Additional MIMIC Preprocessing

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1 Previously preprocessed data

We begin by examining the data that was previously preprocessed. The data frame is titled df and is provided in the file data.Rdata. It is a long file with 22 variables and 1,276 unique patients.

For each 90 min patient's period, there are 60 values for the numerics, corresponding to the *observation* period. The objective is to predict a hypotensive event in the prediction period by using only the previous values of the observation period after a 30 minute gap period.

- Patients are denoted by thier subject_id.
- Baseline variables include bmi, age, gender, sapsi_first, sofa_first, care_unit, admission_type_descr, los_icu, and los_hospital.
- Time-varying treatment variables include the following binaries amine, sedation, ventilation.
- Time-varying "numerics" variables include time_and_date, hr, abpsys, abpdias, abpmean, and spo2.
- Time-varying outcome variable is **event** and is defined as an acute hypotensive episode during the patient's prediction period.

Here is how the gap period appears in the data.

```
## 3021798 20 1 60 F 76 20 8 34.22847
## 302180 20 2 1 F 76 20 8 34.22847
## 302180 20 2 1 F 76 20 8 34.22847
## 302180 4 ELECTIVE 1 5 0 0
## 302180 ventilation time_and_date hr spo2 absys abpdias abpmean ## 302180 1 2567-03-30 90.46559 80 97 128.6 59.8 78.8
## 302180 1 2567-03-30 10:17:59 80 96 109.3 54.6 68.7
## 3021788 0 1 2567-03-30 10:17:59 80 96 109.3 54.6 68.7
```

Later, we will examine the data frame numerics, which is provided in the file data_numerics.Rdata. The numerics data does not contain this gap period (i.e., the 30 minutes of missing data is available)

1.1 Missing outcome and erroneous variable values

First let's summarize each variable in df.

```
## Skim summary statistics
## n obs: 7126365
## n variables: 22
    -- Variable type:character
    n min max empty n_unique
65 6 9 0 4
    p100
    26711
       Variable type:numeric -----variable missing complete n mean abpdias 525 7125840 7126365 58.36
                                                            sd p0
18.7 -22.8
23.84 0.1
34.41 0
17.14 0
                                                                           49.6
69.2
102.3
           abpmean
abpsys
                        525
525
                              7125840 7126365 83.12
7125840 7126365 119.16
                               7126365 7126365
                                                   65.45
                                                                                     67
            amine
                              7126365 7126365
                                                    0.51
                                                             0.5
                              4228009 7126365 117.48 1389.46
7126365 7126365 0.18 0.38
7125840 7126365 85.27 17.44
               bmi 2898356
                                                                      1.97 24.23 28.15
0 0 0
            event
                         525
                                                                             73.5
                                                                                     84.8
    los_hospital 238637
                               6887728 7126365
                                                   21.25
                              7126365 7126365
6008479 7126365
7126365 7126365
                                                             0.5
         sedation
                                                   0.43
       sofa_first 435318
                              6691047 7126365
                                                   7.48
                                                             4.12
                                                  84.93
30.5
0.27
                              7125840 7126365
7125840 7126365
     p75
67
     ventilation
                          0 7126365 7126365
              p100
346.2
                            hist
     91.6
               439.6
     138.5
     33.86 22436.29
     96
28
15
               198
     19
                38
               22
100
      45.25
     -- Variable type:POSIXct -
    variable missing complete n min max mediar
time_and_date 525 7125840 7126365 2500-10-27 3501-01-01 3013-08-01
     n_unique
     3807554
```

We see that the outcome of interest abpmean is missing for 525 observations.

The following erroneous values can be seen from the summary above:

- maximum bmi of 22436.3 and minimum bmi of 4.04
- maximum age of 200

I removed the 525 rows with missing values for abpmean.

1.2 Subject IDs representing multiple patients

Next, for every subject, I calculated the number of distinct levels for every baseline covariate. Each subject should have 1 distinct level for each baseline covariate.

```
## # A tibble: 406 x 10

## subject_id gender age bmi care_unit admission_type_- los_icu

## cint> cint> cint> cint> cint> cint> cint>

## 1 1 124 1 3 3 2 2 2 3 3

## 2 138 1 2 3 2 2 2 3

## 3 177 1 2 1 3 1 4
```

There are 406 patients with more than one distinct level for each baseline covariate. How does this appear in the data?

```
gender age
M 71
M 75
M 70
M 71
   124
                                                                                            NA
   124
124
   124
                                                                                            NA
   124
                                          75
70
                                                             NA
                                                                                            NA
   124
                                                                                2 22.13825
unit admission
                        type_descr los_icu
                                                                                   sedation
MICU
                         EMERGENCY
CSRII
                                 < N A >
CSRU
                                <NA>
                                                                     NA
CSRII
                         EMERGENCY
                                                                     21
       on time_and_date hr spo2
0 3297-08-03 13:57:12 51.1 98.0
                                                                           0.0
                                                                                      41.2
                                                            0.0
0.0
102.5
                                                                                      41.2
41.2
70.9
       0 3297-08-03 13:57:12 51.1 98.0
                                                                           0.0
       0 3297-08-03 13:57:12 51.1 98.0
0 3297-08-03 13:57:12 51.1 98.0
0 3297-08-03 13:58:12 51.4 97.5
0 3297-08-03 13:58:12 51.4 97.5
                                                            102.5
        0 3297-08-03 13:58:12 51.4 97.5
```

It seems like this subject id is representing more than one patient. However, the outcome values are the same across these seemingly different subjects. Because I do not know which covariate information corresponds to the outcome measurements, I removed all subject id's with multiple baseline covariate values.

After removing these 406 subject ids, which appear to represent multiple patients, 870 subjects remained.

1.3 Outcome measurement error

What does it mean when abpmean has a value, but abpsys and abpdias are both zero? See below.

```
subject_id periode time gender age sapsi_first sofa_first
## 3021795
                            20
20
20
                                               27
28
29
                                                               76
76
76
76
                                                                                                   8 34.22847
                                                                                 20
## 3021733
## 3021783
## 3021771
## 3021925
                                                                                 20
20
20
20
                                                                                                   8 34.22847
8 34.22847
                            20
                                                                                                    8 34.22847
## 3021982
                            20
                                               42
                                                               76
                                                                                 20
                                                                                                   8 34.22847
##
## 3021795
                               admission_type_descr los_icu
ELECTIVE 1
                       CSRU
CSRU
## 3021783
                                                 ELECTIVE
## 3021703
## 3021771
## 3021925
## 3021982
                       CSRU
                                                 ELECTIVE
                                           time_and_date hr
##
## 3021795
                 ventilation
                               on time_and_date hr spo2 abpsys abpdias
1 2567-03-30 09:13:59 80 99.9 0 0
                                                                                                   abpmean
224.5
                                                                                                0
    3021783
3021771
                               1 2567-03-30 09:14:59 80 99.8
1 2567-03-30 09:15:59 80 100.0
## 3021925
## 3021982
                               1 2567-03-30 10:25:59 80 95.0
                               1 2567-03-30 12:28:59 80 100.0
##
## 3021795
## 3021783
## 3021771
    3021771
3021925
## 3021982
```

There are 65,022 rows where this is the case. Here's how this looks in the data.

```
## 3021802 20 1 25 F 76 20 8 34.22847 ## 3021781 20 1 28 F 76 20 8 34.22847 ## 3021795 20 1 27 F 76 20 8 34.22847 ## 3021795 20 1 28 F 76 20 8 34.22847 ## 3021783 20 1 28 F 76 20 8 34.22847 ## 3021781 20 1 29 F 76 20 8 34.22847 ## 3021771 20 1 29 F 76 20 8 34.22847
```

```
## 3021772
                              20
                                                  30
                                                                                                         8 34.22847
##
## 3021802
## 3021794
                 care_unit admission_type_descr los_icu los_hospital amine
CSRU ELECTIVE 1 5 0
CSRU ELECTIVE 1 5 0
## 3021795
## 3021783
                         CSRU
                                                    ELECTIVE
                         CSRII
                                                    ELECTIVE
## 3021773
## 3021771
## 3021772
                         CSRU
                                                    ELECTIVE
                                 time_and_date hr
1 2567-03-30 09:11:59 80
1 2567-03-30 09:12:59 80
                                                                        spo2
100.0
100.0
                   ventilation
## 3021802
## 3021794
                                                                                                             72.3
73.2
                                  1 2567-03-30 09:13:59 80
## 3021795
## 3021783
                                                                        99.9
                                                                                     0.0
                                                                                                  0.0
                                                                                                            224.5
                                 1 2567-03-30 09:14:59 80
                                                                         99.8
                                                                                      0.0
                                                                                                   0.0
                                                                                                            169.9
## 3021771
## 3021772
                                 1 2567-03-30 09:15:59 80 100.0
1 2567-03-30 09:16:59 80 100.0
## 3021802
## 3021794
## 3021795
## 3021783
## 3021771
```

The above abpmean values (with abpsys and abpdias both zero) seem to be inconsistent with the other abpmean readings.

However, in other cases (like below), these odd abpmean values are consistent with the normal abpmean readings.

```
## 6654207
## 6654208
                                                                   age
52
52
                                                                          sapsi_first sofa_first
                                                   15
16
                                                                                                           1 24.42046
1 24.42046
## 6654209
## 6654210
## 6654211
## 6654226
                                                                                                            1 24,42046
                              79
                                                   17
                                                                    52
                                                    18
                                                                     52
                                                                                                            1 24,42046
                              79
79
79
79
                                                    19
20
                                                                     52
52
                                                                                                            1 24.42046
1 24.42046
## 6654227
                                                   21
                                                                     52
                                                                                                            1 24.42046
## 6654228
                              79
79
                                                   22
23
                                                                     52
                                                                                                            1 24.42046
## 6654239
##
                                                                     52
                                                                                                               24.42046
                  care_unit admission_type_descr los_icu los_hospital amine sedation
## 6654207
                         CSRU
                                                    EMERGENCY
## 6654208
## 6654209
## 6654210
                         CSRU
                                                    EMERGENCY
                                                    EMERGENCY
                          CSRU
## 6654211
                         CSRU
                                                    EMERGENCY
                                                                                                           0
## 6654226
## 6654227
                         CSRU
                                                   EMERGENCY
EMERGENCY
## 6654228
                         CSRU
                                                    EMERGENCY
## 6654239
                         CSRU
                                                    EMERGENCY
## 6654239
## ## 6654207
## 6654208
## 6654210
## 6654211
                                 on time_and_date hr
0 2756-08-13 16:45:00 102.8
0 2756-08-13 16:46:00 102.7
                                                                                                                     83.2
82.1
                                                                                98.4
99.0
                                                                                           0.0
97.0
                                                                     98.3 99.5
99.3 100.0
100.3 99.3
                                 0 2756-08-13 16:47:00
                                                                                            0.0
                                                                                                          0.0
                                                                                                                     78.4
                                 0 2756-08-13 16:48:00
0 2756-08-13 16:49:00
## 6654226
## 6654227
                                 0 2756-08-13 16:50:00 100.8
                                                                               99.1
                                                                                             0.0
                                                                                                          0.0
                                                                                                                     77.1
                                 0 2756-08-13 16:51:00 101.7
                                                                                98.0
                                                                                           94.2
                                                                                                        68.5
                                                                                                                     79.2
## 6654228
## 6654239
                                 0 2756-08-13 16:52:00 104.4
0 2756-08-13 16:53:00 101.5
##
## 6654207
## 6654207
## 6654208
## 6654210
## 6654211
## 6654226
## 6654227
## 6654228
## 6654239
```

To solve this issue, I calculated subject specific abpmean outlier thresholds as a way to determine if these odd abpmean values are outliers or not.

It's important to note that the subject specific distributions of abpmean values did consider the missing data in the gap period. Section 2.4.1 resolves this issue.

For 2 subjects (ids 25373 and 26209), all of the outcome measurements contain zeros for abpsys and abpdias and values for abpmean, so the outlier thresholds could not be calculated. These two subjects were removed from the data. Now we can see the outliers.

Of the 65,022 rows with this odd outcome measurement, 31,209 rows were deemed outliers and were subsequently removed.

Now 868 subjects remain in the data.

1.4 Missing baseline characteristics

There are still missing values in the data.

##	subject_id	periode	time
##	0	0	0
##	gender	age	sapsi_first
##	- 0	0	55185
##	sofa_first	bmi	care_unit
##	44167	678023	0
##	admission_type_descr	los_icu	los_hospital
##	13671	0	13671
##	amine	sedation	ventilation
##	0	0	0
##	time_and_date	hr	spo2
##	0	0	0
##	abpsys	abpdias	abpmean
##	0	0	0
##	event		
##	0		

Removing all NA values would lead to the omission of 308 subjects, but 258 of these subjects were only missing bmi and no other value. Many subjects had very odd bmi values such as 4, 5, 9, 69, 77, and 22436. Because of this oddity, I only removed subjects that were missing values for other covariates.

818 subjects remain in the data.

1.5 Insufficient patient data

As described in the beginning, the objective is to predict a hypotensive event in the prediction period (i.e., after 90 minutes) using only the values in the observation period (i.e., the first 60 minutes) after a 30 minute gap period.

We need each patient to have data in the prediction period, so we can evaulate the predictions with their actual value. Thus, we need each subject to have data for over 90 minutes to ensure that the data extends into the prediction period.

However, of the 818 remaining subjects, 21 had less than 90 minutes of data available. These subjects were removed from the data.

797 subjects remain in the data.

1.6 Classifying a hypotensive event

The variable **event** is defined as an acute hypotensive episode during the patient's prediction period. In some cases, it appears that a hypotensive event is not classified as one. In other cases, it appears that a non-hypotensive event is misclassified as a hypotensive event. Both scenarios are shown below.

I created a new outcome Y1 using a function Ivana made. This function classifies an event as hypotensive when the current abpmean is less than 62 and at least 5 adjacent *time points* have abpmean less than 65. Adjacent time points are typically 1 minute apart. There is an exception for time points that occur 5 minutes before gap periods and 5 minutes after gap periods. These 5 minutes before and after gap periods consider adjacent time points that are separated by 30 minutes.

See Section 2.4.1 for a resolution to this issue.

We can examine outcomes Y1 and event below.

```
## 86449
## 86450
## 86451
## 86452
                                                                                            14 19.67777
                                    18
                                                                             22
                       906
                                    18
                                                           78
                                                                             22
                                                                                            14 19,67777
                                                                                            14 19.67777
14 19.67777
                                           11
## 86453
                       906
                                    18
                                           12
                                                           78
                                                                             22
                                                                                            14 19,67777
## 86449
## 86450
                    MICU
                                           EMERGENCY
## 86451
## 86452
## 86453
                    MICU
                                           EMERGENCY
                                           EMERGENCY
##
## 86449
             ventilation
                                       time_and_date
                           0 2653-04-21 02:50:25
                                                            95.6
                                                                                         47.9
                                                                                                      60.0
    86450
86451
                              2653-04-21 02:51:25 103.0
2653-04-21 02:52:25 99.8
2653-04-21 02:52:25 97.6
## 86452
                            0 2653-04-21 02:53:25
## 86453
                           0 2653-04-21 02:54:25
## 86450
## 86451
## 86452
## 86453
                                                      age
76
76
## 86
## 87
                    20
20
                                                                        20
20
                                       30
                                                                                          8 34.22847
## 88
## 89
## 90
                                       31
                                                       76
                                                                        20
                                       32
33
                    20
## 91
                                       34
35
                                                      76
                                                                        20
                                                                                          8 34.22847
## 92
                    20
                                                      76
                                                                        20
                                                                                          8 34.22847
## 87
                CSRU
                                        ELECTIVE
                CSRU
                                        ELECTIVE
```

```
ELECTIVE
## 92
##
## 86
                            ELECTIVE
on time_and_date hr
1 2567-03-30 10:45:59 80
          CSRU
ventilation
## 87
## 88
## 89
## 90
## 91
## 92
##
                                                                                   118.7
                            1 2567-03-30 10:46:59 80
                                                                        98.5
                                                                                                   59.8
                                                                                                                 75.9
                            1 2567-03-30 10:47:59 80 99.9
1 2567-03-30 10:48:59 80 100.0
1 2567-03-30 10:49:59 80 100.0
                                                                                  140.3
154.2
                             1 2567-03-30 10:50:59 80 100.0
                                                                                                               106.9
                            1 2567-03-30 10:51:59 80 100.0
## 86
## 87
## 88
## 89
```

1.7 Summary

This updated data frame with 797 subjects is named mimic and does not contain

- NA values for any variable except bmi,
- outcome measurement error outliers,
- single subject id's which represent multiple patients,
- or patients with less than 90 minutes of data.

This mimic data frame does contain

- the gap period of 30 minutes after every hour of data;
- a new hypotensive event outcome Y1 which classifies a hypotensive event as one when the current abpmean is less than 65 and at least 5 adjacent time points have abpmean less than 65;
- and subject's with erroneous baseline characteristic measurements, including
 - $subject_id = 20936$ with age of 200,
 - and many subjects with odd bmi values such as 4, 5, 9, 77, 69, 22436.

The mimic data is used in the Section 2. If you would like to work with data with gap periods, then use the mimic_gap data, which is presented in Section 2.4.1.

2 Data numerics

There is another a data frame numerics saved in data_numerics.Rdata. This data frame contains the outcome data for each subject with no gap periods.

2.1 Omitting 30 minute gap periods

I merged the 30 minute gap period into the mimic data that was created in the previous section. I filled in this missing 30 minutes of data because it is preferred for the simulation, since it provides a more clear representation of the patient data.

The only variables in numerics data frame are subject_id, time_and_date, hr, abpsys, abpdias, abpmean, and spo2.

Before merging the data, I removed any erroneous/outlier outcome measurements according to the method described in Section 1.3.

Here's how the data looks after merging.

```
hr abpsys abpdias abpmean spo2 periode
3.4 105.5 34.8 54.8 92.9 1
5.7 106.4 35.3 55.8 92.6 1
4.0 104.8 34.7 54.6 92.8 NA
                                  time_and_date
                10013 2564-11-02 03:24:18 93.4
10013 2564-11-02 03:25:18 95.7
10013 2564-11-02 03:26:18 94.0
## 61
##
                10013 2564-11-02 03:27:18 93.7
                                                               104.1
                                                                            34.3
                                                                                         53.9 92.0
                                                                                                               NA
                          ## 58
## 59
                                                                8 34.85214
## 60
           NA
                   <NA>
                           NA
                                             NA
                                                              NA
                                                                           NA
                                                                                        <NA>
                   <NA> NA
                                                                                        <NA>
         admission_type_descr
                                                             spital
## 58
## 59
                         EMERGENCY
                        EMERGENCY
## 60
## 61
                                               NA
NA
                                                                        <NA>
## 58
          <NA> NA
```

2.2 Fill in missing values

We can see that missing values need to be filled in for

- 1. the time-varying treatments
- 2. and the baseline covariates.

I filled in the missing baseline covariate information with tidyr::fill. This function fills missing values using the previous entry, so we assume that the 30 minute gap period had the same baseline covariate values as the minute *before* this gap started. This procedure is surely reasonable for the baseline covariates, but probably not for the time-varying treatment, which is why we didn't fill in that information. Let's see how the data shown above looks after filling in this information.

```
## 58
## 59
## 60
## 61
                     20 2567-03-30 09:47:59 80
                                                          129.3
                                                                      60.1
                                                                                 79.2
                                                                                          97
                     20 2567-03-30 09:48:59 80
                                                          128 7
                                                                      59 9
                                                                                 78.8
                                                                                          97
97
97
                                                                                                      NA
                     20 2567-03-30 09:49:59 80
20 2567-03-30 09:50:59 80
                                                                                 78.9
78.9
                                                          129.0
                                                                      60.1
 ##
## 58
## 59
                time
                                                                      bmi ca
                                                             8 34.22847
8 34.22847
 ## 60
## 61
                           76
                                            20
                                                             8 34.22847
            NA
                           76
                                            20
                                                             8 34.22847
                                                                                  CSRU
                                                          spital amine
5 <NA>
5 <NA>
 ## 59
## 60
                          ELECTIVE
                                                                                  <NA>
                                                                                                  <NA>
                          FLECTIVE
                                                                    < N A >
                                                                                 <NA>
                                                                                                  < N A >
## 58 <NA> NA
## 59 <NA> NA
## 60 <NA> NA
## 61 <NA> "'
```

There are some cases when missing values needed to be filled in using a *later* entry, because there are no prior entries without NA. This is the case for 173 subjects. Here's an example of how it looks.

```
## 22919
## 22920
## 22921
## ## 22919
                                                                                              spo2
99.9
                                                                                    bpmean
110.0
                     439 3242-12-28 15:07:01 105.8 166.6
439 3242-12-28 15:08:01 106.7 166.4
                                                                            72.2
                                                                            71.2
                                                                                     109.5 100.0
                      439 3242-12-28 15:09:01 107.1
                       time gender age sapsi_first sofa_first NA \, NA \, NA \, NA \, NA
                                                                            NA
## 22920
                   NA
                        NA
                                <NA> NA
                                                         NA
                                                                        NA
                                                                             NA
                                                                                        <NA>
## 22921
                                       83
                                                         33
                                                                        18
                                                                                         CCU
             admission_type_descr los_icu los_hospital
                                                                             sedation ventilation
## 22919
                                                                 NA
NA
15
                                  <NA>
                                               NA
                                                                      <NA>
## 22920
                                  <NA>
                                               NA
7
                                                                      <NA>
                                                                                  <NA>
                                                                                                   <NA>
## 22921
##
## 22919 <NA> NA
## 22920 <NA> NA
## 22921 0 0
```

The column periode refers to the "number of the patient's period". For these 173 subjects, the minimum periode is always 2 or higher. Thus, the merge filled in something like periode 1 for these subjects.

There was probably a reason for omitting periode 1 for these subjects, maybe the time-varying treatment data was unavailable. I can remove periode 1 from these subjects later if need be.

Something else I noticed about periode when examining this issue. Below I listed all of the unique periode available for the subject above, subject_id = 439.

```
## [1] NA 2 3 4 5 6 6 7 8 12 13 14 16 17 18 19 20 21 ## [18] 22 3 24 25 26 27 28 29 30 31 33 34 36 37 38 39 40 ## [38] 41 42 44 45 46 47 48 49 50 15 52 53 64 55 56 57 58 ## [52] 59 60 61 62 63 64 65 66 67 68 69 70 71 72 73 74 76 ## [58] 77 78 79 80 81 82 83 84 85 68 78 88 89 90 91 29 39 ## [88] 94 95 96 97 98 99 100 101 102 103 104 105 106 107 108
```

Interestingly, this subject is missing some of the periode (e.g., 9-11). The data that would have been periode 9-11 is available in numerics, but this information not included in the preprocessed data df. Perhaps because the time-varying treatment data was not available.

2.3 Classifying a hypotensive event

Like before, I created a new outcome Y1 to classify hypotensive events. The issue with adjacent time points occasionally being 30 minutes apart is not present in this scenario, since there is no gap period.

2.4 Summary

This updated data frame with the full data still contains 797 subjects and is named mimic_nogap. This updated data does not contain

- NA values for any baseline characteristic variable except bmi,
- outcome measurement error outliers,
- single subject id's which represent multiple patients,
- patients with less than 90 minutes of data,
- or the gap period of 30 minutes after every hour of data.

This mimic_nogap data frame does contain

- a new hypotensive event outcome Y1 which classifies a hypotensive event as one when the current abpmean is less than 65 and at least 5 adjacent time points have abpmean less than 65;
- subject's with erroneous baseline characteristic measurements, including
 - $subject_id = 20936$ with age of 200,
 - and many subjects with odd bmi values such as 4, 5, 9, 77, 69, 22436;
- NA values during the merged in 30 minute gap period for
 - time-varying treatments amine sedation ventilation,
 - and periode time event.

The mimic_nogap data is saved in mimic_nogap.Rdata.

2.4.1 Important Note

Because mimic_nogap considered the full data, two of the preprocessing steps in mimic_nogap are more reliable than those in mimic.

1. The new hypotensive event outcome Y1 relies on adjacent time points for classification. Since mimic_nogap does not contain 30-minute gaps with no data, the adjacent time points are closer together in mimic_nogap.

2. There are many instances when abpmean has a value, but abpsys and abpdias are both zero. Sometimes these oddities appear to be consistent with the patients "normal" readings (i.e., readings which have non-zero values for abpmean, abpsys and abpdias). In other cases, these oddities are very different from the patients normal readings. I remove the inconsistent oddities by calculating subject specific abpmean outlier thresholds, which requires calculating subject specific IQRs. These IQRs are more accurate in mimic_nogap, since the full data is used.

For these reasons, I used mimic_nogap to recreate the same gaps that are in mimic. I called this data frame mimic_gap and saved it in mimic_gap.Rdata.

I recommend using the mimic_gap data instead of mimic data.