



Towards Sequential Data Clustering via GBTM and LSTM

Yirui Hu, Ph.D.

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Outline

- Electronic health records (EHR) contain longitudinal data with valuable phenotypic information on disease progression of patients.
 - Weight change trajectories in Roux-en-Y Gastric Bypass (RYGB) patients over 7 years
- Group-based trajectory modeling (GBTM)
 - A specialized application of finite mixture modeling designed to identify clusters of individuals who follow similar trajectories in sequential data
- Long Short Term Memory (LSTM)
 - A special kind of recurrent neural network that can learn the order dependence in sequential data



Weight change trajectories in RYGB patients

ELSEVIER

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Original article

Demographic, clinical, and behavioral determinants of 7-year weight change trajectories in Roux-en-Y gastric bypass patients

Michelle R. Lent Ph.D. ^{a, b} ≈ ⊠, Yirui Hu Ph.D. ^c, Peter N. Benotti M.D. ^a, Anthony T. Petrick M.D. ^a, G. Craig Wood M.S. ^a, Christopher D. Still D.O. ^a, H. Lester Kirchner Ph.D. ^c

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Weight change trajectories in RYGB patients

- To assess long-term weight trajectories in a large cohort of RYGB patients over 7 years
 - Primary outcome: Weight changes (% percent change from baseline) post-operatively over time
- Hypothesis: Patients respond differently to the same surgical intervention
 - Identify heterogenous subgroups
 - Identify high-risks patients / guide clinical care
 - Inform patient-provider conversations about treatment expectations

 $BMI > 40 \text{ kg/m}^2 \text{ or } > 35$ kg/m² + comorbidity

RYGB

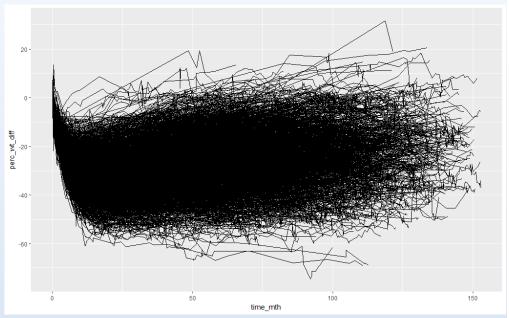
Up to 7 years postsurgery

Weight data in the EMR pre and 7 years postop

Study Population

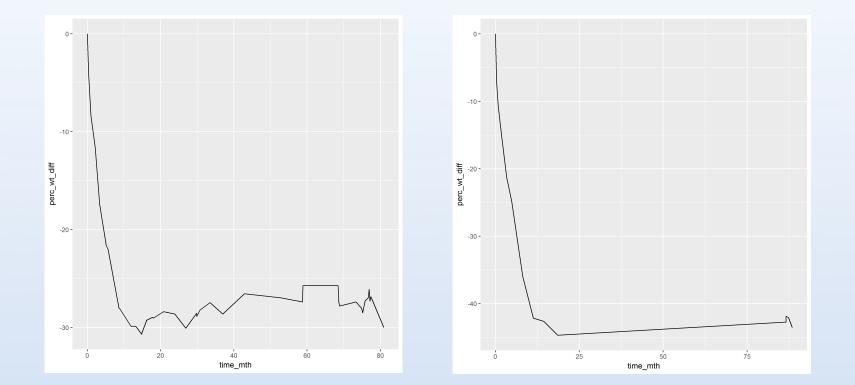
• RYGB bariatric surgery (2004-2016)

- N=3,215
- 80% female, >95% White
- Preoperative BMI = $49.4 \pm 8.8 \text{ kg/m}^2$
- Data visualization
 - X-axis: Months post-surgery
 - Y-axis: Weight changes (% from baseline)



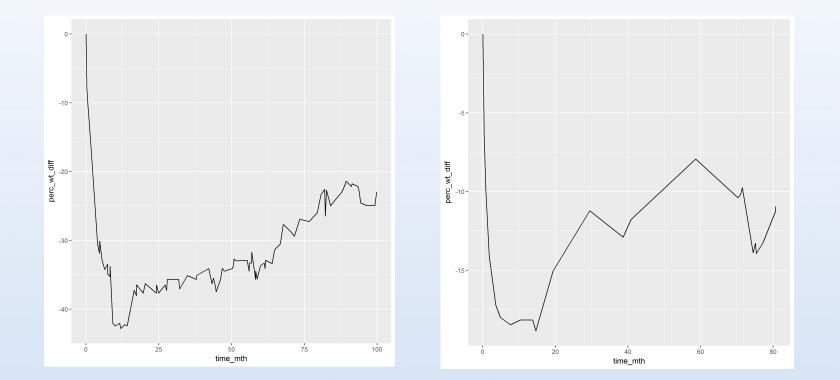
Individual-Level Data Visualization

• Pattern 1: Remain stable in weight change after the first major weight loss



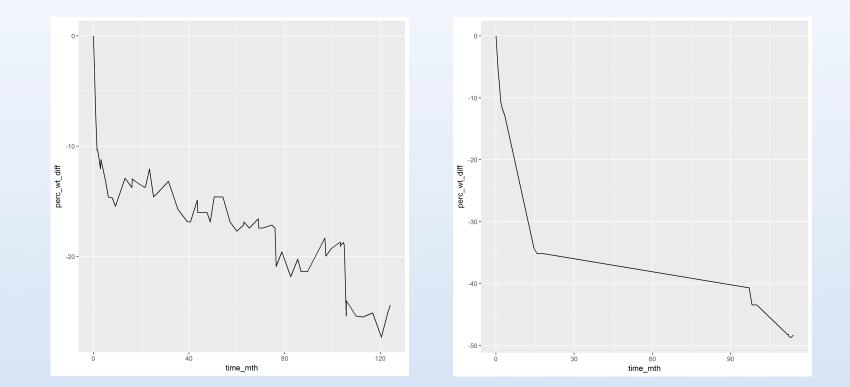
Individual-Level Data Visualization

• Pattern 2: Regain in weight change after the first major weight loss



Individual-Level Data Visualization

• Pattern 3: Lose more in weight change after the first major weight loss



Summary of Individual-Level Patterns

- First stage: major weight loss patterns
 - Almost everyone lose weight to some degree
 - within 12-18 months post-surgery
- Second stage: different weight change patterns (assumes heterogeneous subpopulations)
 - Remain stable -> Average
 - Regain -> Suboptimal
 - Further weight loss -> Optimal

Group-Based Trajectory Modeling (GBTM)

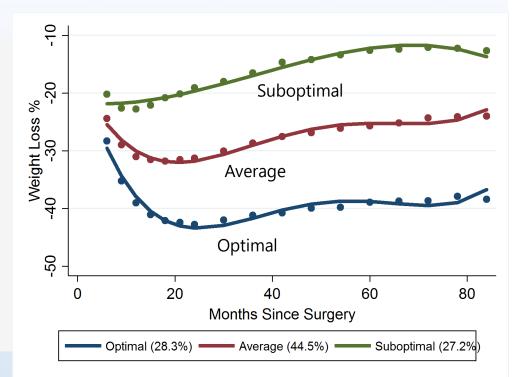
- A statistical method that is designed to identify a finite number of groups of individuals following similar trajectories over time of a single outcome
 - Identify distinctive trajectories: groups of individuals with similar trajectories
 - Estimate the proportion of the population following each such trajectory group
 - Relate group membership probability to individual characteristics
- Two key outputs of the GBTM
 - the shape of the trajectory, typically defined by a polynomial function of time,
 - the probability of trajectory group membership



Summary results of GBTM

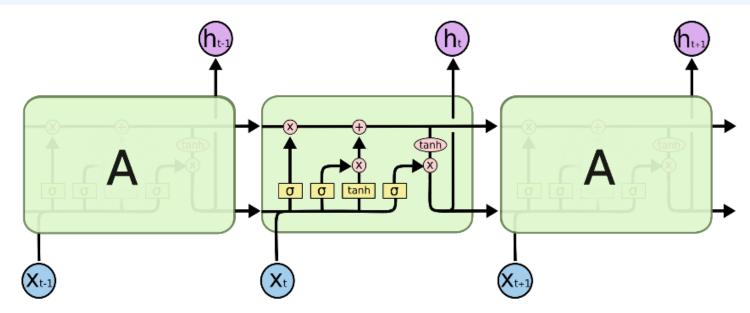
Highlights

- Weight loss trajectories up to seven-years following Roux-en-Y Gastric Bypass (RYGB) were not uniform, with 27% of patients maintaining <20% body weight loss.
- Patients that experienced poorer long-term weight loss were more likely to be male and have diabetes, but less likely to have a smoking history or take sleep medications.
- Lower BMI at the time of surgery and less early postoperative weight loss were associated with poorer weight outcomes.



Long Short Term Memory (LSTM)

- Recurrent Neural Network (RNN) and its special kind, Long Short Term Memory (LSTM) network have successfully tackled many time-relevant problems.
 - LSTM (introduced by Hochreiter & Schemidhuber 1997) is capable of learning long-term dependencies.

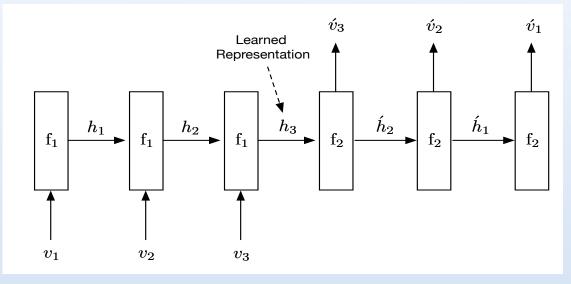


The repeating module in an LSTM contains four interacting layers.

https://colah.github.io/posts/2015-08-Understanding-LSTMs/

LSTM Autoencoder

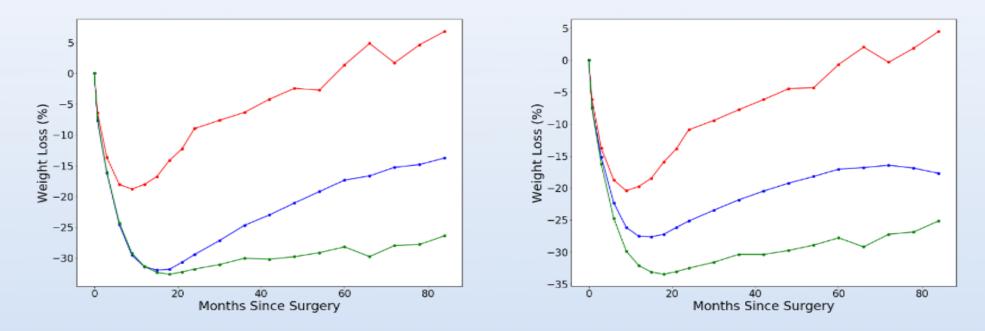
- Autoencoders are a type of self-supervised learning model that can learn a compressed representation of input data.
 - For a given dataset of sequences, an encoder-decoder LSTM is configured to read the input sequence, encode it, decode it, and recreate it. The performance of the model is evaluated based on the model's ability to recreate the input sequence.





Clustering (K-Means and GMM)

Subgroup	Red	Blue	Green
K-means Count (%)	82 (2.9%)	1128 (39.6%)	1641 (57.6%)
GMM Count (%)	63 (2.2%)	438 (15.4%)	2350 (82.4%)



Concluding remarks

- Cluster analysis of longitudinal EHR data
 - Group-based trajectory modeling
 - Long Short Term Memory networks
- Potential applications
 - Identify heterogenous subgroups in longitudinal data
 - Link covariates to identified trajectory membership
 - Relate trajectories to subsequent outcomes in predictive modeling



Our Predictive Modeling Pipeline

Data Preparation

- Import data from EHR
 - Input: features and label
- Impute missing values
 - MICE
- Detect outliers

Model Development

- Feature selection and engineering
- Run algorithms
 - Logistic regression
 - Random forest
 - XGBoost
- Cross-Validation
 - Tune hyperparameters

Model Evaluation

- Performance report
 - AUC, etc.
 - Feature importance
- Risk prediction
 - Probability of developing disease
 - High risk vs. low risk

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 - <u>yhu1@geisinger.edu</u>





- Lent MR, Hu Y, Benotti PN, Petrick AT, Wood GC, Still CD, Kirchner HL. Demographic, clinical, and behavioral determinants of 7-year weight change trajectories in Roux-en-Y gastric bypass patients. Surgery for Obesity and Related Diseases. 2018 Nov 1;14(11):1680-5
- <u>https://colah.github.io/posts/2015-08-Understanding-LSTMs/</u>
- <u>https://machinelearningmastery.com/lstm-autoencoders/</u>



Missing Data Imputation

- In the imputation step, we learned from the series of past observations to predict the next value in the sequence.
- Mean Absolute Error (MAE)
 - MAE of Median imputation: 0.28;
 - MAE of LSTM imputation: 0.003.

