#### **Korea Advanced Institute of Science and Technology**

School of Electrical Engineering

# Machine Learning Technique for Survival Analysis

July 22, 2021

**GWANGSU KIM** 



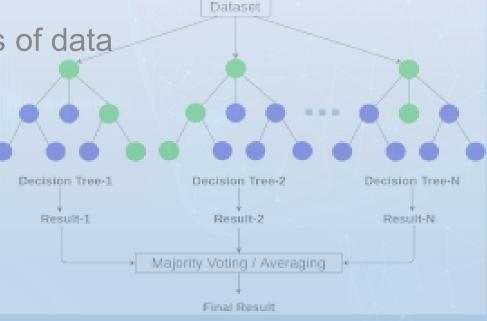
### Contents



- Data, Model and Loss function
- Tree and Random Forests
- Survival Tree (Random Forests)

XG Boosting

 Not providing examples of data analysis.





# Data, Model and Loss function



Data:  $\{X_i, T_i, \delta_i\}_{i=1}^N, T_i = \min(Y_i, C_i)$ 

Model: 
$$\lambda(t \mid X) = \lim_{\delta \downarrow 0} P(t \le Y < t + \delta \mid t \le Y, X)/dt$$

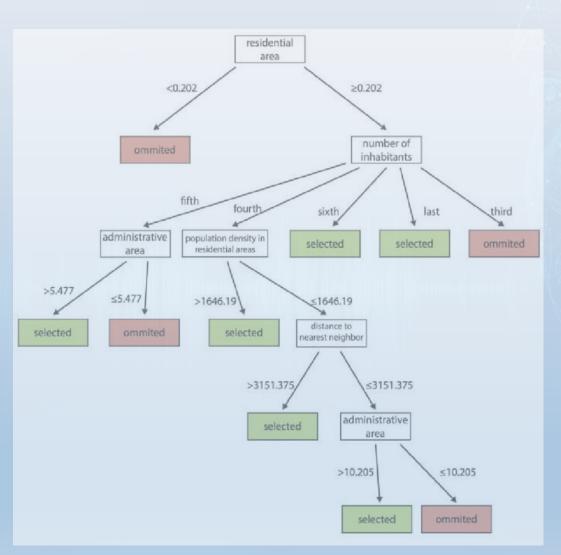
Loss function: Negative log likelihood

$$-\sum_{i=1}^{N} \left\{ \delta_i \log \lambda(T_i \mid X_i) - \int_0^{T_i} \lambda(s \mid X_i) ds \right\}$$



### Tree





Node 1	Node 2	
	Node 3	Node 4

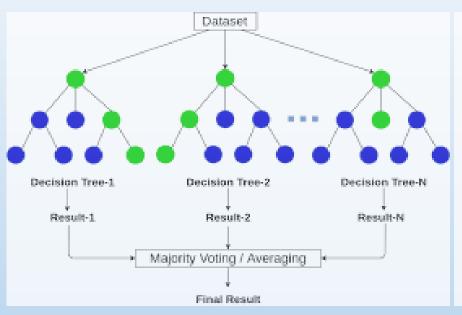
- 1. Sequentially selecting variable and splitting to minimize the criteria.
- 2. Applicable in highdimensional features.
- Most critical point is the rule of splitting.
- 4. Pruning (deleting nodes ) to avoid the over-fitting.
- 5. Non-parametric prediction

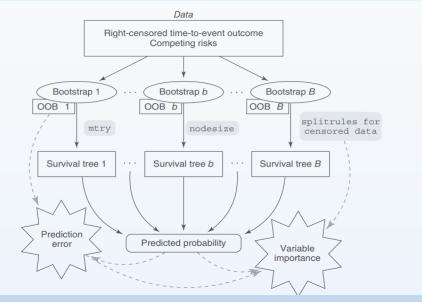


google

### Random Forests







google

Helen et al, 2018

- Making many trees (bootstrap sampling and subset selection of feature dimensions)
- As independent as possible
- Final prediction is a voting.

- Survival Tree: criteria rule
- Constraints: mtry-trying subset size of feature, nodesize, splitrules for censored data
- OOB: out-of-bag



### **Survival Trees**



Splitting rule

When  $\mathcal{X}_l$  and  $\mathcal{X}_r$  are supports for the left and right nodes respectively. Log-rank test (equivalence of CHFs) statistic:

$$\frac{1}{\hat{\sigma}^{LR}} \int_0^t \frac{\bar{Y}_l(s)\bar{Y}_r(s)}{\bar{Y}_l(s) + \bar{Y}_r(s)} \left( \frac{d\bar{N}_l(s)}{\bar{Y}_l(s)} - \frac{d\bar{N}_r(s)}{\bar{Y}_r(s)} \right)$$
 where  $\bar{N}_r(s) = \sum_{i=1}^N \mathbb{I}(X_i \in \mathcal{X}_r) \mathbb{I}(T_i \leq t, \delta_i = 1)$  
$$\bar{Y}_r(t) = \sum_{i=1}^N \mathbb{I}(X_i \in \mathcal{X}_r) \mathbb{I}(T_i > t)$$

Constraints

mtry: size of random subset in splitting

nodesize: number of nodes split rules for censored data



# Radom Forests for Survival Analysis



 How to estimate/predict the CHF using random forests (Ishwaran et al., 2008)

1. Combining N-A estimators  $H_n(t)$  of all nodes.

OOB ensemble 
$$H_e(t \mid X) = \frac{\sum_{n=1}^{ntree} H_n(t \mid X)(X \notin \mathcal{L}_n)}{\sum_{n=1}^{ntree} (X \notin \mathcal{L}_n)}$$

- PredictionOOB ensemble for new X
- 3. Ensemble KM estimator (Hothorn, 2004)

$$\hat{S}(t \mid X) = \prod_{s \le t} \left( 1 - \frac{\sum_{i=1}^{N} \sum_{n=1}^{ntrees} n_{i,n} L_{i,n}(X) N_i(ds)}{\sum_{i=1}^{N} \sum_{n=1}^{ntrees} n_{i,n} L_{i,n}(X) Y_i(s)} \right)$$



### Prediction error and Importance of variables



#### Prediction error

#### Modification of the C-index,

the C-index compare the pairs (small uncensored, large all)s. If the estimated hazard of large observation is larger than the other, then it is considered as error.

#### OOB prediction error

When 
$$\sum_{j=1}^{J} H_e(t_j^* \mid X_i) \ge \sum_{j=1}^{J} H_e(t_j^* \mid X_j)$$
 and  $T_i \ge T_j$ 

Importance of variables (VIMP)

For trees concerning a specific variable, convert the split by the variable to random splitting. Measure the ration of PE increasing



### Issues



- Censoring ratio in each node
- Can variable selection be possible? In conventional random forests, some papers exist.
- Can high-dimensional analysis be available with certain levels?
- More explainable approach / extension such as cure model, some papers exist.





- XG Boosting (Chen and Guestrin, 2016) is the best in many competitions for the prediction of structure datasets.
- Basic concept is to use the Talyor series expansions and gradient boosting.

$$\mathcal{L}^{t} = \sum_{i=1}^{N} \ell(y_{i}, \hat{y}_{i}^{t-1} + f_{t}(X_{i})) + \Omega(f_{t})$$





$$\mathcal{L}^{t} = \sum_{i=1}^{N} \ell(y_{i}, \hat{y}_{i}^{t-1} + f_{t}(X_{i})) + \Omega(f_{t})$$

$$\approx \sum_{i=1}^{N} \left\{ \ell(y_{i}, \hat{y}_{i}^{t-1}) + g_{i}f_{t}(X_{i}) + \frac{1}{2}h_{i}f_{t}(x_{i})^{2} \right\} + \Omega(f_{t})$$

$$\approx \sum_{i=1}^{N} \left\{ \ell(y_{i}, \hat{y}_{i}^{t-1}) + g_{i}f_{t}(X_{i}) + \frac{1}{2}h_{i}f_{t}(x_{i})^{2} \right\} + \gamma T + \frac{1}{2}\lambda \sum_{j=1}^{T} w_{j}^{2}$$

$$= \sum_{j=1}^{T} \left\{ w_{j} \sum_{i \in I_{j}} g_{i} + \frac{1}{2}w_{j}^{2} \left( \sum_{i \in I_{j}} h_{i} + \lambda \right) \right\} + \gamma T$$

$$w_{j}^{*} = -\frac{\sum_{i \in I_{j}} g_{i}}{\sum_{i \in I_{j}} h_{i} + \lambda}, \quad \mathcal{L}_{split} = \frac{1}{2} \left[ \frac{(\sum_{i \in I_{L}} g_{i})^{2}}{\sum_{i \in I_{L}} h_{i} + \lambda} + \frac{(\sum_{i \in I_{R}} g_{i})^{2}}{\sum_{i \in I_{R}} h_{i} + \lambda} - \frac{(\sum_{i \in I} g_{i})^{2}}{\sum_{i \in I} h_{i} + \lambda} \right]$$

$$\mathcal{L}(w_{j}^{*})^{t} = -\frac{1}{2} \sum_{j=1}^{T} \frac{(\sum_{i \in I_{j}} g_{i})^{2}}{\sum_{i \in I_{j}} h_{i} + \lambda} + \gamma T$$





 Based on the previous additive model, simple tree (only one split) is added until at most several hundreds or thousands trees.

#### Other Tricks:

- 1) Greedy search for splitting point: using percentile of features.
- 2) Weighted quantile (varying Hessian based)
- 3) Sparsity-aware split: ignoring the missing points.
- 4) Programming level techniques.





- Gradient and Hessian are required.
- Partial likelihood and other likelihood can be candidates.
- Sparsity can be implemented with the boosting procedure.
- Feature extraction can be used for the conventional inference.



### References



Chen, Tianqi, and Carlos Guestrin. "Xgboost: A scalable tree boosting system." *Proceedings of the 22nd acm sigkdd international conference on knowledge discovery and data mining*. 2016.

Genuer, Robin, Jean-Michel Poggi, and Christine Tuleau-Malot. "Variable selection using random forests." *Pattern recognition letters* 31.14 (2010): 2225-2236.

Ishwaran, Hemant, et al. "Random survival forests." *The annals of applied statistics* 2.3 (2008): 841-860.

Ma, Li, and Suohai Fan. "CURE-SMOTE algorithm and hybrid algorithm for feature selection and parameter optimization based on random forests." *BMC* bioinformatics 18.1 (2017): 1-18.

Rytgaard, Helene C., and Thomas A. Gerds. "Random forests for survival analysis." *Wiley StatsRef: Statistics Reference Online* (2014): 1-8.

