Imputation of SEER Data using Artificial Neural Networks

A comparison of technique performance

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Outline

Background

- Imputation
- Neural networks
- SEER selection criteria

Performance Analysis

- Overfitting indicators
- Cross validation scores
- Train/Test distribution comparison

Next Steps/Discussion & Questions

- NOS reclassification
- More comparative techniques
- Alternative architectures

Imputation:

- The process of replacing missing data using various techniques
- Missing completely at random (MCAR), missing at random (MAR), and missing not at random (MNAR)
- Assumptions, bias compounding, and effects on statistical power
- Techniques, nested imputation, and multiple imputation

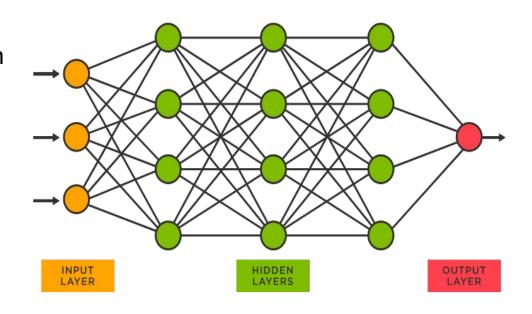


Artificial Neural Networks (ANNs) and Multilayer Perceptrons (MLPs):

- Input data (features) with normalization and embedding.
- Artificial Neurons containing non-linear activation function with weighted input and output edges.
- Output layer maps to categorical classification
- Loss function to train against
- Gradient descent optimization with learning rate factor
- Dropout probability to guard against overfitting

Parameters of MLP used in this project:

- 43 Predictors/features, 15 numerical, 28 categorical embedded with dimensions = ceiling(n/2)
- ReLU activation function, dropout p = 0.4, hidden layer dimensions [200, 100, 50], output layer size =
 4, learning rate = 0.001



SEER Stage Variables:

- "The Staging Over Time Project."
- AJCC 3rd (1988-2004), AJCC 6th (2004-2015), AJCC 7th (2016-2017), AJCC 8th (2018+)
- Target is time merged AJCC with 6th > 7th > 8th hierarchy excluding substages, created by NCI

<u>Selection Criteria (N = 1, 947, 359):</u>

- SEER 18 Incidence (Not delay adjusted)
- 2004-2018
- Non-Cardia Gastric cases
 - Intestinal histologies: <u>8140</u>, 8143-8144, 8210-8211, 8221, 8260-8263
 - Diffuse histologies: 8141-8142, 8145, 8490
- Known age, race, diagnostic confirmation



Predictor Variable Groupings and Stage Missingness:

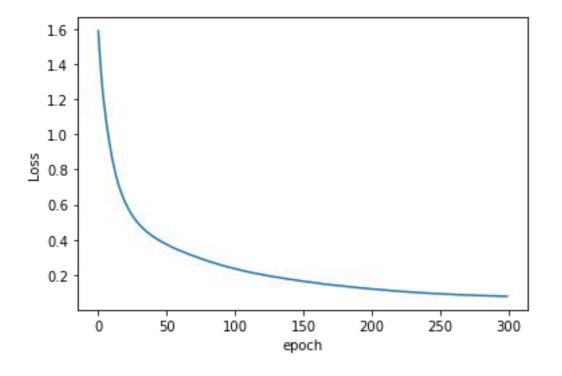
- Demographic (Age, Sex, Race/Origin, Year, n = 4)
- AJCC Stage (AJCC Editions, n = 4)
- Histology (Intestinal vs Diffuse, n = 1)
- Summary Stages (L/R/D, n = 8)
- Extent of Disease (T/N/M, n = 19)
- CS Variables (Tumor size, extension etc, n = 7)

Group Total	Staged Count 1771115	Unstaged Count 176244	% Missing 9.05
Race			
NH White	1234824	117585	8.69
AAPI	122566	13384	9.84
AI/AN	8997	1000	10.00
Hispanic	176240	23396	11.72
NH Black	228488	20879	8.37
Sex			
Female	525981	65870	11.13
Male	1245134	110374	8.14
Histology			
Intestinal	1743261	171850	8.97
Diffuse	27854	4394	13.63

Year	Staged Count	Unstaged Count	% Missing
2004	109140	11724	9.70
2005	108701	10754	9.00
2006	115619	10697	8.47
2007	122315	11462	8.57
2008	121148	11096	8.39
2009	122818	10976	8.20
2010	122087	10032	7.59
2011	124073	10655	7.91
2012	117458	10152	7.96
2013	117226	9921	7.80
2014	116121	9871	7.83
2015	119458	9789	7.57
2016	118719	14183	10.67
2017	120399	15164	11.19
2018	115833	19768	14.58

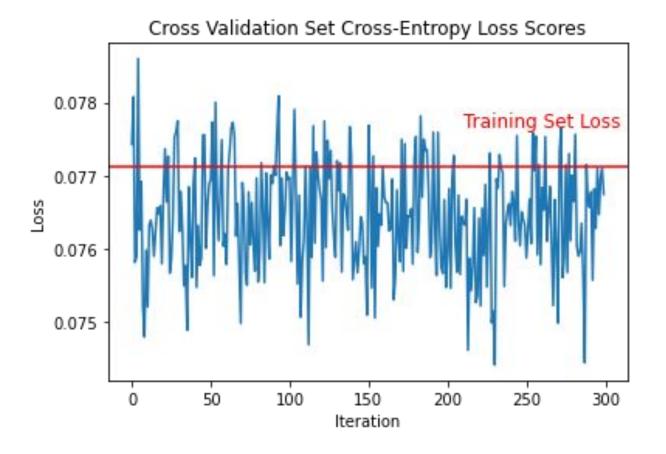
Cross Entropy Loss:

- Cost function, measure of performance
- Training set Cross Entropy =
 0.07712 bits after 300 epochs
- Noisy training curve tail indicates overfitting



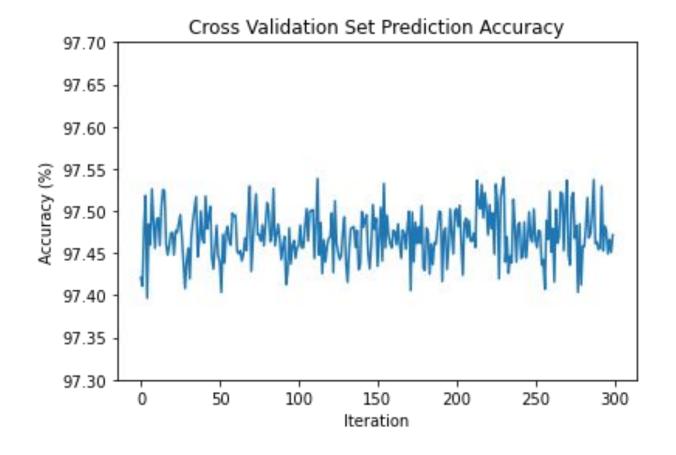
Cross Validation Performance

- Higher Cross Entropy loss indicates overfitting
- Every iteration with the same cross validation set (20% of Train Set)

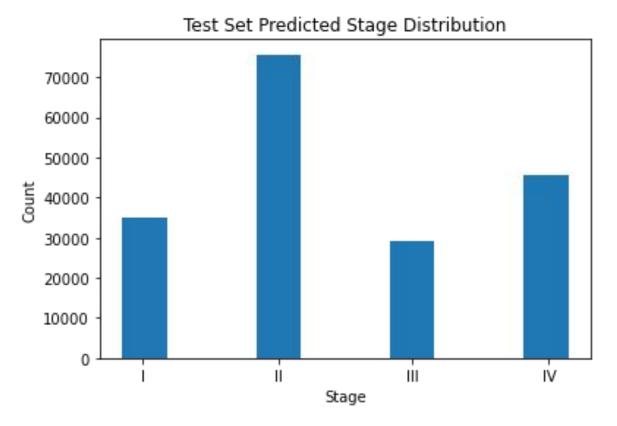


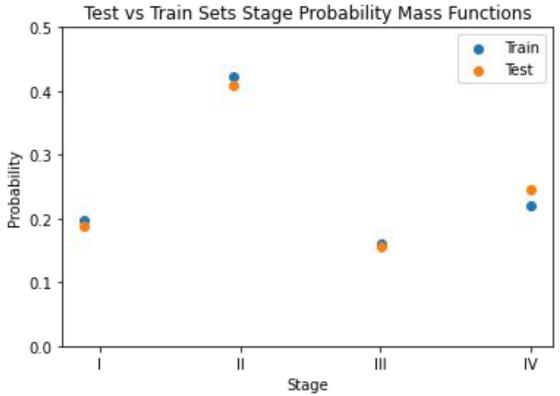
Cross Validation Performance

 Accuracy records the percentage of correctly labelled observations in the Cross Validation set



Test Prediction Distribution





Next Steps

- New Model Architectures
 - Properly encode the time validity of certain predictors
 - Finetune hyperparameters
- Sensitivity Analysis
 - Dimension reduction, pruning predictors
 - Multifold CV scores
- Further Applications
 - Other sites
 - NOS histology reclassification



Closing Discussion

Is this too much of a "black box" technique?

Is this too dangerous given that we are forcing changes to our evidence?

How much uncertainty will this introduce to modelling based on the imputed data?

Additional Questions from the Audience

Citations

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- 3. Marek Smieja, Łukasz Struski, Jacek Tabor, Bartosz Zielinski. Przemysław Spurek, Processing of missing data by neural networks, 32nd Conference on Neural Information Processing Systems (NeurIPS 2018), Montreal, Canada.
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