Predicting Cognitive Decline

Group 8: Brett Castro, Luke Napolitano, Sam Remmey, Lucy Shichman, Sanjana Srivastava



Introduction

- Alzheimer's Disease: Progressive neurodegenerative disease that deteriorates memory and other important mental functions
 - Highly prevalent in patients 85+
 - Earlier detection allows for earlier treatment
 - Clinical diagnostic tools are expensive and invasive
- Research Questions
 - O How can we use health indicators to predict cognitive decline?
 - Mini-Mental State Exam (MMSE) via MLR
 - Diagnosis via Logistic Regression
- Dataset summary:
 - Health information of 2149 patients
 - O Symptoms, demographics, medical assessments, and medical history
 - 14 numerical, 22 categorical variables



Healthy Brain



Mild Alzheimer's Disease

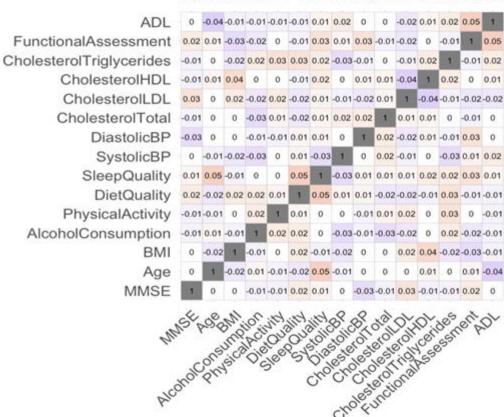


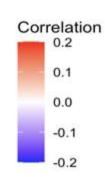
Severe Alzheimer's Disease

Exploratory Data Analysis - Numerical

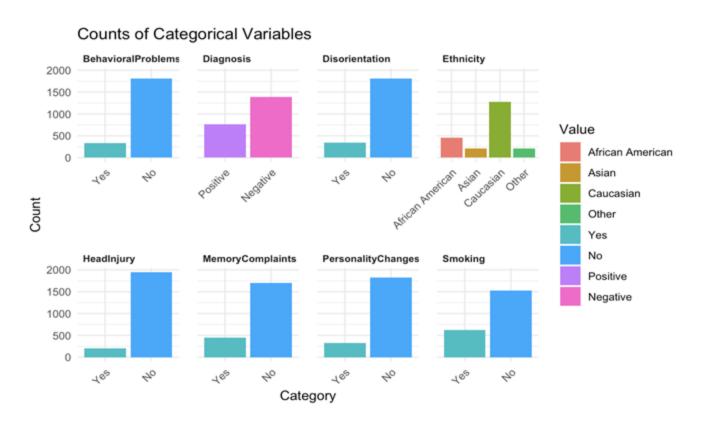
- Low correlations
- Strongest correlation: 0.05

Correlation Plot of Numeric Variables





Exploratory Data Analysis - Categorical

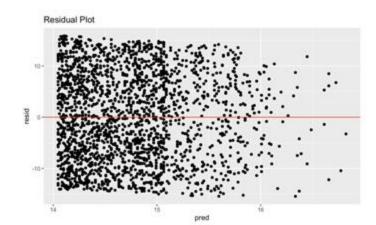


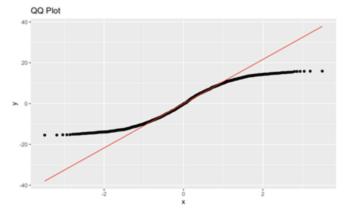
- Key categorical variables
- More negative diagnoses
- Majority white patients

MLR: How can we use health indicators to predict MMSE score?

- Building the model:
 - checking assumptions
 - log transformations
 - trimming influential/leverage points
 - o stepAIC
 - o lasso

- o P-value = 0.03919, Adjusted R^2 = 0.002528, Residual SE = 8.533
- Cross-validation
 - o 5 folds, 10 repetitions
 - o RMSE = 8.539161, $R^2 = 0.004267931$
- Prediction Interval
 - O Given a diastolic BP of 72, no personality changes, and no disorientation, we are 95% confident that MMSE score is between -211.6598 and 34.95187
 - o actual: 21.46353





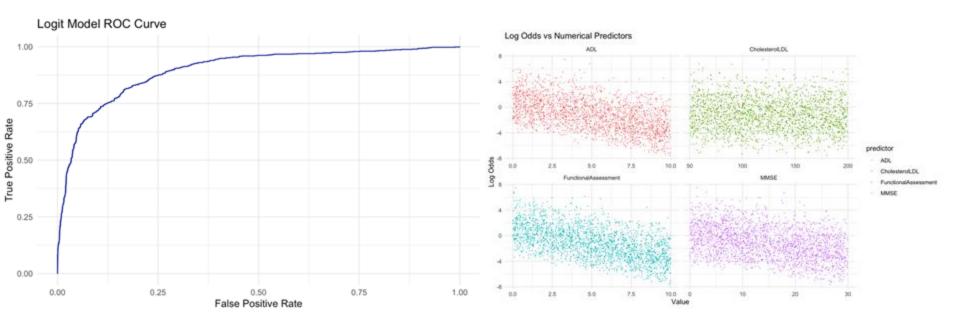
Logistic Regression: How can we use health indicators to predict Alzheimer's diagnosis?

- Building the model
 - Including MMSE, trying to predict diagnosis
 - No transformations
 - Reducing model to only include predictors significant at α =0.05
 - stepAIC (AIC:1835.6 -> AIC:1615), reducing information loss
 - O Lasso: kept same variables as stepAIC model

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Log odds = 4.268 - 0.003(CholesterolLDL) - 0.107(MMSE) - 0.445(FunctionalAssessment) + 2.595(I_MemoryComplaintsYes) + 2.487(I_BehavioralProblemsYes) - 0.416(ADL)
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- O AUC = 0.903, Accuracy = 0.841, F-1 score = 0.798
- Cross-validation
 - o 10 folds
 - o AIC = 1617.2
- Prediction
 - O Based on this model, the predicted probabilities of a positive diagnosis are **0.396** and **0.929** for two different patients who have an actual diagnosis of negative and positive, respectively.

Logistic Regression: How can we use health indicators to predict Alzheimer's diagnosis?



AUC = 0.903, Accuracy = 0.841, F-1 score = 0.798

Conclusion & Future Work

- We found that the logistic model predicting diagnosis was far more useful than the multilinear regression model predicting MMSE score
- Other explorations:
 - predicting MMSE:
 - Random Forest: 8.207 RMSR, 9.17% variance explained
 - o predicting diagnosis:
 - Support Vector Machine: Overall accuracy (94%)
- Developing non-invasive alzheimer's diagnostic modeling:
 - Machine Learning with clinical data
 - NLP with speech testing data



Research Citation: Moradi, E., Prakash, M., Hall, A., Solomon, A., Strange, B., Tohka, J., & Alzheimer's Disease Neuroimaging Initiative (2024). Machine learning prediction of future amyloid beta positivity in amyloid-negative individuals. Alzheimer's research & therapy, 16(1), 46. https://doi.org/10.1186/s13195-024-01415-w

Thank You! Questions?

