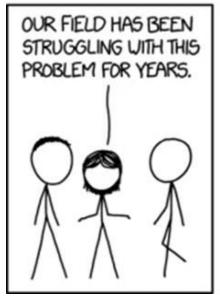
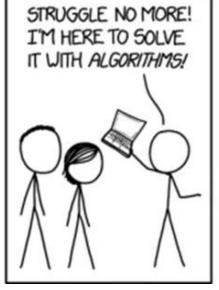


Machine Learning is not Magic









Here to Help: https://xkcd.com/1831

The problem with the "magical" metaphor

Why is Machine Learning So Magical?

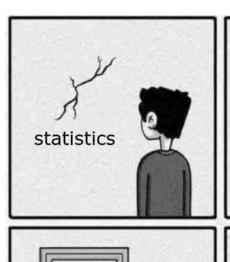
And the struggles of practising the tricks

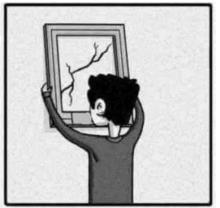


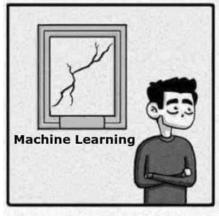


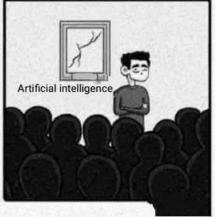


On the flip side, are some too cynical?











Epidemiologists use tools for different purposes

Questionnaire Development

CLINICAL Biological Assays
TEST PROTOCOLS Propens

Assays
Propensity
Scores

EXPOSURE MODELING

Community Engagement

Regression

Machine Learning??

AGENT-BASED MODELS



Expect Magic and You Will Be Disappointed....



Original Investigation | Cardiology

Comparison of Machine Learning Methods With Traditional Models for Use of Administrative Claims With Electronic Medical Records to Predict Heart Failure Outcomes

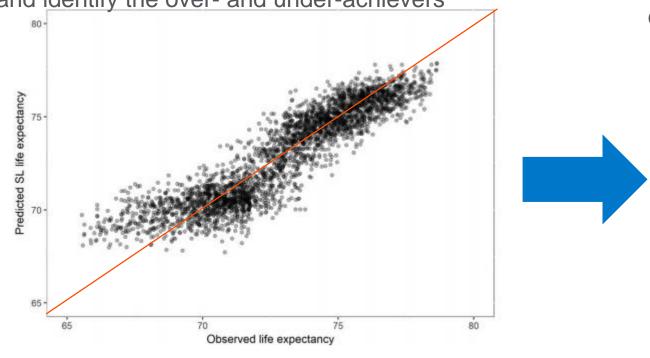
6113 obs in training 3389 obs for testing

54 variables from Medicare claims 8 variables from EHR "In our study, we observed that when using only claims-based predictors, many of which are binary variables indicating presence or absence of medical conditions or use of specific medications, the performance improvement with machine learning approaches was minimal for prediction of most outcomes. However, when the predictor set was expanded to include EMR-based information, which included numerous laboratory test results as continuous variables, we noted that machine learning continuous variables, we noted that machine learning approaches generally fared better than logistic regression. This observation follows the intuition that, because tree-based machine learning approaches, such as GBM or random forests, are nonparametric and do not assume linearity for a predictor-outcome association, they are usually more adept at generating predictions based on continuous variables."

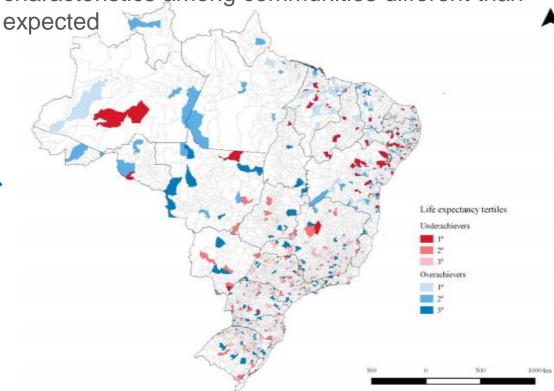
Machine Learning for Precision Public Health

"use better and more precise data to target disease prevention and control"-CDC

Step 1: Predict life expectancy based on local features and identify the over- and under-achievers



Step 2: Investigate modifiable public health characteristics among communities different than



Source: Chiavegatto Filho et al Epidemiology 2018; 29:836-840.

Machine Learning for Novel Data Generation

Image Analysis and Text Mining of Large Resources



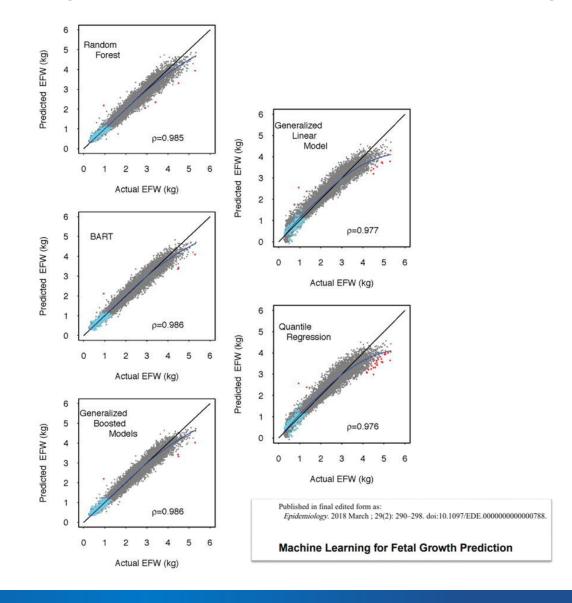
Environment International

Volume 122, January 2019, Pages 3-10



A picture tells a thousand...exposures: Opportunities and challenges of deep learning image analyses in exposure science and environmental epidemiology

Scott Weichenthal ^a ♀ ☑, Marianne Hatzopoulou ^b, Michael Brauer ^c



Overview

- What is "machine learning"? Does it matter?
- How have these methods been traditionally used?
 - Prediction vs Explanation
- Types of Machine Learning and Analytic Pipelines
- Common Algorithms
- Overview of Caret Package in R

How "machine learning" is defined often depends on who you ask.....

Computational methods using experience to improve performance or to make accurate predictions. Here, experience refers to the past information available to the learner, which typically takes the form of electronic data....In all cases, its quality and size are crucial to the success of the predictions made by the learner.

-Foundations of Machine Learning Mohri, Rostamizadeh, Talwalkar, The MIT Press

A program or system that builds (trains) a predictive model from input data. The system uses the learned model to make useful predictions from new (never-before-seen) data drawn from the same distribution as the one used to train the model. Machine learning also refers to the field of study concerned with these programs or systems.

-Google

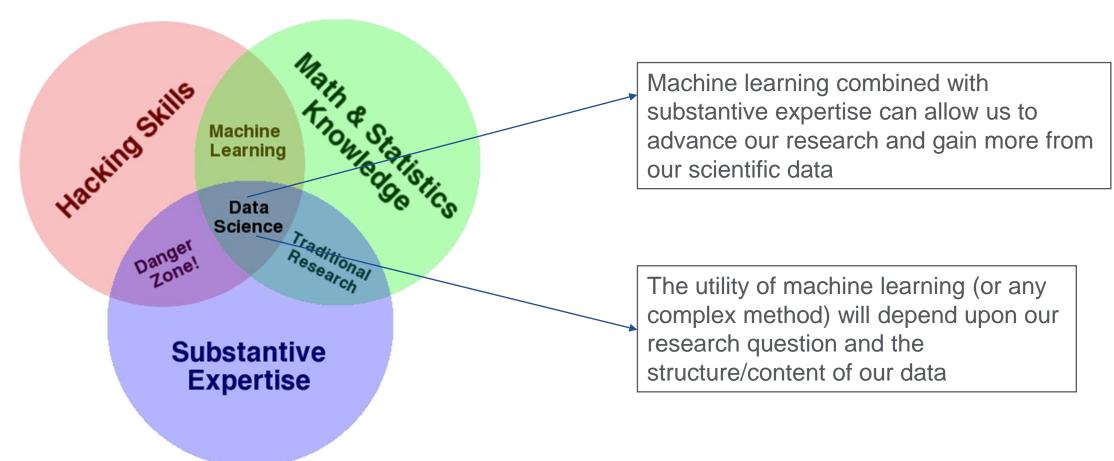
..an umbrella term for techniques that **fit models algorithmically by adapting to patterns in data**-Mooney and Pejaver, Annual Review of Public Health

Scientific study of algorithms and statistical models that computer systems use to effectively **perform a specific task without using explicit instructions, relying on patterns and inference instead**. - Wikipedia

Machine Learning: Intersection between Computational and Mathematical/Statistical Knowledge



Machine Learning: Intersection between Computational and Mathematical/Statistical Knowledge



To Explain or To Predict: What is the question...and what is the difference?

Explanatory Modeling: use of statistical models to test (or estimate) hypothesized causal associations; requires pre-existing causal model

Predictive Modeling: use of data to develop model that can predict new or future observations

Machine learning approaches traditionally used **AND** developed for prediction goals.

- Note there are questions of prediction within explanatory modelling
 - construction of propensity scores
 - use of risk scores to account for confounding
 - predicting the counterfactual
- ➤ If goal is not prediction, do we need to adapt machine learning approaches for our goal?

But what if my goal is explanation, but I don't have a good pre-existing causal model.....

➤ "By capturing underlying complex patterns and relationships, predictive modeling can suggest improvements to existing explanatory models" ---Shmueli 2010

Identifying "Predictors" using machine learning

Factors Related to Pediatric Unintentional Burns: The Comparison of Logistic Regression and Data Mining Algorithms

Abbas Aghaei, PhD, Hamid Soori, PhD, Azra Ramezankhani, PhD, Yadollah Mehrabi, PhD ▼

Journal of Burn Care & Research, irz066, https://doi.org/10.1093/jbcr/irz066

Published: 27 April 2019

Excerpt from the Abstract: The majority of the burn-related variables were related to individuals' social welfare status and their environments. Lessening the effects of these factors could reduce the incidence of pediatric burns.

...Reliant on the assumption that a good predictor is a good explainer.....

How can Epidemiologists Benefit from Training in Machine Learning?

- Facilitate use of large and/or complex data where relationships cannot be easily visualized
 - ➤ Use of ML approaches can identify patterns in data; potentially generate hypotheses, refine metrics of exposure and/or outcome
- ➤ Make exploratory data analysis and model selection more formal
 - ➤ Similar to use of DAGs to explicitly represent assumptions of relationships between variables
 - ➤ Don't just publish the final model, show how you arrived there.
- ➤ Greater consideration of questions of prediction and how they can benefit public health

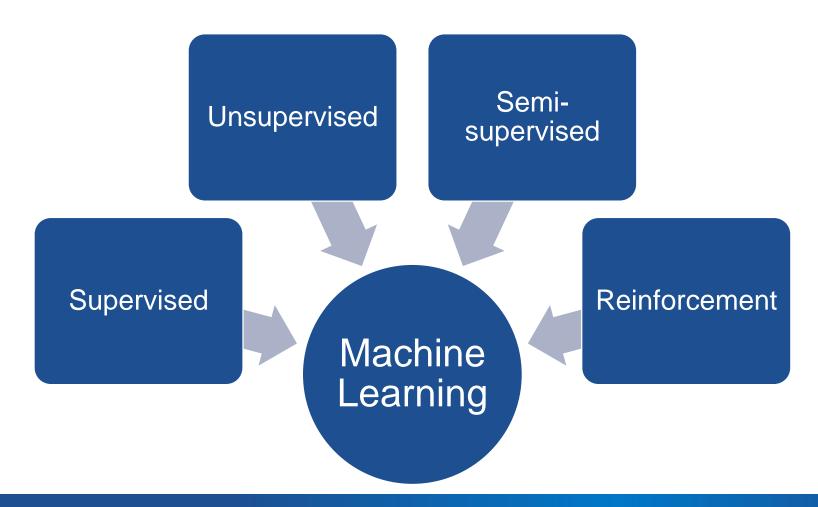
Prediction in Epidemiology & Public Health

Interested in predicting some exposure/outcome or predictions as intermediate product toward another aim

- Disease/Outcome Forecasting
 - > Predicting peak demand days for healthcare services associated with specific events
- Creation of Disease/Outcome Risk Scores
 - > Triage in the ED for who needs supplemental treatments
- Construction of Propensity Scores/Inverse Probability of Treatment Weights
- > Imputation of Missing Data; Construction of Synthetic Data
- Predicting toxicity of chemicals based on structure
- > Exposure Modelling
 - Image analysis to predict exposures
 - Can calculating traffic density allow you to estimate PM emissions in areas with little air monitoring?

What are the different types of machine learning?

Types of Machine Learning



Unsupervised

Context: for each observation of the inputs (predictor/exposure/independent variables), there is no associated output (response measurement); also described as data are "unlabeled"

Algorithm identifies patterns within the vector of inputs and generates an output that seeks to understand or represent the relationships between variables and/or observations.

Addresses: Clustering and Dimension Reduction Problems

Clustering to refine the outcome classification



Cluster analysis identifying clinical phenotypes of preterm birth and related maternal and neonatal outcomes from the Brazilian Multicentre Study on Preterm Birth

Clustering for Exposure Assessment

Vol. 127, No. 10 | Research

Air Pollution, Clustering of Particulate Matter Components, and Breast Cancer in the Sister Study: A U.S.-Wide Cohort

Alexandra J. White 🖂, Joshua P. Keller, Shanshan Zhao, Rachel Carroll, Joel D. Kaufman, and Dale P. Sandler

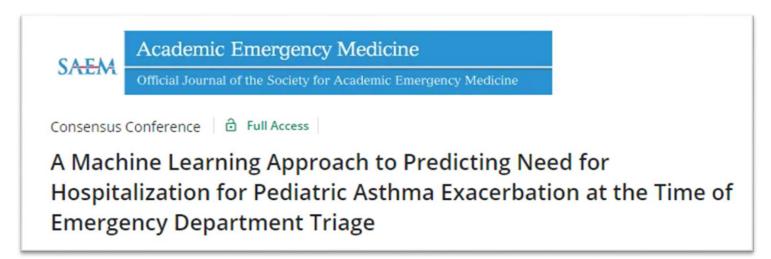
Published: 9 October 2019 | CID: 107002 | https://doi.org/10.1289/EHP5131 | Cited by: 12

Supervised

Context: for each observation of the inputs (predictor/exposure/independent variables), there is an associated output (response measurement); also described as data are "labeled"

Algorithm learns how to use inputs to generate outputs through training and receives feedback by looking at actual outcomes; process is "supervised"

Addresses: Regression, Classification and Estimation Problems



Semi-supervised

Context: for each observation of the inputs (predictor/exposure/independent variables), only a (typically) small subset has an associated output (response variable); combination of labeled and unlabeled data

Algorithm learns from subset of labeled data and then applies what it has learned to the unlabeled data.

Addresses: Useful when it is overly human or resource intensive to obtain sufficient amounts of labeled data to train a supervised algorithm



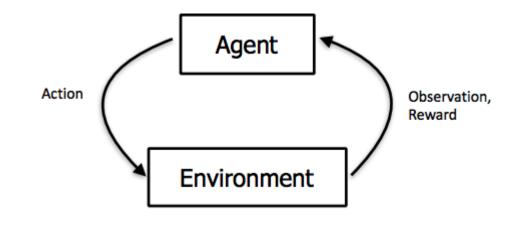
Reinforcement

Algorithm learns how to act in a given environment through maximization of reward; needs to anticipate future rewards from short-term actions

Original Paper

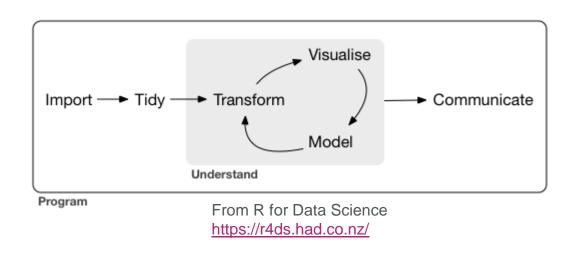
Encouraging Physical Activity in Patients With Diabetes:
Intervention Using a Reinforcement Learning System

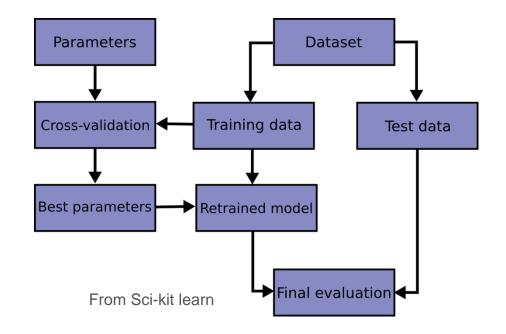
Elad Yom-Tov¹, PhD; Guy Feraru², MD, PhD; Mark Kozdoba³, PhD; Shie Mannor³, PhD; Moshe Tennenholtz⁴, PhD; Irit Hochberg⁵, MD, PhD



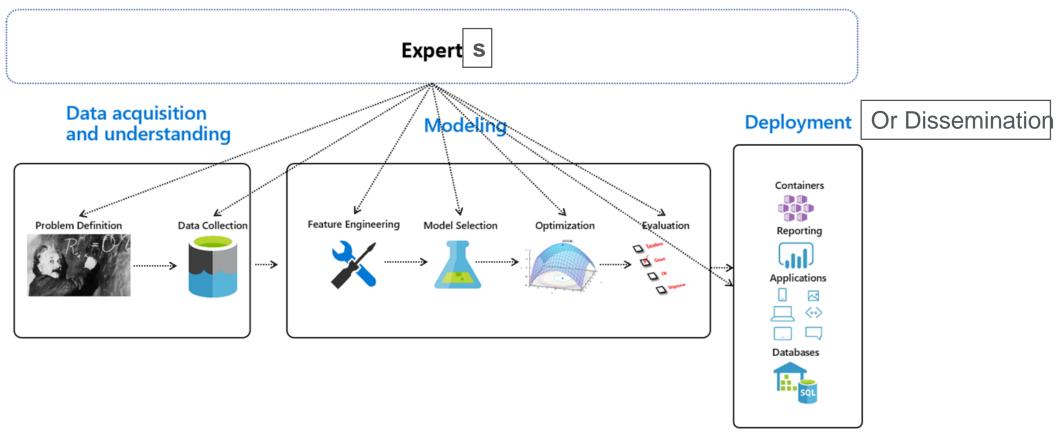
Pipelines

Ordered set of tasks to accomplish a specific task or goal; Visual study protocol Specific definition varies by field and their perspective on data analytics





Pipeline of entire Data LifeCycle



Commonly used Terms in Machine Learning pipelines

Small n, large-p vs Small p, large-n problem

- o n-number of individuals in dataset, p-number of features for each individual
- Refers to shape of dataset (wide vs long) with each having specific set of challenges

Parameters

- o a variable, internal to the model, and derived from the data; often saved as part of final model
- Example: β in a regression model

Hyperparameters

- o a variable, external to the model and often set by the programmer/analyst; used to estimate model parameters or to optimize the algorithm; can also be called tuning parameter
- Example: number of trees in a random forest

Tuning

 Customization of a model by varying the hyperparameters to determine the values that provide the optimal performance

Evaluation: Focus on Prediction

ClassificationConfusion MatrixROC C ROC Curves and Area Under the Curve Calibration Mean Square ErrorR-squared Regression



How we Evaluate (again focus on prediction): Differentiating Training, Validation and Testing

Data Partitioning

- Splitting a dataset into random subsets for use in either training, validating or testing the machine learning model
- Use of different subsets
 - Training: used by algorithm to learn the resulting model
 - Validation: used to compare performance of models produced by different algorithms, hyperparameters, ...
 - Test/Hold Out: used to obtain final metrics of performance and results of the model

Sample size typically dictates how data are partitioned.

More data used for training than testing in the context of prediction.

Also includes creation of K-folds for cross-validation. Folds are equal sized.

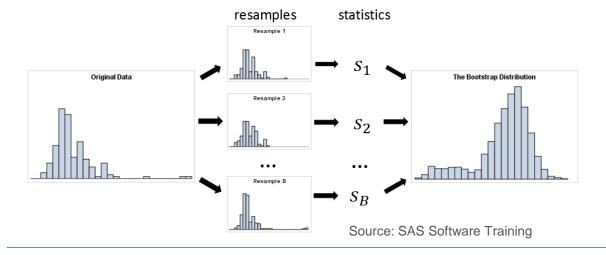
Resampling Methods

Bootstrapping

- Iteratively sampling with replacement
- Used to estimate parameters and draw inferences on a population
- Used in ensemble methods e.g. bagging

Cross-validation

- Validation technique
- Partition data into k non-overlapping subsets
- Estimate model parameters on k-1 subsets (training) then apply model in the held-out subset for evaluation metrics
- Repeat k times
- Similar Approach for Leave-one-out Cross-Validation



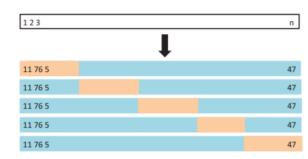


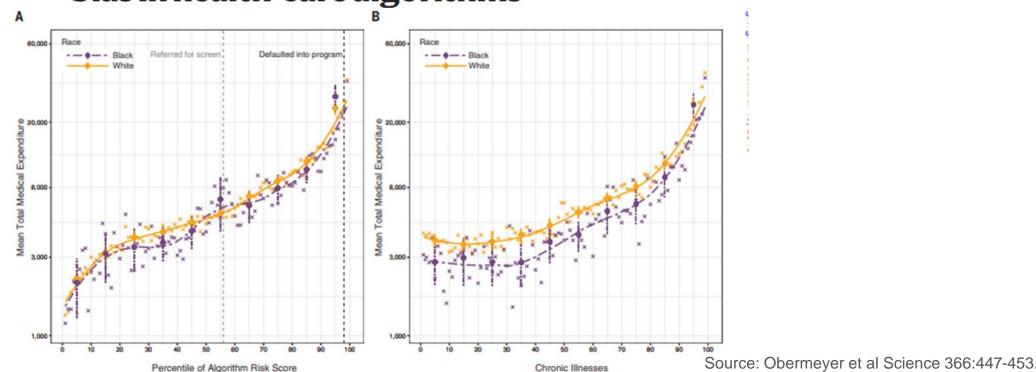
FIGURE 5.5. A schematic display of 5-fold CV. A set of n observations is randomly split into five non-overlapping groups. Each of these fifths acts as a validation set (shown in beige), and the remainder as a training set (shown in blue). The test error is estimated by averaging the five resulting MSE estimates.

Source: Introduction to Statistical Learning in R

Evaluating in the Real World: Do models behave as expected?

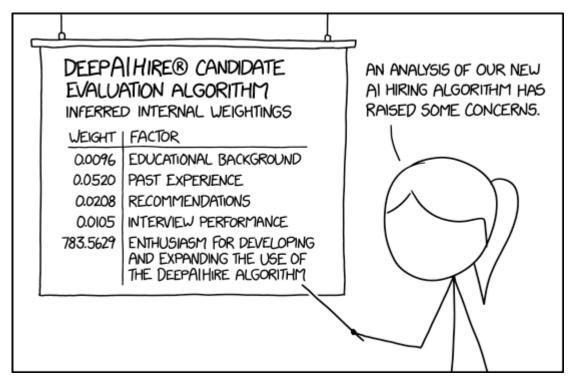
IEWS · 24 OCTOBER 2019 · UPDATE 26 OCTOBER 2019

Millions of black people affected by racial bias in health-care algorithms



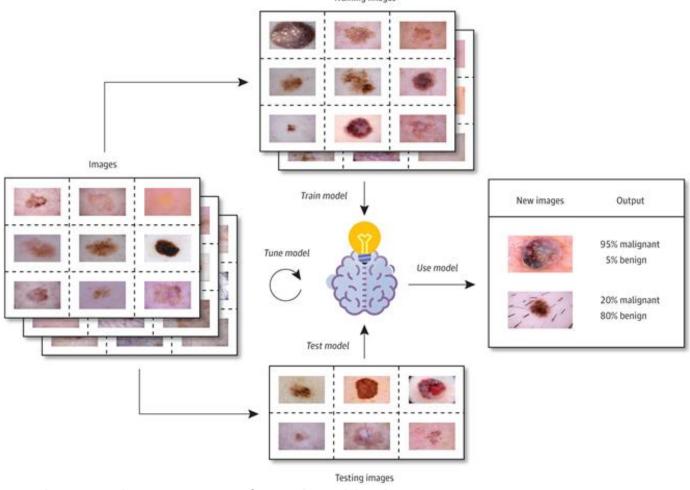
Know the Data that Drives Your Predictions

Are there biases in our past actions?



Source: XKCD https://xkcd.com/2237/

Bias doesn't have to be explicitly programmed



From: Machine Learning and Health Care Disparities in Dermatology JAMA Dermatol. 2018;154(11):1247-1248. doi:10.1001/jamadermatol.2018.2348

Commonly-used Algorithms

Unsupervised

K-Means*

Hierarchical clustering*

PCA

Self Organizing Maps

Gaussian Mixture

Models

Supervised

Support Vector Machines

Naïve Bayes

K-Nearest Neighbors*

Regularized Regression*

Decision Trees*

Neural Networks

Ensemble*

Bagging

Random Forest

Boosting

XGBoost

Stacking

SuperLearner

*To be reviewed in more detail in later slides

Clustering

Broad set of techniques for finding subgroups or clusters in a dataset-ISLR

Difference between clusters>>> difference within clusters

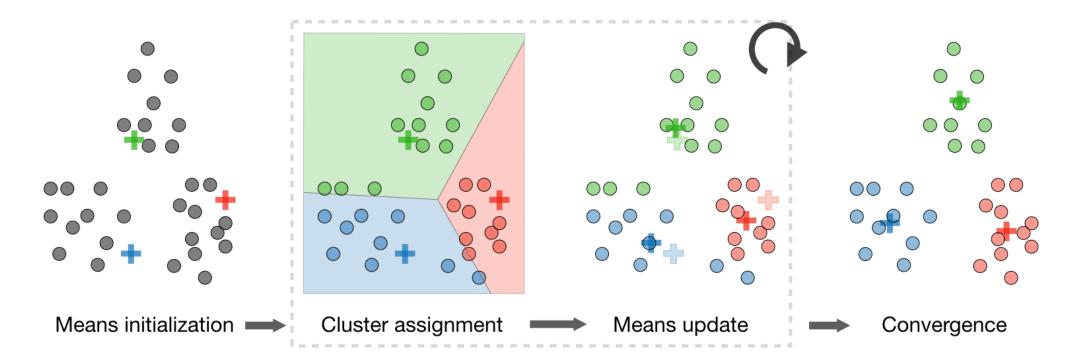
All observations are forced into a cluster

Two common algorithms: K-means and Hierarchical, but also k-prototypes, PAM, etc.

Applications

Identify clusters of observations based on features: phenotypic subgroup identification Identify clusters of features based on observations: identifying genetic expression patterns

K-means clustering: user must specify k

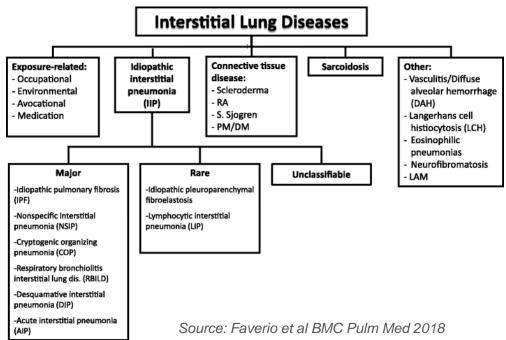


Source: CS-229 Course Material

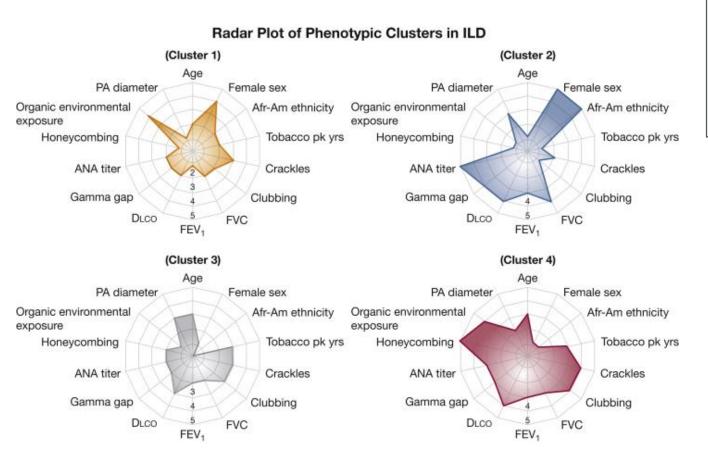
Application: Identify Phenotypic Subtypes



Goal: "Identify distinct clinical phenotypes in heterogeneous diseases"



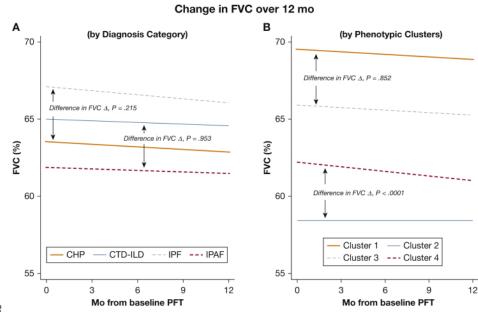
Method: Partitioning around Medoids (PAM)



Why PAM?

Similar to K-means, but more robust to outliers

Relies on median distances rather than means



Hierarchical Clustering

Builds nested clusters in a successive manner

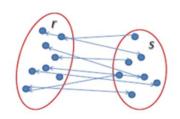
Agglomerative: each observation starts in its own cluster, and pairs of clusters are merged successively; better in discovering small clusters

Divisive: all observations start in the same cluster and splits are performed recursively; better in discovering large clusters

Merges and splits decided by cluster **dissimilarity**. Dissimilarity computed by **distance** and **linkage**.

Multiple distance metrics for calculating dissimilarity

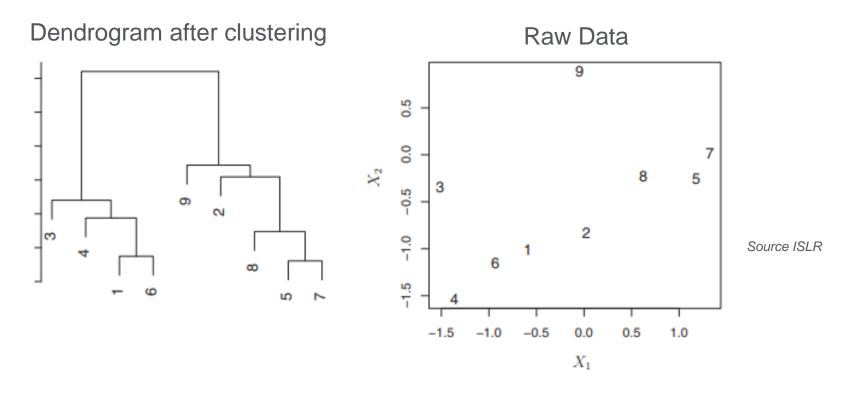
- Euclidian
- Squared Euclidian
- Manhattan
- Maximum
- Mahalanobis



Multiple criterion for linkage

- Complete
- Single
- Average

Dendrogram: visualization of hierarchical clustering



Again, need to determine optimal number of clusters

Evaluating Number of Clusters

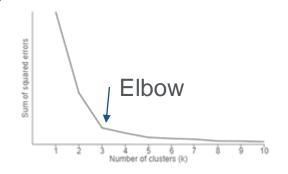
Goal: minimize the intra-cluster variation (i.e. homogeneous clusters)

Elbow: Plot the within-cluster sum of squares. Optimal clusters is identified at the bend in the plot

Silhouette: measure of the quality of the clustering, maximize s

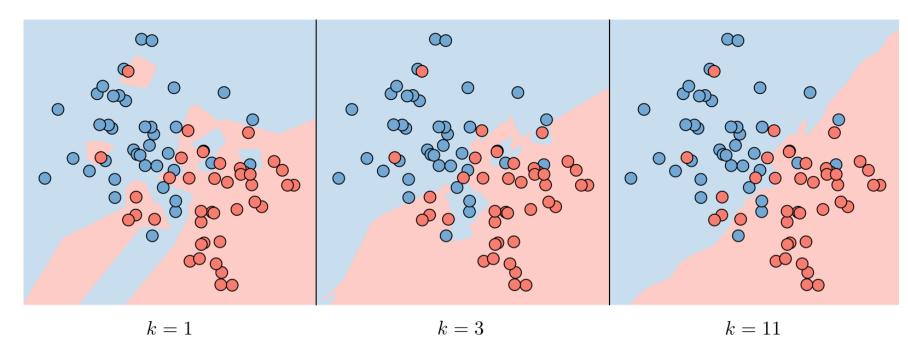
$$s=rac{b-a}{\max(a,b)}$$
, where a is mean distance between a sample and other samples within the same class and b is the mean distance between a sample and other points in the next nearest cluster

Gap-statistic: compares total within cluster variation with their expected values under the null reference distribution. Requires boostrapping to generate reference distribution. Maximize the gap-statistic to identify optimal number of clusters.



K-nearest neighbors

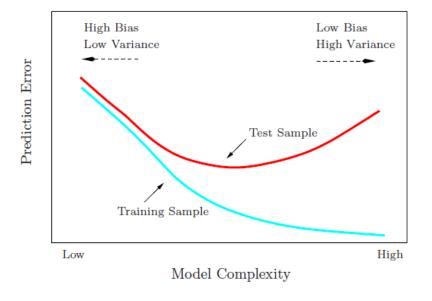
Non-parametric approach where the predicted value of an observation is determined by the nature of its k-neighbors from the training set.



Source: CS229 Course Material

Regularized Regression (shrinkage)

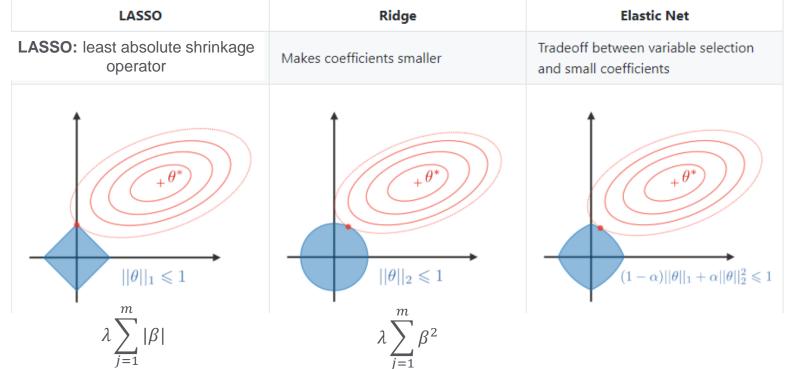
Addresses limitation of regression: Increase flexibility/complexity by including power terms, interactions, etc. decreases bias but can increase variance (overfitting)



$$\sum_{i=1}^{N} \left(y_i - \theta_0 - \sum_{j=1}^{p} \theta_j \cdot x_{ij} \right)^2 + \lambda \|\theta\|$$
 Penalty function

Residual sum of squares
Typical least squares loss function

Difference in the algorithms: penalty



L1 Regularization

L2 Regularization

Combines L1 and L2 Regularization

Elastic Net: $\frac{\sum_{i=1}^{n}(y_i-x_i^J\hat{\beta})^2}{2n} + \lambda \left(\frac{1-\alpha}{2}\sum_{j=1}^{m}\hat{\beta_j^2} + \alpha\sum_{j=1}^{m}|\hat{\beta_j}|\right)$

Alpha is the hyperparameter that controls the loss function that differentiates LASSO, Ridge and Elastic Net

Varying Lambda (λ)

Controls the penalty function

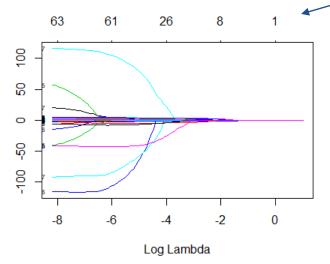
 $\lambda = 0$: penalty=0 and produces standard regression coefficients

λ -> ∞: penalty=large and regression coefficients shrunk toward 0

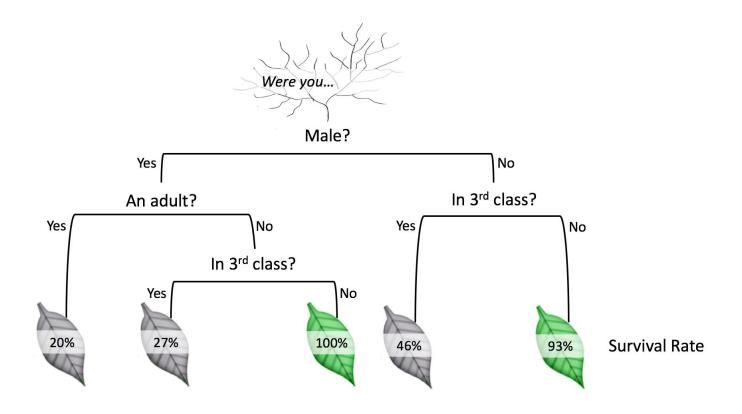
λ : chosen through **cross-validation**

Number of features

Plot of coefficients from LASSO when varying λ

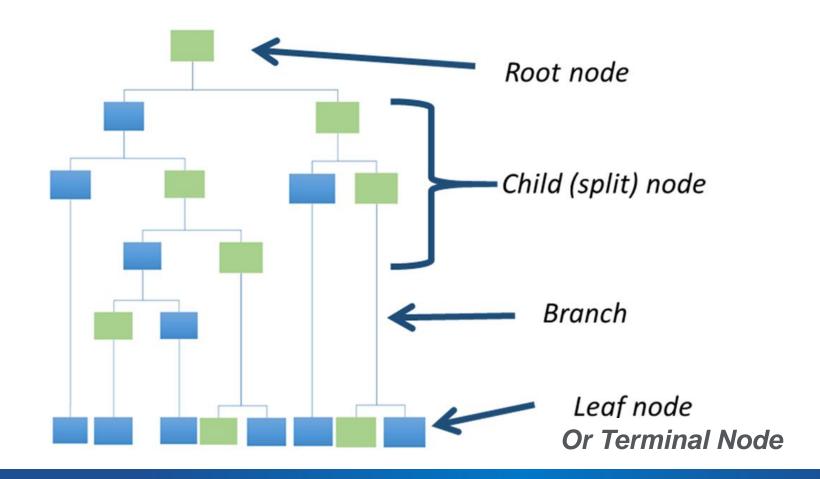


Intro to Tree-Based Methods: Titanic Example



Structure of a Tree

Trees generated through recursive partitioning



Key Terms

"Greedy" Algorithm: makes the optimal choice at each step

Makes "best" first split without consideration of subsequent splits

Node purity: homogeneity of a node in relation to the labels of the observations contained

Goal is often to maximize node purity to obtain optimal classification or prediction

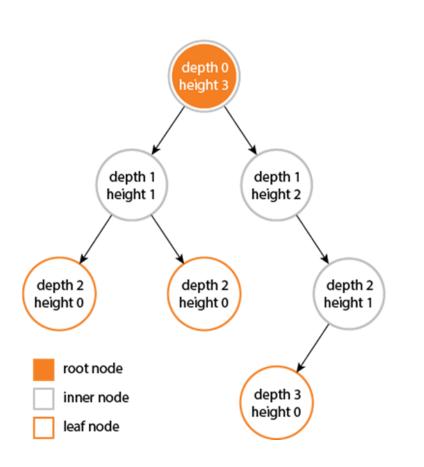
Measures of purity

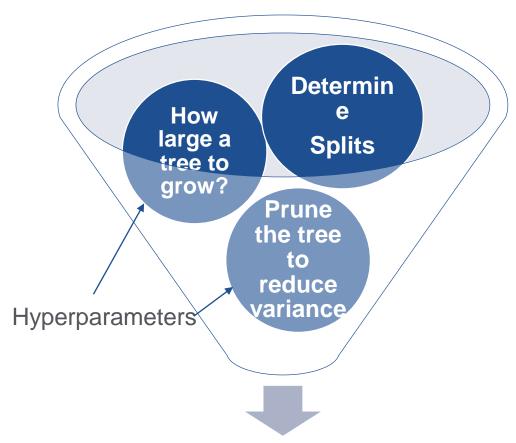
Gini coefficient, entropy, variance, mean square error.... and others

Surrogate split: split using another feature that most closely resembles the consequences of the original split

Often how tree-based methods handle missing data

Growing a Tree: Analytic Considerations



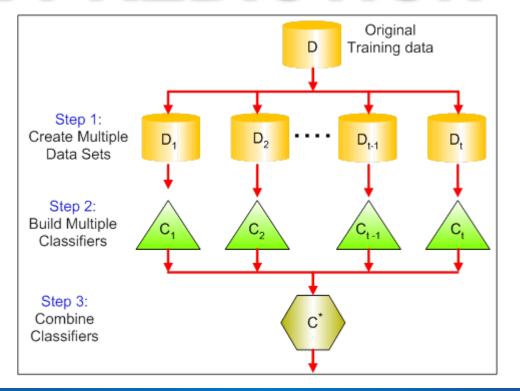


Constructing a tree-based model

Ensemble Methods

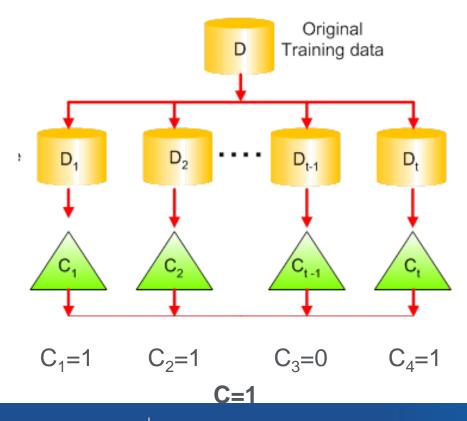
IMPROVE PREDICTION

Ensembles **combine** several base models in order to produce an optimal prediction

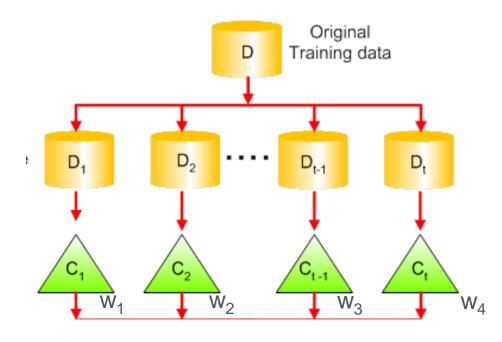


Simple Techniques to Combine: Classification

Majority Voting



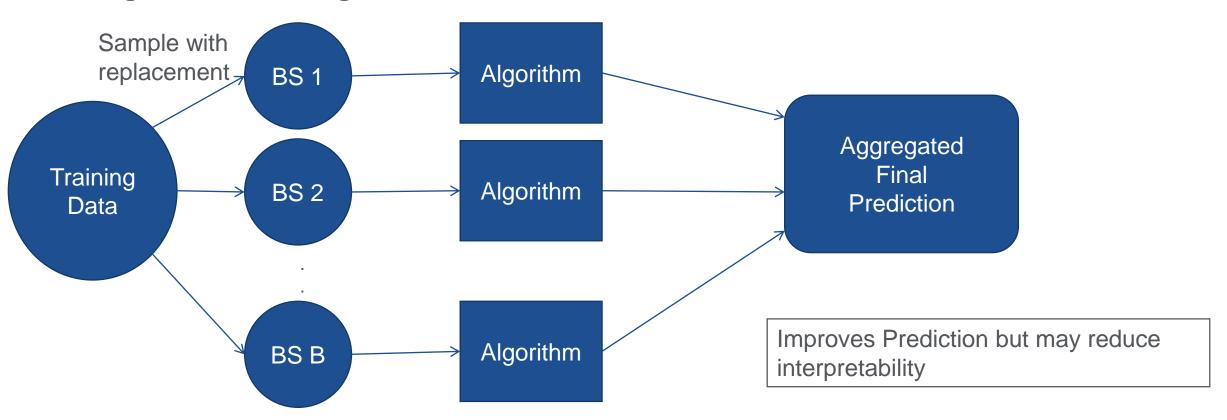
Weighted Voting



Final Prediction (C) a weighted combination with weights calculated based on predictive ability

Advanced Techniques: Bagging

Bagging: Bootstrap Aggregation; average results (or voting) across bootstrapped samples of data



Random Forest: Extension of Bagging

At each split, we choose a random sample of *m* predictors from the full set of *p* features

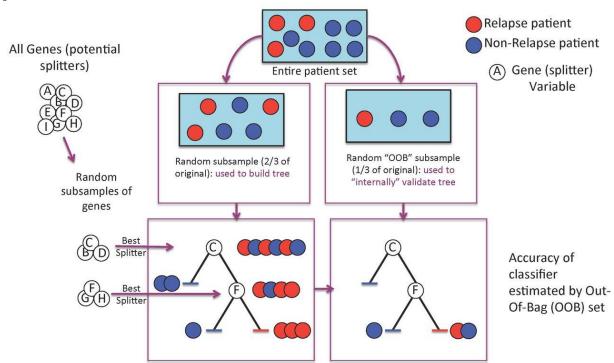
When m=p, then bagging and RF are equivalent

Randomly sampling from features decorrelates the trees

Provides opportunity for all features to contribute to prediction

Improves prediction overall, even though individual trees may be weaker

For prediction, want m to be small, typically m=√p



Source: BioInformatics Handbook

Variable Importance Factors

Measure of Individual Variable Contribution to the Overall Prediction

Accuracy-based Importance

Within OOB, record the prediction/classification error.

Permute the feature and recalculate prediction/classification

Difference in two errors are averaged over trees and normalized.

Node purity based Importance (Gini importance for classification)

Within OOB, total decrease in node impurities from splitting on variable averaged over all trees.

Advanced Techniques: Boosting

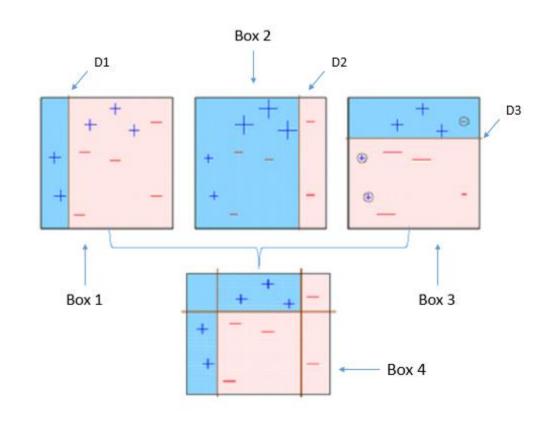
General technique that can be applied to diverse algorithms; popular with trees as base

Goal: Convert series of weak learners to a strong learner by learning algorithms sequentially

Example: Results from first algorithm provide information to second, etc.

Both provide an initial strong learner and then add to the initial learner with weaker learners

In adaptive boosting, information is weights of data points. If classified correctly, gets smaller weight so algorithm focuses on misclassified datapoints.



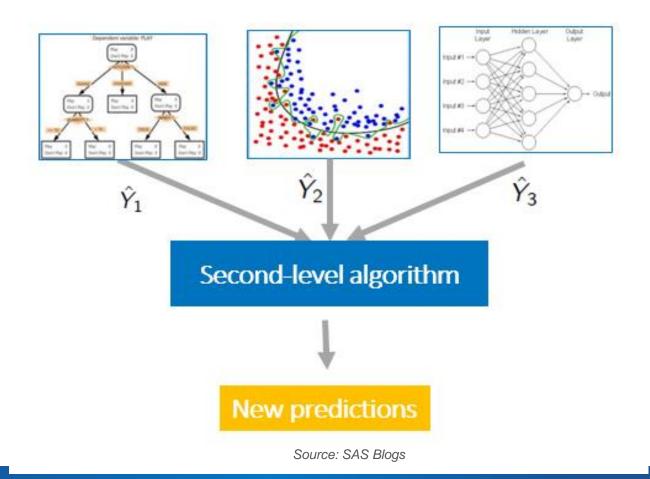
Source: Desarda Understanding AdaRoost

Advanced Techniques: Stacking

Rather than combining results of multiple learners, stacking uses predictions from learners as inputs to additional learners.

Optimal performance when diverse learners are used in the initial predictions

Common Example in Epidemiology: SuperLearner



Overview of Super Learner

Data Partitionin g

- Partition into K-folds
- Within each fold, partition into training and testing

Fit with library of algorithms

- Select library consisting of 'm' algorithms
- For each fold, fit each algorithm on the training set. Apply in test set. Calculate evaluation metric (L-2 squared error loss but equivalent to MSE)

Combine across folds

- Average evaluation metrics across all folds to obtain one measure for each algorithm
- Discrete SuperLearner: choose algorithm which minimizes loss

Combine across algorithms

- Regress actual outcome against predictions from each algorithm with certain constraints (depending upon type of data)
- Obtain weights for each algorithm by normalizing coefficient values

Osain final predicted outcome

• Use weights in combination with predictions from each algorithm to generate predictions for newly observed data

Critical thinking is not optional....



Credit: XKCD

Consider needs of research question....



FIGURE 2.7. A representation of the tradeoff between flexibility and interpretability, using different statistical learning methods. In general, as the flexibility of a method increases, its interpretability decreases.

What types of epidemiologic questions/tasks benefit from high flexibility?

What types require more interpretability?

What does interpretability mean in the context of machine learning?



Source: ISLR

Consider particulars of the data...

Are data highly correlated?

Do you anticipate non-linear effects?

Are you interested in interactions between features/exposures?

Multiple avenues to integrate machine learning into epidemiologic research and practice..

Complex algorithms won't solve all our problems, but they can enhance our research if applied *thoughtfully*

Largely underutilized potential of using machine learning for other tasks like data harmonization, data-linkage and knowledge modelling



Funding and Institutional Support:

NIEHS Career Development Award (R00ES027022); HHEAR Data Science Center-Subcontract (U2CES026555) Columbia Calderone Early Faculty Prize Columbia DSI Scholars Program