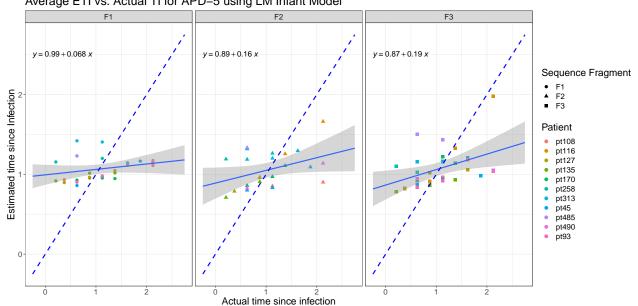
```
for (ty in c("LM", "LM_origin", "LAD", "LAD_origin", "LM_GEE")){
   together = compile_data(train_data = training_data, new_data = testing_data, apd = 5, type = ty)
   plot_APD_TI_regression(together, apd = 5, type = ty)
   show_model_statistics(apd = 5, type= ty)
   plot_ETI_TI_regression(together, apd = 5, type = ty)
}
```



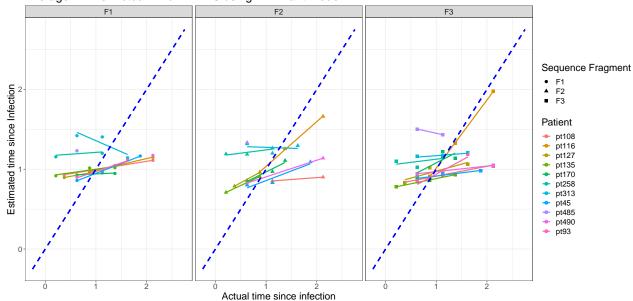


```
## [1] "LM Model for APD5 and Fragment F1 Summary Statistics:"
                 Estimate Std. Error t value
## (Intercept) 0.8595803 0.1880881 4.570094 0.0001135604
               30.9880608 22.9409226 1.350777 0.1888674317
  avg_apd5
## [1] "LM Model for APD5 and Fragment F2 Summary Statistics:"
##
                 Estimate Std. Error t value
                                                 Pr(>|t|)
               0.7097526  0.2042171  3.475481  0.002146657
## (Intercept)
               48.2872289 23.6799716 2.039159 0.053625692
## avg_apd5
## [1] "LM Model for APD5 and Fragment F3 Summary Statistics:"
##
                 Estimate Std. Error t value
                                                  Pr(>|t|)
## (Intercept) 0.7723701 0.1457767 5.298309 1.107161e-05
## avg_apd5
              63.0511604 23.8721359 2.641203 1.316952e-02
```

Average ETI vs. Actual TI for APD-5 using LM Infant Model

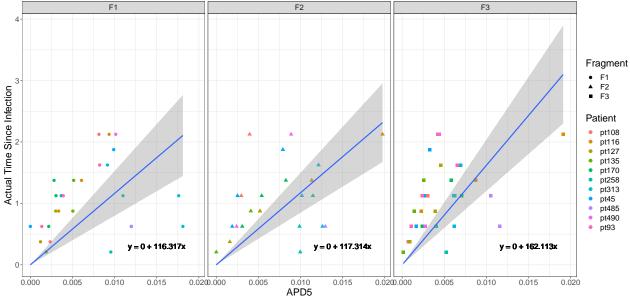


Average ETI vs. Actual TI for APD-5 using LM Infant Model

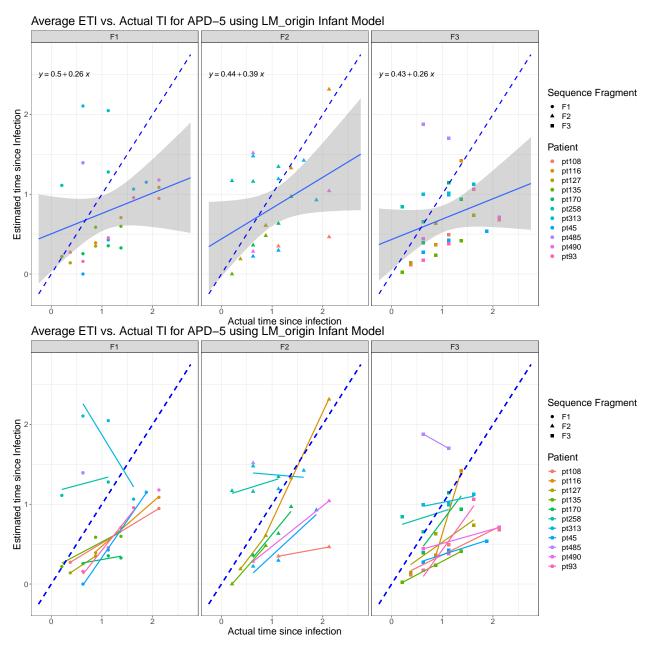


Saving 16 x 8 in image

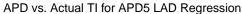


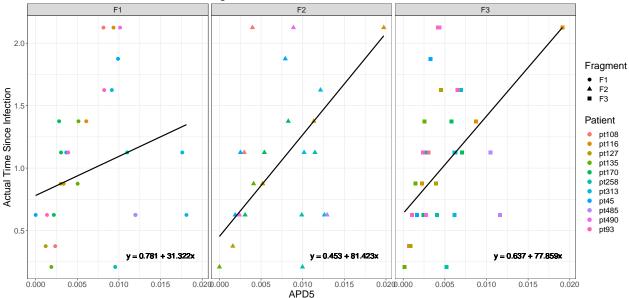


```
## [1] "LM_origin Model for APD5 and Fragment F1 Summary Statistics:"
## Estimate Std. Error t value Pr(>|t|)
## avg_apd5 116.3167 17.70788 6.568642 5.772489e-07
## [1] "LM_origin Model for APD5 and Fragment F2 Summary Statistics:"
## Estimate Std. Error t value Pr(>|t|)
## avg_apd5 117.3145 15.69606 7.474136 1.349512e-07
## [1] "LM_origin Model for APD5 and Fragment F3 Summary Statistics:"
## Estimate Std. Error t value Pr(>|t|)
## avg_apd5 162.1126 20.47231 7.918627 7.73886e-09
```



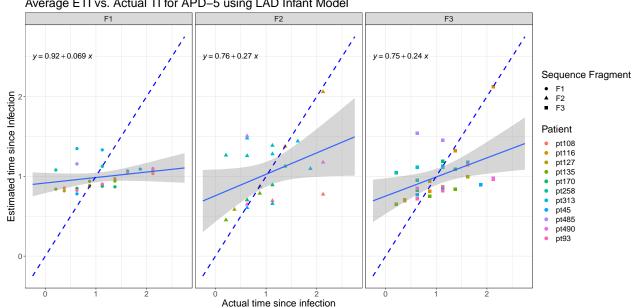
Saving 16 x 8 in image



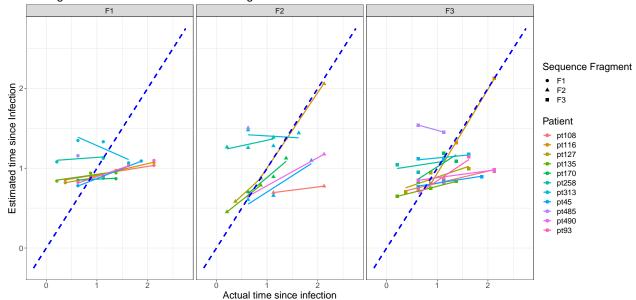


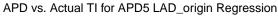
```
[1] "LAD Model for APD5 and Fragment F1 Summary Statistics:"
               Estimate Std.Error
                                    Z value
                                               p-value
                        0.2995866 2.6057532 0.009167249
## (Intercept) 0.7806487
              31.3216953 36.5402881 0.8571825 0.391344035
  avg_apd5
## [1] "LAD Model for APD5 and Fragment F2 Summary Statistics:"
##
               Estimate Std.Error Z value
                                             p-value
              ## (Intercept)
             81.4233551 33.3622682 2.440582 0.01466361
## avg_apd5
## [1] "LAD Model for APD5 and Fragment F3 Summary Statistics:"
##
               Estimate Std.Error Z value
                                              p-value
## (Intercept) 0.6373845 0.2151962 2.961876 0.003057712
## avg_apd5
             77.8592682 35.2401532 2.209391 0.027147475
```

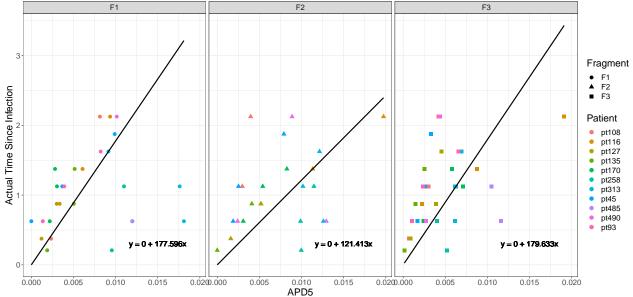
Average ETI vs. Actual TI for APD-5 using LAD Infant Model



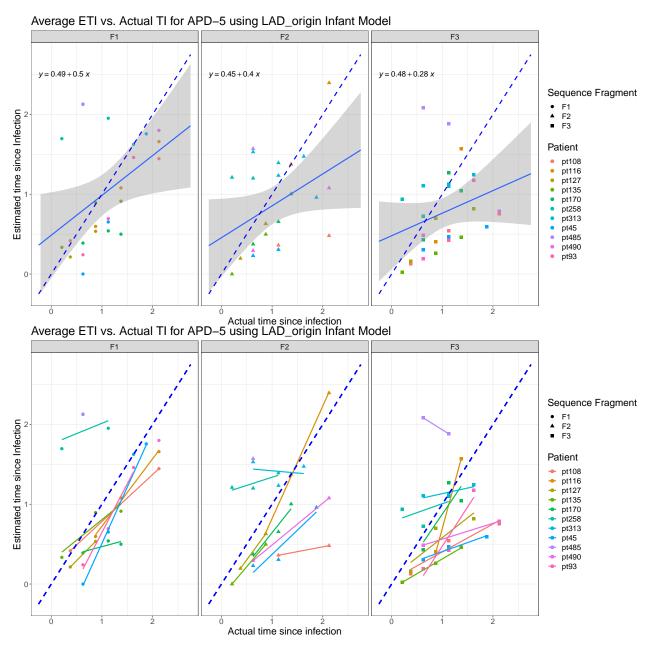






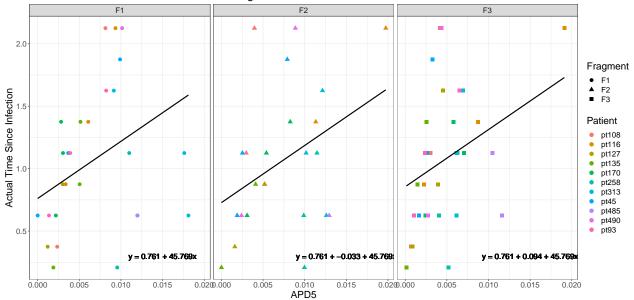


```
## [1] "LAD_origin Model for APD5 and Fragment F1 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd5 177.5964 29.5629 6.007409 1.885118e-09
## [1] "LAD_origin Model for APD5 and Fragment F2 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd5 121.4133 24.63826 4.927837 8.314501e-07
## [1] "LAD_origin Model for APD5 and Fragment F3 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd5 179.6327 32.27193 5.566221 2.603229e-08
## Saving 16 x 8 in image
```



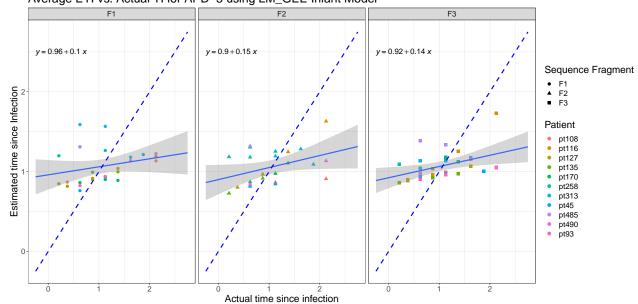
Saving 16 x 8 in image



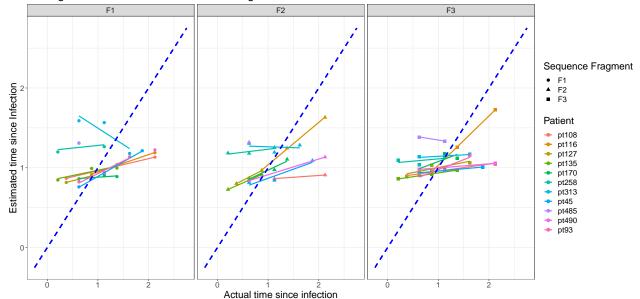


```
## [1] "LM_GEE Model for APD5 Summary Statistics:"
                      Estimate Naive S.E.
##
                                           Naive z Robust S.E.
                                                              Robust z
## (Intercept)
                     8.057242
## avg_apd5
                    45.76861490 13.3981106 3.4160500 20.27227328 2.257695
## factor(fragment)F2 -0.03298185 0.1500156 -0.2198561 0.06374178 -0.517429
## factor(fragment)F3 0.09407500 0.1428497 0.6585594 0.04668233 2.015216
         (Intercept)
                             avg_apd5 factor(fragment)F2
##
        8.881784e-16
                         2.396467e-02
                                          6.048567e-01
##
## factor(fragment)F3
##
        4.388199e-02
```

Average ETI vs. Actual TI for APD-5 using LM_GEE Infant Model





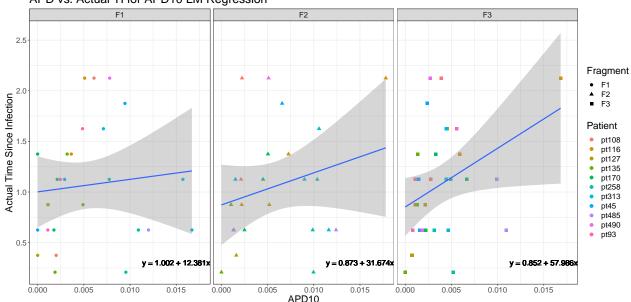


Here, again, you will see the same plotting scheme except, now, the following plots show the models created for the data using an APD cutoff of 0.1.

```
for (ty in c("LM", "LM_origin", "LAD", "LAD_origin", "LM_GEE")){
   together = compile_data(train_data = training_data, new_data = testing_data, apd = 10, type = ty)
   plot_APD_TI_regression(together, apd = 10, type = ty)
   show_model_statistics(apd = 10, type= ty)
   plot_ETI_TI_regression(together, apd = 10, type = ty)
}
```

Saving 16 x 8 in image

APD vs. Actual TI for APD10 LM Regression



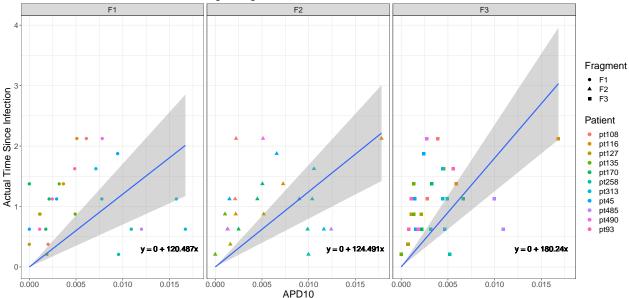
```
## [1] "LM Model for APD10 and Fragment F1 Summary Statistics:"
## Estimate Std. Error t value Pr(>|t|)
## (Intercept) 1.001571 0.1707357 5.8662067 4.045402e-06
```

```
## avg_apd10
                    12.380843 24.5210732 0.5049063 6.180467e-01
 ## [1] "LM Model for APD10 and Fragment F2 Summary Statistics:"
                       Estimate Std. Error t value
 ##
 ## (Intercept) 0.8727834
                                     0.191634 4.554429 0.0001557378
                    31.6735168 25.771600 1.229009 0.2320527190
 ## avg_apd10
 ## [1] "LM Model for APD10 and Fragment F3 Summary Statistics:"
                       Estimate Std. Error t value
 ## (Intercept) 0.8519093 0.1397386 6.096450 1.222884e-06
 ## avg_apd10
                    57.9861322 27.0534250 2.143393 4.060003e-02
 ## Saving 16 x 8 in image
    Average ETI vs. Actual TI for APD-10 using LM Infant Model
                                                                                   F3
      y = 1.1 + 0.01 x
                                      y = 0.99 + 0.064 x
                                                                      y = 0.93 + 0.14 x
                                                                                                      Sequence Fragment
Estimated time since Infection _{
m S}
                                                                                                      ■ F3
                                                                                                      Patient
                                                                                                        pt108
                                                                                                        pt116
pt127
                                                                                                        pt135
                                                                                                        pt170
pt258
                                                                                                        pt313
                                                                                                        pt45
pt485
pt490
                                                                                                        pt93
    Actual time since infection
Average ETI vs. Actual TI for APD-10 using LM Infant Model
                                                                                                      Sequence Fragment
Estimated time since Infection
                                                                                                      ■ F3
                                                                                                      Patient
                                                                                                      pt108
                                                                                                      pt116pt127
                                                                                                      pt135
                                                                                                      pt170pt258
                                                                                                      • pt313
                                                                                                      pt45pt485pt490
                                                                                                      • pt93
```

Saving 16 x 8 in image

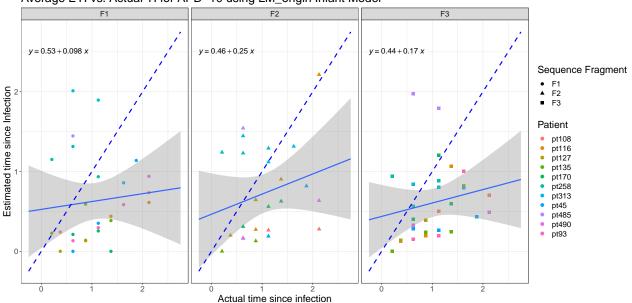
Actual time since infection

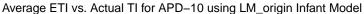


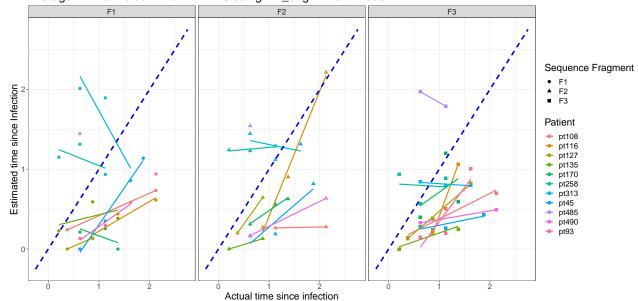


```
## [1] "LM_origin Model for APD10 and Fragment F1 Summary Statistics:"
             Estimate Std. Error t value
                        24.45261 4.927385 4.070286e-05
## avg_apd10 120.4874
## [1] "LM_origin Model for APD10 and Fragment F2 Summary Statistics:"
             Estimate Std. Error t value
##
                                              Pr(>|t|)
## avg_apd10 124.4912
                        21.50463 5.789045 6.741141e-06
## [1] "LM_origin Model for APD10 and Fragment F3 Summary Statistics:"
             Estimate Std. Error t value
                                              Pr(>|t|)
##
                         26.9682 6.683442 2.100821e-07
## avg_apd10 180.2404
```

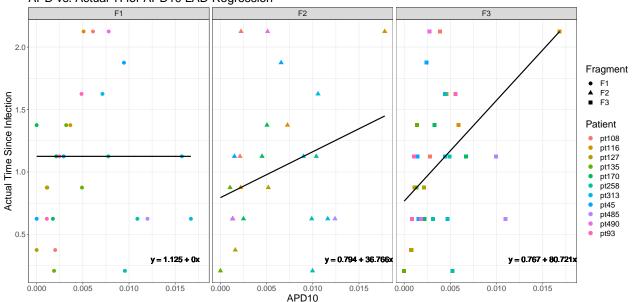
Average ETI vs. Actual TI for APD-10 using LM_origin Infant Model



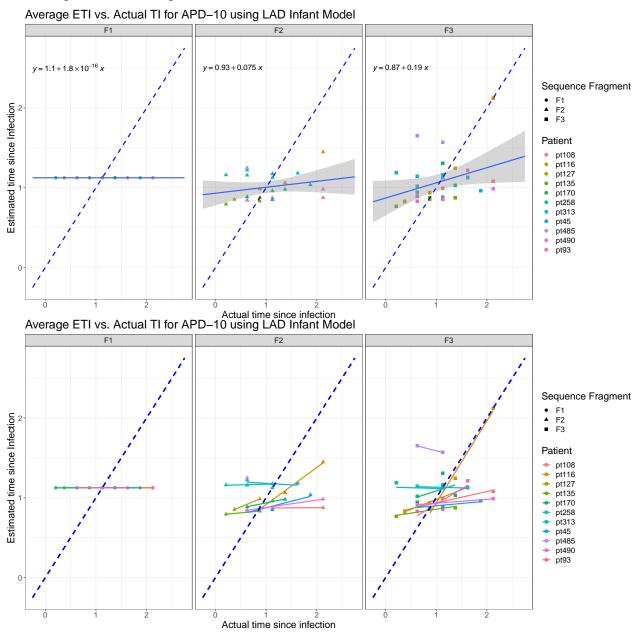




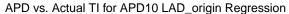


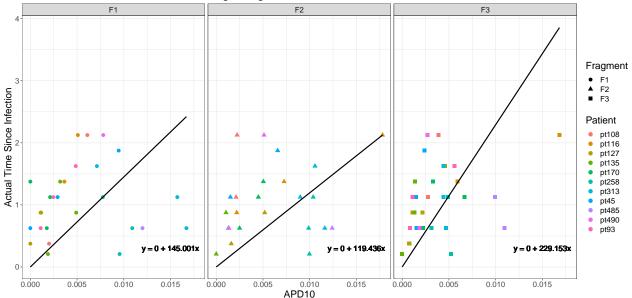


```
[1] "LAD Model for APD10 and Fragment F1 Summary Statistics:"
              Estimate Std.Error Z value
                                             p-value
                 1.125 0.2719231 4.1372 3.5157e-05
## (Intercept)
                 0.000 39.0536182 0.0000 1.0000e+00
## avg_apd10
  [1] "LAD Model for APD10 and Fragment F2 Summary Statistics:"
##
               Estimate Std.Error
                                     Z value
                                                 p-value
## (Intercept) 0.794464 0.2896571 2.7427742 0.006092256
              36.765869 38.9540844 0.9438258 0.345258683
## avg_apd10
## [1] "LAD Model for APD10 and Fragment F3 Summary Statistics:"
               Estimate Std.Error Z value
## (Intercept) 0.766699 0.2127834 3.603190 0.0003143358
## avg_apd10
              80.721062 41.1949189 1.959491 0.0500553444
```



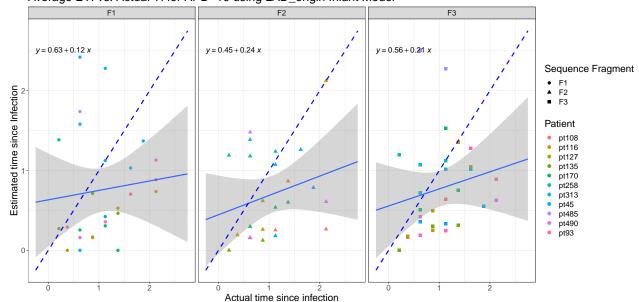
Saving 16 x 8 in image

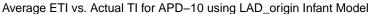


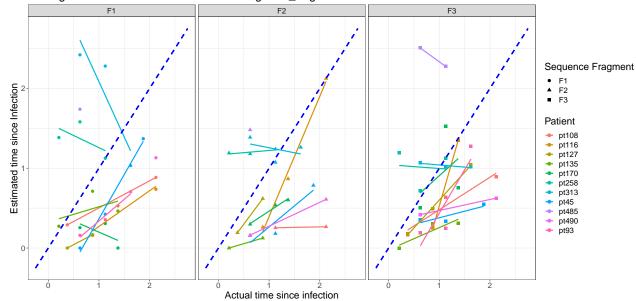


```
## [1] "LAD_origin Model for APD10 and Fragment F1 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd10 145.0006 42.65267 3.399566 0.0006749289
## [1] "LAD_origin Model for APD10 and Fragment F2 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd10 119.4363 34.54921 3.45699 0.0005462442
## [1] "LAD_origin Model for APD10 and Fragment F3 Summary Statistics:"
## Estimate Std.Error Z value p-value
## avg_apd10 229.1527 43.69364 5.244531 1.5668e-07
```

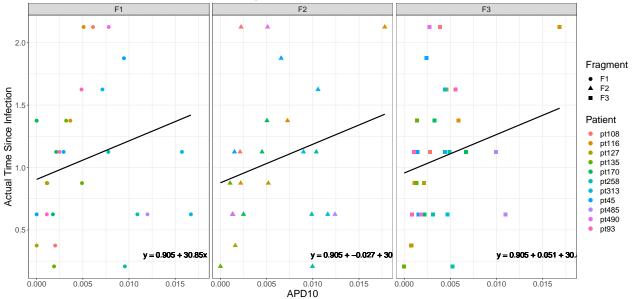
Average ETI vs. Actual TI for APD-10 using LAD_origin Infant Model











```
## [1] "LM_GEE Model for APD10 Summary Statistics:"
##
                         Estimate Naive S.E.
                                                Naive z Robust S.E.
## (Intercept)
                       0.90492138
                                  0.1318987
                                              6.8607305
                                                          0.08967801
## avg_apd10
                      30.85048485 14.6829337
                                              2.1011118 22.73931647
## factor(fragment)F2 -0.02729840
                                   0.1565716 -0.1743509
                                                          0.05492514
## factor(fragment)F3 0.05088403
                                   0.1480864 0.3436104
                                                          0.03893931
##
                        Robust z
## (Intercept)
                      10.0907840
## avg_apd10
                       1.3567024
## factor(fragment)F2 -0.4970111
## factor(fragment)F3
                      1.3067523
##
          (Intercept)
                               avg_apd10 factor(fragment)F2
```

0.000000 0.1748758 0.6191812 ## factor(fragment)F3 ## 0.1912968

