Predictive Modeling of Mice Behavior through Neural Activity

2022/4/6

Introduction

Cruz-Martin, Alberto from the Center for Systems Neuroscience at Boston University is dedicated to unraveling the mysteries of how mice's brain circuits control their behaviors. The central puzzle revolves around understanding the roles of specific cell types and their contributions to the visual processing and perception in mice. Recognizing the complexity of this endeavor, Alberto sought our expertise in data analysis and prediction.

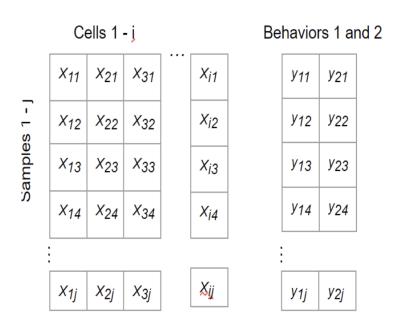
Objective

Our primary objective is to analyze the data from the Zero Maze experiment and predict the behaviors of mice based on their cell activation status.

Data Collection and Methodology

Dataset Structure

Alberto's team tested 13 mice in the Zero Maze experiment. Data for each mouse is stored in distinct datasets. For every 10 HZ interval during the 10-minute experiment, data points are recorded, detailing the behavior of the mouse and the pertinent cell information.



Data Features

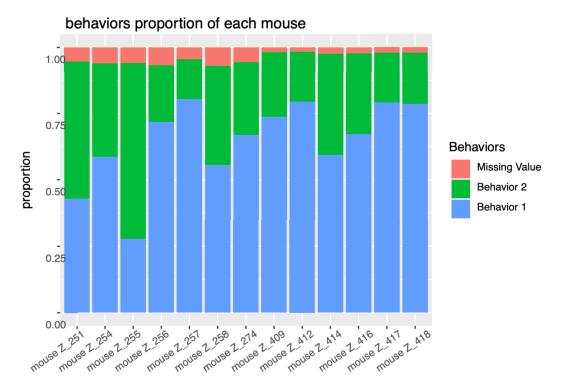
Location of Mice: The dataset identifies the mouse's location with two distinct columns. A '1' in a column indicates the mouse's presence in the respective arm, while a '0' denotes its absence. Notably, during times when a mouse is transitioning between arms, both columns reflect a '0'. After a comprehensive discussion with Alberto, we've decided to treat these instances as edge cases, excluding them from our analysis. This simplifies our interpretation, allowing us to view the mice behaviors as binary.

Cell Activation: The dataset records the calcium flow amount for each mouse cell as a measure of its activation, with all values being positive. It's crucial to note that the specific cells recorded vary for each mouse. For instance, the data documents 112 cells for mouse Z409, whereas only 28 cells are recorded for mouse Z416.

Armed with this data, we aim to establish a clear link between cell activation and mice behavior, delving deep into the intricacies of neuroscience.

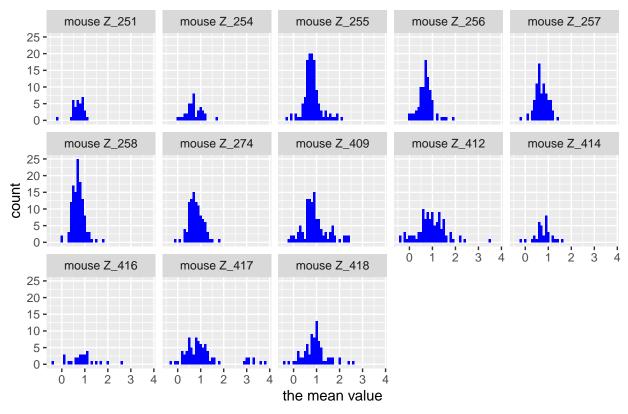
Exploratory Data Analysis

Then we explore the proportion of two types of behaviors of each mouse. We can see that most mice conduct behavior 1 at least 50% of the time except for mice Z251 and Z255. Mouse Z257, Z417, and Z418 even conduct behavior 1 about 75%. We also notice the existence of missing values where both behaviors are recorded as 0, but there are not many of them.

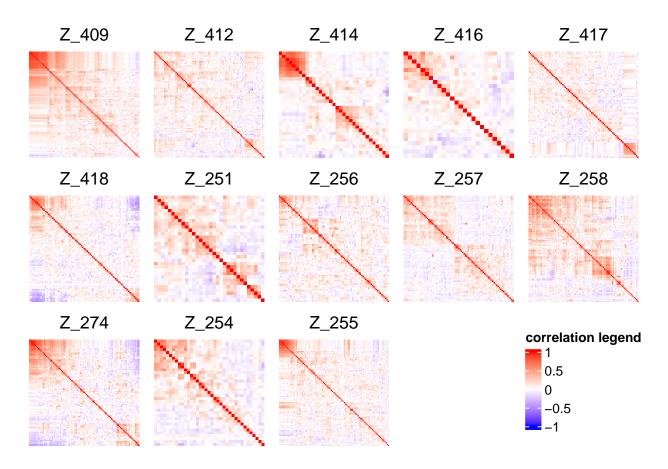


This graph shows us the distribution of the mean value of calcium transition in cells of each mouse. As we can see, most cells have an average value between 0 to 2. Moreover, most of them are t distributed except for the mice Z_416 and Z_251, the two mice that have fewer data (25 cells and 34 cells, respectively).

the distributions of mean value of calcium in cells of each mouse



In this heat map, we plot 13 mice. The heat map illustrates the correlations of cells of each mouse, as we can see that there exist some strong correlations between certain cells. Thus we need to check correlations in detail before constructing any models.



PCA

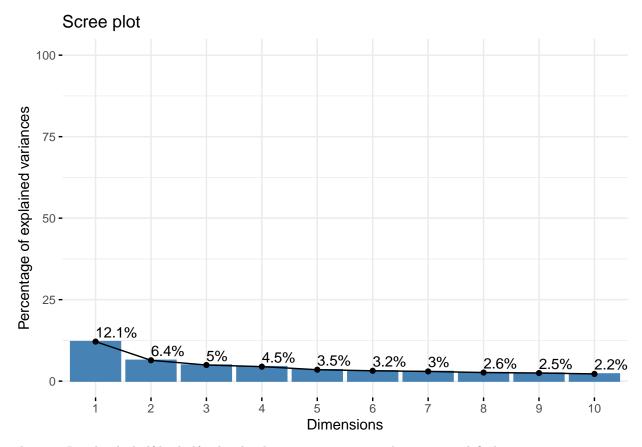
According to the correlation map, we found there is a high correlation between variables, so we considered conducting PCA. At first, we calculated VIF values. Large VIF suggests predictors almost completely explained by the other variables in the equation. The VIF values we calculate for Mouse 255 of the Zero Maze study are as follows:

```
##
          Х1
                     X2
                               ХЗ
                                          Х4
                                                     Х5
                                                               Х6
                                                                          Х7
                                                                                    Х8
##
    4.443693 32.799587 11.122254
                                    3.406167
                                              5.946836
                                                         2.626223
                                                                    5.854133
                                                                              4.427542
##
          Х9
                    X10
                              X11
                                         X12
                                                    X13
                                                              X14
                                                                         X15
                                                                                    X16
## 10.108495 18.600526
                         5.731881 30.105783 40.978332 17.056840
                                                                   3.807708
                                                                              5.079167
##
         X17
                    X18
                              X19
                                         X20
## 16.882012 3.652280 14.019159
                                   7.706484
```

We find strong colinearity in our predictors, so we conduct PCA and then choose the first 20 PCs as new predictors.

Training model

The scree plot shows the tenth dimension only explained the 2.2% information of the data.



Then we Randomly half-by-half split the data into train set and test set and fit logistic regression again using the 20 principle components. Then we calculate the VIF values again, and we found there is no strong colinearity.

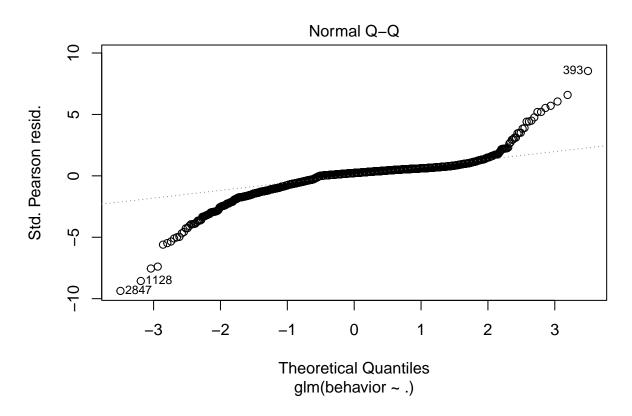
```
##
               Dim.2
                         Dim.3
                                  Dim.4
                                            Dim.5
                                                                        Dim.8
      Dim.1
                                                     Dim.6
                                                               Dim.7
## 1.236405 1.602410 1.588525 1.500733 1.707871 1.514087 1.558070 1.329879
##
      Dim.9
              Dim.10
                        Dim.11
                                 Dim.12
                                           Dim.13
                                                    Dim. 14
                                                              Dim.15
                                                                       Dim. 16
## 1.505733 1.354067 1.481117 1.202244 1.306605 1.205972 1.202887 1.314359
     Dim.17
              Dim.18
                        Dim.19
                                 Dim.20
## 1.173397 1.080312 1.141499 1.163684
```

Then we did the prediction on the test set using the threshold of 0.5 and got a result of about 0.83 accuracies (0.81, 0.85), 0.8 recall, and 0.9 specificities. The ROC plot also shows that the best threshold may be around 0.3, where we can have both high recall and specificity.

```
##
  Confusion Matrix and Statistics
##
##
      р
##
          0
                1
##
        444
             158
##
     1
        161 1356
##
                   Accuracy: 0.8495
##
                     95% CI: (0.8335, 0.8644)
##
       No Information Rate: 0.7145
##
##
       P-Value [Acc > NIR] : <2e-16
##
                      Kappa: 0.6305
##
```

```
##
    Mcnemar's Test P-Value : 0.9108
##
##
               Sensitivity: 0.7339
##
##
               Specificity: 0.8956
            Pos Pred Value: 0.7375
##
##
            Neg Pred Value: 0.8939
                Prevalence: 0.2855
##
##
            Detection Rate: 0.2095
##
      Detection Prevalence: 0.2841
##
         Balanced Accuracy: 0.8148
##
##
          'Positive' Class : 0
##
```

The QQ plot shows that the residual follows a normal distribution.



Model for the whole dataset

Finally, based on the model building in the training set and test by test set, we fit the model again in the whole dataset.

```
##
## Call:
## glm(formula = behavior ~ ., family = "binomial", data = new_predictors1)
##
## Deviance Residuals:
```

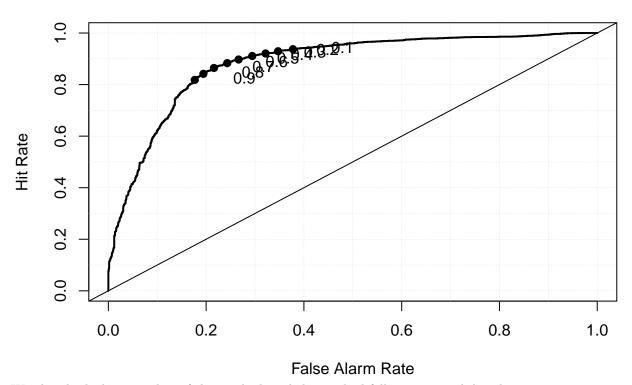
```
Median
##
       Min
                 10
                                    3Q
                                            Max
## -3.2308
           -0.4357
                      0.3118
                                0.6522
                                         3.1049
##
## Coefficients:
##
               Estimate Std. Error z value Pr(>|z|)
## (Intercept)
                1.42835
                           0.05539
                                     25.786
                                            < 2e-16 ***
## Dim.1
               -0.12589
                           0.01220 -10.317 < 2e-16 ***
## Dim.2
                0.19036
                           0.02455
                                      7.754 8.88e-15 ***
## Dim.3
                0.26253
                           0.02606
                                     10.075
                                            < 2e-16 ***
## Dim.4
               -0.12443
                           0.02910
                                     -4.276 1.90e-05 ***
## Dim.5
                0.76711
                           0.03514
                                     21.830
                                            < 2e-16 ***
## Dim.6
               -0.40561
                           0.03277 -12.376 < 2e-16 ***
## Dim.7
               -0.24334
                           0.03134
                                     -7.764 8.24e-15 ***
## Dim.8
                0.39169
                           0.02877
                                     13.614 < 2e-16 ***
## Dim.9
                0.12280
                           0.03444
                                      3.565 0.000364 ***
## Dim.10
                0.22221
                           0.03562
                                      6.238 4.42e-10 ***
## Dim.11
                0.33747
                           0.03580
                                      9.427 < 2e-16 ***
## Dim.12
                0.15524
                           0.03040
                                      5.106 3.28e-07 ***
## Dim.13
                0.48362
                           0.03400
                                     14.223 < 2e-16 ***
## Dim.14
                0.04326
                           0.03325
                                      1.301 0.193205
## Dim.15
                0.02575
                           0.03400
                                      0.757 0.448877
## Dim.16
               -0.28052
                           0.03960
                                     -7.083 1.41e-12 ***
## Dim.17
               -0.22576
                           0.03353
                                     -6.733 1.67e-11 ***
## Dim.18
               -0.03125
                           0.03342
                                     -0.935 0.349739
## Dim.19
               -0.28814
                           0.03842
                                     -7.500 6.37e-14 ***
## Dim.20
                0.14868
                           0.03965
                                      3.750 0.000177 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Dispersion parameter for binomial family taken to be 1)
##
##
       Null deviance: 5140.9
                              on 4237
                                        degrees of freedom
## Residual deviance: 3311.6
                             on 4217
                                        degrees of freedom
  AIC: 3353.6
## Number of Fisher Scoring iterations: 6
```

The confusion matrix shows the accuracy of the model for the whole dataset is 0.85 (0.8376, 0.8594) and with a Sensitivity of 0.74 and Specificity of 0.88.

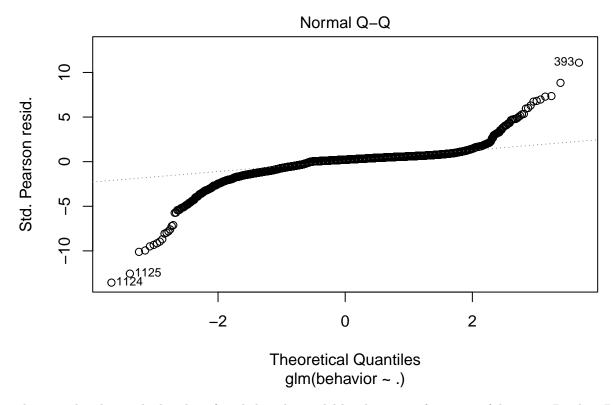
```
## Confusion Matrix and Statistics
##
##
      р
##
          0
               1
##
        916
             334
        307 2681
##
     1
##
##
                  Accuracy : 0.8487
##
                     95% CI: (0.8376, 0.8594)
       No Information Rate: 0.7114
##
##
       P-Value [Acc > NIR] : <2e-16
##
##
                      Kappa: 0.634
##
##
    Mcnemar's Test P-Value: 0.3044
```

```
##
               Sensitivity: 0.7490
##
               Specificity: 0.8892
##
##
            Pos Pred Value : 0.7328
            Neg Pred Value: 0.8973
##
##
                Prevalence: 0.2886
##
            Detection Rate: 0.2161
      Detection Prevalence : 0.2950
##
##
         Balanced Accuracy: 0.8191
##
          'Positive' Class : 0
##
##
```

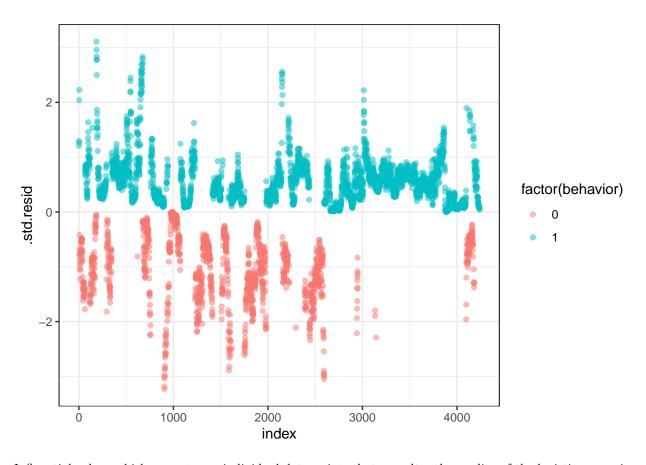
ROC Curve



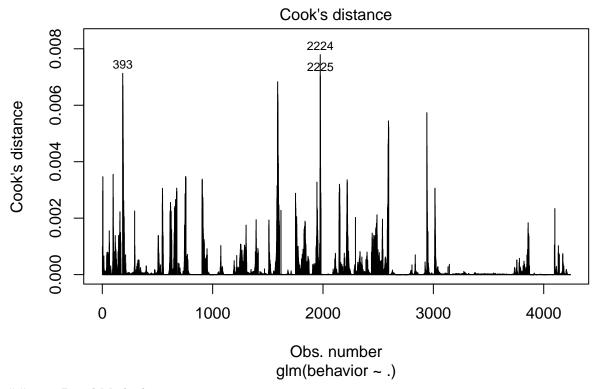
We also check the normality of the residual, and the residual follows a normal distribution.



Then we plot the residual and we found that the model has better performance of detecting B1 than B0.



Influential values which are extreme individual data points that can alter the quality of the logistic regression model. The most extreme values in the data can be examined by visualizing the Cook's distance values. Here we label the top 3 largest values.



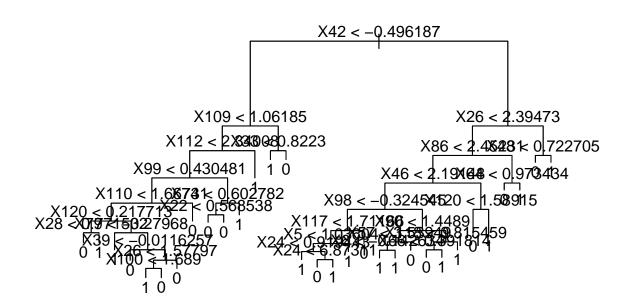
Tree-Based Method

We extract 20 PC to fit the logistic regression, while the price for that is that we lose some useful signal information and noise since the top 20 PC only explained an accumulative 61% of the data, and we only get the 83% accuracy. Considering this data includes many cells, we do not need to make dimensionality reduction before the random forest—the tree-based method without distribution restrictions.

Binary Decision tree

Therefore, we use a classification tree to improve our accuracy of prediction. We put all variables in the training set and fit a binary decision tree.

```
##
## Classification tree:
## tree(formula = behavior ~ ., data = rf[train, ])
  Variables actually used in tree construction:
   [1] "X42" "X109" "X112" "X99"
                                    "X110" "X120"
                                                                        "X26"
   [11] "X100" "X74"
                      "X22"
                             "X33"
                                    "X86"
                                           "X46"
                                                  "X98"
                                                         "X117" "X5"
                                                                        "X24"
## [21] "X90"
              "X57"
                      "X6"
                             "X118" "X66"
## Number of terminal nodes: 31
## Residual mean deviance: 0.2537 = 529.7 / 2088
## Misclassification error rate: 0.042 = 89 / 2119
```



The predicted result shown by the confusion matrix, the accuracy is 0.89(0.88, 0.90). And the Sensitivity is 0.840, and the Specificity is 0.9164.

```
Confusion Matrix and Statistics
##
##
##
          0
               1
##
        528
              79
        112 1400
##
##
##
                  Accuracy : 0.9099
##
                    95% CI: (0.8969, 0.9217)
##
       No Information Rate: 0.698
       P-Value [Acc > NIR] : < 2e-16
##
##
##
                     Kappa: 0.783
##
##
    Mcnemar's Test P-Value: 0.02059
##
##
               Sensitivity: 0.8250
##
               Specificity: 0.9466
##
            Pos Pred Value: 0.8699
##
            Neg Pred Value: 0.9259
                Prevalence: 0.3020
##
##
            Detection Rate: 0.2492
##
      Detection Prevalence: 0.2865
##
         Balanced Accuracy: 0.8858
```

Random Forest

A random forest is the average of many different single trees that only contain different subsets of original predictors. So a random forest will typically be helpful when we have many correlated predictors. We use a random forest model, which can be a universal method for all 13 zero maze data sets. We choose predictor subset size for each data set according to the number of cells. Then we build a reproducible function 'rf' and apply it to all 13 data sets.

This is the confusion matrix for Mouse 255, this is one result of our function.

```
## Confusion Matrix and Statistics
##
##
  yhat.rf
##
                   1
##
         0
            586
                  10
             54 1469
##
##
##
                  Accuracy : 0.9698
##
                    95% CI: (0.9616, 0.9767)
##
       No Information Rate: 0.698
##
       P-Value [Acc > NIR] : < 2.2e-16
##
##
                     Kappa: 0.9269
##
    Mcnemar's Test P-Value: 7.658e-08
##
##
##
               Sensitivity: 0.9156
               Specificity: 0.9932
##
##
            Pos Pred Value: 0.9832
            Neg Pred Value: 0.9645
##
##
                Prevalence: 0.3020
##
            Detection Rate: 0.2765
##
      Detection Prevalence: 0.2813
##
         Balanced Accuracy: 0.9544
##
##
          'Positive' Class: 0
##
```

Result for 13 mice in random forest function:

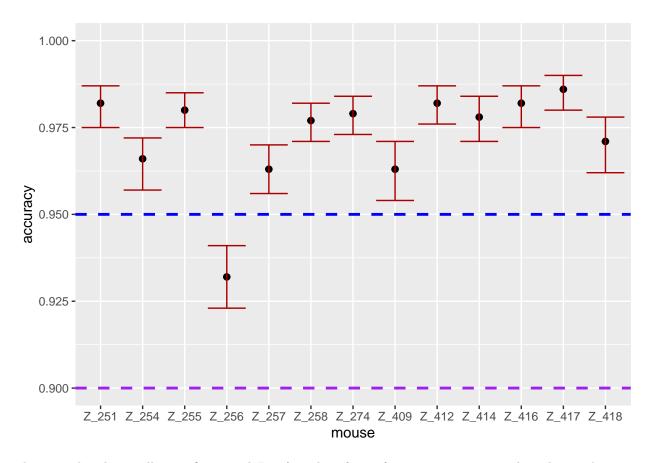
mouse	accuracy low	er_boundip	per_bound	record	specificity	F1
Z_409	0.963	0.954	0.971	0.954	0.970	0.956
Z_412	0.982	0.976	0.987	0.991	0.969	0.986
Z_414	0.978	0.971	0.984	0.942	0.994	0.963
Z_416	0.982	0.975	0.987	0.996	0.935	0.988
Z_417	0.986	0.980	0.990	0.998	0.914	0.992
Z_418	0.971	0.962	0.978	0.993	0.937	0.976
Z_251	0.982	0.975	0.987	0.993	0.956	0.987

mouse	accuracy low	er_boundip	per_bound	record	specificity	F1
Z_254	0.966	0.957	0.972	0.996	0.874	0.976
Z_255	0.980	0.975	0.985	0.999	0.901	0.988
Z_256	0.932	0.923	0.941	0.976	0.863	0.946
Z_257	0.963	0.956	0.970	0.990	0.904	0.974
Z_258	0.977	0.971	0.982	0.995	0.902	0.986
Z_274	0.979	0.973	0.984	0.998	0.896	0.987

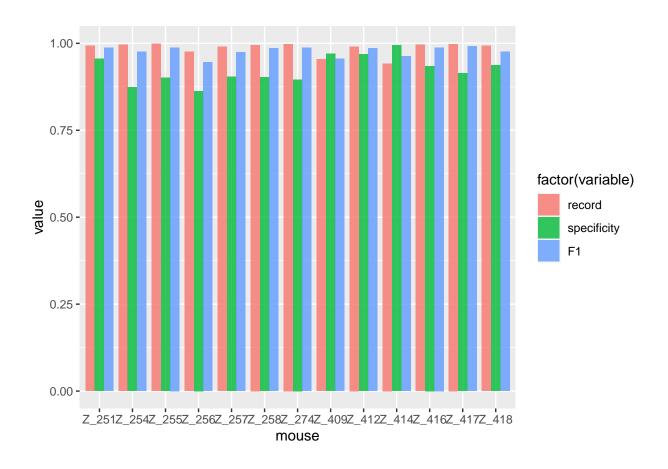
The average accuracy of random forest for 13 mice is 97.23%.

##	mouse	accuracy	lower_bound	upper_bound
##	Length:13	Min. :0.9320	Min. :0.9230	Min. :0.9410
##	Class :character	1st Qu.:0.9660	1st Qu.:0.9570	1st Qu.:0.9720
##	Mode :character	Median :0.9780	Median :0.9710	Median :0.9840
##		Mean :0.9724	Mean :0.9652	Mean :0.9783
##		3rd Qu.:0.9820	3rd Qu.:0.9750	3rd Qu.:0.9870
##		Max. :0.9860	Max. :0.9800	Max. :0.9900
##	record	specificity	F1	
##	Min. :0.9420	Min. :0.8630	Min. :0.9460	
##	1st Qu.:0.9900	1st Qu.:0.9010	1st Qu.:0.9740	
##	Median :0.9930	Median :0.9140	Median :0.9860	
##	Mean :0.9862	Mean :0.9242	Mean :0.9773	
##	3rd Qu.:0.9960	3rd Qu.:0.9560	3rd Qu.:0.9870	
##	Max. :0.9990	Max. :0.9940	Max. :0.9920	

We plot the accuracy with a 95% confidence interval. The dot in the figure is the accuracy for each mouse, and the red line corresponds 95% confidence interval. The blue dashed line is baseline at 0.95, while the purple dashed line is 0.9. We can see almost mice got an accuracy of more than 95%, and all mice had more than 90% accuracy.

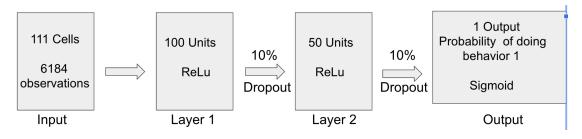


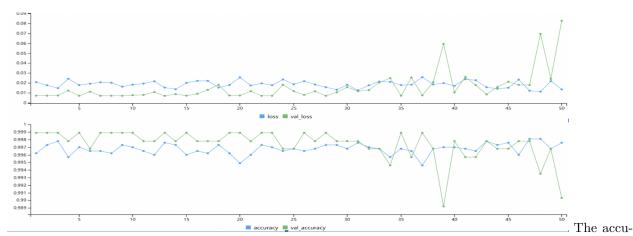
Then we plot the recall, specificity, and F1 of random forest for 13 mice, respectively. These values are generally close to 1, which means our model has excellent performance.



Multilayer Neural Network

We also have built a simple neural network model to predict the mice's behaviors. In the model, we construct two hidden layers, which contain 100 units and 50 units respectively, and both with ReLu activation function. In the output layer, we use the Sigmoid activation function to show the probability of whether the mouse is conducting behavior 1. The following graph shows the model structure of mouse Z409 as an example.





racies of our Neural Network models are very high; most of them are around 97%. We set the epochs to be 50, batch size to be 32, and validation set to be random 20% of the data, and we plot the model accuracy of mouse Z409 as an example below.

Conclusion

There exist correlations between many cells of each mouse, including all the cells as predictors in a model is not wise, dimension reduction is necessary.

Compared with the logistic_PCA model, the random forest classification model not only is easier to visualize and interpret, but also provides a higher prediction accuracy that can even compete with the Neural Network model.

Discussion

For our logistic-PCA model, we select the first 20 principal components as predictors. The choice of this number is not rigorously verified, so 20 may not be the optimal choice. And in the random forest, we pick the predictor subset size to be the square root of the number of all predictors, this is a common convention, and it is not guaranteed to be the optimal choice. In the neural network model, the number of hidden layers and units in each layer are also not verified to be the optimal choice. In the validation process, since there are spatial and time relationships between each row of the data, randomly selecting observations to form the validation set may not be the best choice.