# Machine Learning for Survival Analysis: A New Approach

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ABSTRACT: We have applied a little-known data transformation on subsets of the Surveillance, Epidemiology, and End Results (SEER) publically available data of the National Cancer Institute (NCI) to make it suitable input to standard machine learning classifiers. This transformation properly treats the right-censored data in the SEER data and the resulting Random Forest and Multi-Layer Perceptron models predict full survival curves. Treating the 6, 12, and 60 months points of the resulting survival curves as 3 binary classifiers, the 18 resulting classifiers have AUC values ranging from .765 to .885. Further evidence that the models have generalized well from the training data is provided by the extremely high levels of agreement between the random forest and neural network models predictions on the 6, 12, and 60 month binary classifiers.

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# 1 Introduction and Background

Extracting actionable information from data is changing the fabric of modern business. A class of techniques that transforms data into actionable information goes by the name of Machine Learning [13]. Machine Learning has recently become a popular method to answer questions and solve problems that are too complex to solve via traditional methods. The Surveillance, Epidemiolgy, and End Results (SEER) Program of the National Cancer Institute (NCI) has been collecting data because intuitively researchers feel confident that this

data is capturing information that has buried within it useful information in the form of relationships between the types of data collected (demographic as well as staging information) and the survival outcomes. Though this relationship evades capture by traditional methods, it is possible to surface it with the two machine learning techniques known as **Random Forests** and **Neural Networks**. These two methods produce very similar results when applied to the SEER dataset, and are based on two almost diametrically opposed learning philosophies, which lends confidence in the validity of the results.

The Surveillance, Epidemiolgy, and End Results (SEER) Program of the National Cancer Institute (NCI) is the most recognized authoritative source of information on cancer incidence and survival in the United States. SEER currently collects and publishes cancer incidence and survival data from population-based cancer registries covering approximately 28 percent of the US population.

Quoting directly from the SEER website [11]:

The SEER program registries routinely collect data on patient demographics, primary tumor site, tumor morphology and stage at diagnosis, first course of treatment, and follow-up for vital status. This program is the only comprehensive source of population-based information in the United States that includes stage of cancer at the time of diagnosis and patient survival data. The mortality data reported by SEER are provided by the National Center for Health Statistics. The population data used in calculating cancer rates is obtained periodically from the Census Bureau. Updated annually and provided as a public service in print and electronic formats, SEER data are used by thousands of researchers, clinicians, public health officials, legislators, policymakers, community groups, and the public.

One characterstic of the SEER data and that is shared by many datasets in the medical field goes by the name of "censored data." The SEER data contains the number of months each patient survived, as well as an indicator variable showing whether or not the patient is still alive at the end of the data collection period. Methods to deal effectively with this kind of "right-censored data" include Kaplan-Meier curves and Cox's Proportional Hazard models [4]. The Kaplan-Meier techniques only give estimates for cohorts of patients and are not applicable for predicting the surival curve for a single patient, and the Cox Proportional Hazard models require a fairly restrictive set of assumptions to be satisfied in order to yield reliable results. In addition, the Cox Proportional Hazard models are not able to capture the nonlinear relationships between the given data fields that go into making predictions; they can only capture the first-order linear relationships.

Previous work applying machine learning methods to subsets of the SEER data include creative attempts to deal with the problems presented by "right-censored data." The authors of [14] use semi-supervised learning techniques to predict 5 year survival, essentially imputing values for SEER records where the survival months infomation is censored at a value less than 5 years. The authors of [15] investigate the effects of comordbidities; i.e., patients with two different cancer diagnosises, but their treatment of the censored data

underestimates the survival probabilities. All records representing patients who survived at least 60 months as well as all those who died earlier than 60 months were considered, but patients alive prior to 60 months but censored out of the study before 60 months were not included. This treatment biases the data and the predictions, leading to overly pessimistic survival probabilities predicted by the trained models.

To overcome these limitations of the traditional methods, IOBS has applied a little-known technique to transform the SEER data to make it amenable to more powerful machine learning methods. The essential idea is to recast the problem to an appropriate discrete classification problem instead of a regression problem (predicting survival months). Treating months after diagnosis as just another discrete feature, the SEER data (or any other right-censored data) can be transformed simply so as to make predictions for the hazard function, probability of dying in the next month, given that the patient has not yet died. The full survival function can then be derived from the hazard function. Details of this transformation can be found in this blog post [1].

## 2 Methodology

### 2.1 Data acquisition

We used the publically available 1973-2012 SEER incidence data files corresponding to colon, breast and lung cancer contained in the following list. SEER requires that researchers submit a request for the data, which includes an agreement form. Detailed documentation explaining the contents of both the incidence data files used in this study as well as a data dictionary for the 1973-2012 SEER incidence data files are available without the need to register or submit a data request [12].

- incidence\yr1973 2012.seer9\COLRECT.txt
- incidence\yr1973 2012.seer9\BREAST.txt
- incidence\yr1973  $2012.seer9\RESPIR.txt$
- incidence\yr1992 2012.sj la rg ak\COLRECT.txt
- incidence\yr1992 2012.sj la rg ak\BREAST.txt
- incidence\yr1992 2012.sj la rg ak\RESPIR.txt
- incidence\yr2000 2012.ca ky lo nj ga\COLRECT.txt
- incidence\yr2000\_2012.ca\_ky\_lo\_nj\_ga\BREAST.txt
- incidence\yr2000 2012.ca ky lo nj ga\RESPIR.txt
- incidence\vr2005.lo 2nd half\COLRECT.txt
- incidence $\yr2005.lo_2nd_half\BREAST.txt$
- $incidence \yr2005.lo_2nd_half \RESPIR.txt$

# 2.2 Data preparation and preprocessing

A great deal of data munging is necessary before using these SEER incidence files as input into machine learning algorithms. A preprocessing step common to each of three cancer types studied involves the STATE-COUNTY RECODE. The STATE-COUNTY RECODE field is a state-county combination where the first two characters represent the state FIPS code

Code	Description
1	Male
2	Female

**Table 1**. Encoding of gender in the SEER incidence files. These types of categorical variables need to be transformed via one-hot-encoding.

and the last three digits represent the FIPS county code. This particular field illustrates an important feature of machine learning, that between *categorical features* and *numeric features*. All input into a machine learning algorithm must be numeric, but real numbers carry with them the usually extremely useful property known as the well-ordering property of the real numbers. But if one is tasked with encoding a categorical feature into suitable numeric format for machine learning, it is necessary to do so in a way that removes the well-ordering property [2].

As a simple example of how to correctly treat categorical variables in a mchine learning context, consider the SEER variable SEX. This variable is encoded with a numeric 1 for males and a numeric 2 for females as shown in Table 1. Values such as "Male" and "Female" encoded as numbers are dangerous because if not handled properly, they can generate bogus results [7]. The proper way to transform the SEER SEX variable is to create two additional variables: sex\_Male and sex\_Female, and then to eliminate the variable SEX. For example,

$$\left[\frac{\text{SEX}}{1}\right] \to \left[\frac{\text{sex\_Male} | \text{sex\_Female}}{1}\right]$$
(2.1)

and

$$\left[\frac{\text{SEX}}{2}\right] \to \left[\frac{\text{sex\_Male} | \text{sex\_Female}}{0 | 1}\right]$$
(2.2)

The procedure outlined in Equations (2.1, 2.2) needs to be applied to all of the nominal categorical variables in the SEER data that we wish to include in our predictive models. In particular, in order to include the geophgraphical information contained in the SEER categorical variable STATE-COUNTY RECODE, it becomes necessary to create a new feature variable for each of the distinct (state,county) pairs in the data. In the United States, there are approximately 3,000 counties. Clearly, transforming the STATE-COUNTY RECODE data representation into distinct (state\_county) columns will explode the data to become wider than is optimal for machine learning. Adding extra columns to your dataset, making it wider, requires more data rows (making it taller) in order for machine learning algorithms to effectively learn [2]. Because one-hot coding STATE-COUNTY RECODE would cause such drastic shape changes in our data, we wish to avoid doing so. Fortunately, this variable, though given as a categorical variable, is actually an ordinal variable. There is an ordering among the (state\_county) columns, name longitude, latitude, and elevation. We can trans-

STATE-COUNTY RECODE	address	elevation	lat	lng
35001	Bernalillo+county+NM	5207.579772	35.017785	-106.629130
35003	${\tt Catron+county+NM}$	8089.242628	34.151517	-108.427605
35005	${\tt Chaves+county+NM}$	3559.931671	33.475739	-104.472330
35006	${\it Cibola+county+NM}$	6443.415570	35.094756	-107.858387
35007	$_{\rm Colfax+county+NM}$	6147.749089	36.579976	-104.472330

**Table 2**. Example of the transformation of STATE-COUNTY RECODE to elevation, lat, and lng.

form the data in STATE-COUNTY RECODE into three new numerical columns: lat, lng, and elevation.

For example, Table (2) shows how five entries of STATE-COUNTY RECODE corresponding to counties within New Mexico would can be represented by the elevation, lat, and lng features.

It is a simple exercise to construct the full lookup table from the SEER STATE-COUNTY RECODE variable to the corresponding three values elevation, lat, and lng. Using the publically available dafafile from the United States Census Bureau [3] to construct query strings like the values of the address field in Table (2), it is possible to then programmatically query the Google Maps Geocoding API for the latitude and longitude [6], and the Google Maps Elevation API for the corresponding elevation [5]. An added benefit of this shift from the single categorical variable STATE-COUNTY RECODE to the three continuous numerical variables lat, lng, and elevation is that input into the web applications described later are not restricted to the states and counties covereed in the SEER registries. The full lookup table analogous to Table 2 is available from a GitHub repository containing supplemental information for this study [8].

## 2.3 Colon Cancer Data

In this section we describe the data processing steps that were specific to the colon cancer model development. The four COLRECT.txt files were imported into a pandas DataFrame object. This data was then filtered according to the conditions in Table 3.

The following categorical features were one-hot encoded as described in section 2.2:

- SEX,
- MARITAL STATUS AT DX,
- RACE/ETHNICITY,
- SPANISH/HISPANIC ORIGIN,
- GRADE,
- PRIMARY SITE,
- LATERALITY,
- SEER HISTORIC STAGE A,

Column	Filter
SEQUENCE NUMBER-CENTRAL	eq "Unspecified"
AGE AT DIAGNOSIS	eq "Unknown age"
BIRTHDATE-YEAR	eq "Unknown year of birth"
YEAR OF DIAGNOSIS	$\geq 2004$
SURVIVAL MONTHS FLAG	= "1"
CS TUMOR SIZE EXT/EVAL	≠ ""
CS TUMOR SIZE	$\neq 999$
SEER RECORD NUMBER	= 1
PRIMARY SITE	= "LARGE INTESTINE, (EXCL. APPENDIX)"
SEQUENCE NUMBER-CENTRAL	= 0

**Table 3**. Filters applied to the Colon Cancer data.

- HISTOLOGY RECODE-BROAD GROUPINGS,
- MONTH OF DIAGNOSIS,
- VITAL STATUS RECODE.

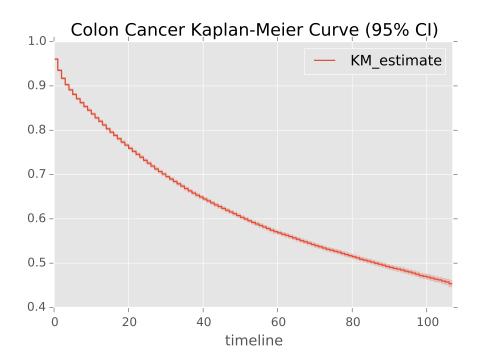
The STATE-COUNTY RECODE variable was dropped and replaced with the elevation, lat, and lng variables as illustrated in Table 2.

With just the above data preparation, it is possible to construct traditional Kaplan-Meier estimates of the survival curves for the colon cancer population represented by this subset of the data. After the above one-hot encoding procedure, the new variable vital\_status\_recode\_Dead indicates that the patient is deceased if this variable = 1, or else that the patient's record is right-censored if this variable = 0. SURVIVAL MONTHS and vital\_status\_recode\_Dead are all that is needed to construct the Kaplan-Meier estimate shown in Figure (1).

#### 2.4 Lung Cancer Data

In this section we describe the data processing steps that were specific to the lung cancer model development. The four RESPIR.txt files were imported into a pandas DataFrame object. This data was then filtered according to the conditions in Table 4. The same list of categorical features as in the colon cancer case were then one-hot encoded.

With just the above data preparation, it is possible to construct traditional Kaplan-Meier estimates of the survival curves for the colon cancer population represented by this subset of the data. After the above one-hot encoding procedure, the new variable vital\_status\_recode\_Dead indicates that the patient is deceased if this variable = 1, or else that the patient's record is right-censored if this variable = 0. SURVIVAL MONTHS and vital\_status\_recode\_Dead are all that is needed to construct the Kaplan-Meier estimate shown in Figure (2).



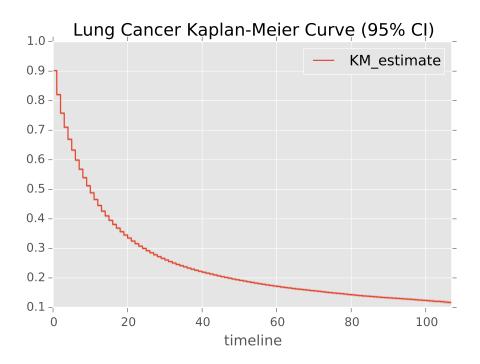
**Figure 1**. Traditional Kaplan-Meier estimate of the survival curve for all colon cancer patients. Fitted with 113072 observations, 71804 censored.

Column	Filter
SEQUENCE NUMBER-CENTRAL	eq "Unspecified"
AGE AT DIAGNOSIS	eq "Unknown age"
BIRTHDATE-YEAR	eq "Unknown year of birth"
YEAR OF DIAGNOSIS	$\geq 2004$
SURVIVAL MONTHS FLAG	= "1"
CS TUMOR SIZE EXT/EVAL	≠ ""
CS TUMOR SIZE	$\neq 999$
SEER RECORD NUMBER	=1
PRIMARY SITE	= "LUNG & BRONCHUS"
SEQUENCE NUMBER-CENTRAL	= 0

Table 4. Filters applied to the Lung Cancer data.

#### 2.5 Breast Cancer Data

In this section we describe the data processing steps that were specific to the lung cancer model development. The four BREAST.txt files were imported into a pandas DataFrame object. This data was then filtered according to the conditions in Table 5. The same list of categorical features as in the colon cancer case were then one-hot encoded.

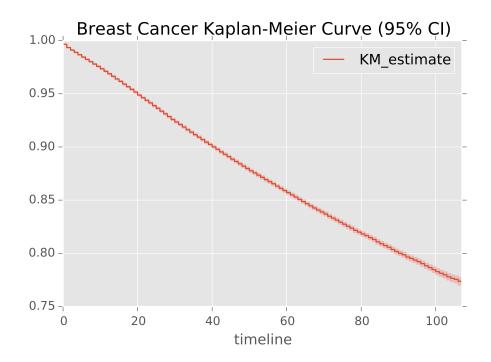


**Figure 2**. Traditional Kaplan-Meier estimate of the survival curve for all lung cancer patients. Fitted with 177089 observatins, 47409 censored.

Column	Filter
SEQUENCE NUMBER-CENTRAL	eq "Unspecified"
AGE AT DIAGNOSIS	eq "Unknown age"
BIRTHDATE-YEAR	eq "Unknown year of birth"
YEAR OF DIAGNOSIS	$\geq 2004$
SURVIVAL MONTHS FLAG	= "1"
CS TUMOR SIZE EXT/EVAL	≠ " "
CS TUMOR SIZE	$\neq 999$
SEER RECORD NUMBER	=1
SEQUENCE NUMBER-CENTRAL	=0

**Table 5**. Filters applied to the Breast Cancer data.

With just the above data preparation, it is possible to construct traditional Kaplan-Meier estimates of the survival curves for the colon cancer population represented by this subset of the data. After the above one-hot encoding procedure, the new variable vital\_status\_recode\_Dead indicates that the patient is deceased if this variable = 1, or else that the patient's record is right-censored if this variable = 0. SURVIVAL MONTHS and vital\_status\_recode\_Dead are all that is needed to construct the Kaplan-Meier estimate



**Figure 3**. Traditional Kaplan-Meier estimate of the survival curve for all breast cancer patients. Fitted with 329949 observatins, 292279 censored.

shown in Figure (3).

# 3 Machine Learning Survival Analysis with Censored Data

## 4 Prediction Models

atex

## 4.1 Decision Trees and Random Forests

## 4.2 MLP Neural Networks

### 5 Performance Metrics

# 5.1 Model Agreement

TO DO: Check if comorbidites are contributing to the outliers in the agreement boxplots that follow. Could mesh with the prevous work [15].

# 6 Web Applications

## 7 Further Directions

Discussion of causality. A certain Marital status is not a "cause" of a better prognosis; c.f. Simpson's Paradox. Implementation of Judea Pearl's Causality Calculus.

Model	6 Months AUC	12 Months AUC	60 Months AUC
Breast RF	.846	.885	.844
Breast $NN$	.855	.867	.836
Colon RF	.804	.806	.828
Colon NN	.797	.804	.841
Lung RF	.772	.796	.874
${\rm Lung}~{\rm NN}$	.765	.796	.875

**Table 6**. AUC values for the Random Forest and Neural Networks model binary classifiers derived from the full survival curve predictions; see text for details.

Cancer Type	% agreement 6 months	% agreement 12 months	% agreement 60 months
Colon	.981	.971	.915
$\operatorname{Breast}$	.994	.984	.938
Lung	.861	.883	.900

**Table 7**. Percentage agreement for the Random Forest and Neural Network classifiers for 6, 12, and 60 month survival predictions on the test data for each cancer type.

#### A Selected Features

In this Appendix we explicitly list the features chosen for each of the Colon, Breast and Lung cancer predictive models. For each cancer type, the features chosen for the random forest and neural network models were the same, so as to be best able to compare the two models. IPython notebooks explicitly providing all code, as well as html versions of the notebooks, are available from a GitHub repository providing supplemental material for thus study [8].

#### A.1 Colon Cancer Feature Selection

The feature set used as input into both the Random Forest and Neural Network models, after the transformation described in Section is given below and also available in full detail in the file NewPatientColonML.html.

- cs\_tumor\_size
- elevation
- grade\_cell type not determined
- grade moderately differentiated
- grade poorly differentiated
- grade undifferentiated; anaplastic
- grade well differentiated
- histology recode broad groupings acinar cell neoplasms

- histology recode broad groupings adenomas and adenocarcinomas
- histology recode broad groupings blood vessel tumors
- histology recode broad groupings complex epithelial neoplasms
- histology recode broad groupings complex mixed and stromal neoplasms
- histology recode broad groupings cystic, mucinous and serous neoplasms
- histology recode broad groupings ductal and lobular neoplasms
- histology recode broad groupings epithelial neoplasms, NOS
- histology recode broad groupings fibromatuos neoplasms
- histology recode broad groupings germ cell neoplasms
- histology recode broad groupings lipomatous neplasms
- $\bullet \ histology\_recode\_broad\_groupings\_miscellaneous \ bone \ tumors$
- histology recode broad groupings myomatous neoplasms
- $\bullet \ \ histology\_recode\_broad\_groupings\_neuroepitheliomatous\ neoplasms$
- histology\_recode\_broad\_groupings\_nevi and melanomas
- histology\_recode\_broad\_groupings\_paragangliomas and glumus tumors
- histology recode broad groupings soft tissue tumors and sarcomas, NOS
- histology recode broad groupings squamous cell neoplasms
- histology recode broad groupings synovial-like neoplasms
- histology recode broad groupings transistional cell papillomas and carcinomas
- histology recode broad groupings unspecified neoplasms
- lat
- laterality Left: origin of primary
- laterality Not a paired site
- laterality Only one side involved, right or left origin unspecified
- laterality\_Paired site, but no information concerning laterality; midline tumor
- laterality Right: origin of primary
- lng
- $\bullet \ \ marital\_status\_at\_dx\_Divorced$
- marital status at dx Married (including common law)
- $\bullet \ marital\_status\_at\_dx\_Separated$
- marital status at dx Single (never married)
- marital status at dx Unknown
- marital\_status\_at\_dx\_Unmarried or domestic partner
- $\bullet$  marital\_status\_at\_dx\_Widowed
- month of diagnosis Apr
- month\_of\_diagnosis\_Aug
- month of diagnosis Dec
- month\_of\_diagnosis\_Feb
- month\_of\_diagnosis\_Jan
- month of diagnosis Jul
- month of diagnosis Jun
- month of diagnosis Mar
- month of diagnosis May

- month of diagnosis Nov
- month\_of\_diagnosis\_Oct
- month of diagnosis Sep
- number of primaries
- race ethnicity Amerian Indian, Aleutian, Alaskan Native or Eskimo
- race ethnicity Asian Indian
- race ethnicity Asian Indian or Pakistani
- race ethnicity Black
- race ethnicity Chinese
- race ethnicity Fiji Islander
- race ethnicity Filipino
- race ethnicity Guamanian
- race ethnicity Hawaiian
- race\_ethnicity\_Hmong
- $\bullet$  race\_ethnicity\_Japanese
- $\bullet$  race\_ethnicity\_Kampuchean
- race ethnicity Korean
- race ethnicity Laotian
- race ethnicity Melanesian
- race ethnicity Micronesian
- race ethnicity New Guinean
- race ethnicity Other
- race ethnicity Other Asian
- race ethnicity Pacific Islander
- race\_ethnicity\_Pakistani
- race ethnicity Polynesian
- race ethnicity Samoan
- race ethnicity Thai
- race ethnicity Tongan
- $\bullet$  race\_ethnicity\_Unknown
- $\bullet$  race\_ethnicity\_Vietnamese
- $\bullet \ \ race\_ethnicity\_White \\$
- seer historic stage a Distant
- seer\_historic\_stage\_a\_In situ
- seer historic stage a Localized
- seer\_historic\_stage\_a\_Regional
- seer historic stage a Unstaged
- sex Female
- spanish hispanic origin Cuban
- spanish hispanic origin Dominican Republic
- spanish hispanic origin Mexican
- spanish hispanic origin Non-Spanish/Non-hispanic
- spanish hispanic origin Other specified Spanish/Hispanic origin (excludes Domini-

can Repuclic)

- spanish hispanic origin Puerto Rican
- spanish\_hispanic\_origin\_South or Central American (except Brazil)
- spanish hispanic origin Spanish surname only
- spanish hispanic origin Spanish, NOS; Hispanic, NOS; Latino, NOS
- spanish hispanic origin Uknown whether Spanish/Hispanic or not
- year of birth
- year\_of\_diagnosis
- month
- newtarget

## A.2 Lung Cancer Feature Selection

The feature set used as input into both the Random Forest and Neural Network models, after the transformation described in Section is given below and also available in full detail in the file NewPatientLungML.html.

- cs tumor size
- elevation
- grade cell type not determined
- grade moderately differentiated
- grade poorly differentiated
- grade undifferentiated; anaplastic
- grade well differentiated
- histology recode broad groupings acinar cell neoplasms
- histology\_recode\_broad\_groupings\_adenomas and adenocarcinomas
- histology recode broad groupings blood vessel tumors
- histology recode broad groupings complex epithelial neoplasms
- histology recode broad groupings complex mixed and stromal neoplasms
- histology recode broad groupings cystic, mucinous and serous neoplasms
- histology recode broad groupings ductal and lobular neoplasms
- histology recode broad groupings epithelial neoplasms, NOS
- $\bullet \ histology\_recode\_broad\_groupings\_fibroepithelial\ neoplasms$
- $\bullet \ \ histology\_recode\_broad\_groupings\_fibromatuos\ neoplasms$
- histology\_recode\_broad\_groupings\_germ cell neoplasms
- histology recode broad groupings gliomas
- histology\_recode\_broad\_groupings\_granular cell tumors & alveolar soft part sarcomas
- histology\_recode\_broad\_groupings\_lipomatous neplasms
- histology recode broad groupings miscellaneous bone tumors
- histology recode broad groupings miscellaneous tumors
- histology recode broad groupings mucoepidermoid neoplasms
- histology recode broad groupings myomatous neoplasms
- histology recode broad groupings myxomatous neoplasms

- histology recode broad groupings nerve sheath tumors
- histology recode broad groupings neuroepitheliomatous neoplasms
- histology recode broad groupings nevi and melanomas
- histology recode broad groupings osseous and chondromatous neoplasms
- histology recode broad groupings paragangliomas and glumus tumors
- histology\_recode\_broad\_groupings\_soft tissue tumors and sarcomas, NOS
- histology recode broad groupings squamous cell neoplasms
- histology recode broad groupings synovial-like neoplasms
- histology recode broad groupings thymic epithelial neoplasms
- $\bullet \ histology\_recode\_broad\_groupings\_transistional\ cell\ papillomas\ and\ carcinomas$
- $\bullet \ histology\_recode\_broad\_groupings\_trophoblastic\ neoplasms$
- histology\_recode\_broad\_groupings\_unspecified neoplasms
- lat
- laterality\_Bilateral involvement, lateral origin unknown; stated to be single primary
- laterality\_Left: origin of primary
- laterality Not a paired site
- laterality\_Only one side involved, right or left origin unspecified
- laterality Paired site, but no information concerning laterality; midline tumor
- laterality\_Right: origin of primary
- lng
- marital\_status\_at\_dx\_Divorced
- marital status at dx Married (including common law)
- marital status at dx Separated
- marital status at dx Single (never married)
- marital\_status\_at\_dx\_Unknown
- marital status at dx Unmarried or domestic partner
- marital status at dx Widowed
- month of diagnosis Apr
- month of diagnosis Aug
- month of diagnosis Dec
- month of diagnosis Feb
- $\bullet$  month\_of\_diagnosis\_Jan
- month of diagnosis Jul
- $\bullet \hspace{0.1cm} \text{month\_of\_diagnosis\_Jun}$
- $\bullet \hspace{0.1cm} \text{month\_of\_diagnosis\_Mar}$
- month\_of\_diagnosis\_May
- month\_of\_diagnosis\_Nov
- $\bullet \hspace{0.1cm} month\_of\_diagnosis\_Oct$
- $\bullet$  month\_of\_diagnosis Sep
- number of primaries
- race ethnicity Amerian Indian, Aleutian, Alaskan Native or Eskimo
- race ethnicity Asian Indian
- race ethnicity Asian Indian or Pakistani

- race ethnicity Black
- race ethnicity Chamorran
- race ethnicity Chinese
- race ethnicity Fiji Islander
- race ethnicity Filipino
- race ethnicity Guamanian
- race ethnicity Hawaiian
- race\_ethnicity\_Hmong
- race ethnicity Japanese
- $\bullet$  race\_ethnicity\_Kampuchean
- ullet race\_ethnicity\_Korean
- race ethnicity Laotian
- race ethnicity Melanesian
- $\bullet$  race\_ethnicity\_Micronesian
- race\_ethnicity\_New Guinean
- race\_ethnicity\_Other
- race ethnicity Other Asian
- race ethnicity Pacific Islander
- race ethnicity Pakistani
- race ethnicity Polynesian
- race ethnicity Samoan
- race ethnicity Thai
- $\bullet$  race\_ethnicity\_Tongan
- race ethnicity Unknown
- race\_ethnicity\_Vietnamese
- race ethnicity White
- seer historic stage a Distant
- seer historic stage a In situ
- seer historic stage a Localized
- seer historic stage a Regional
- seer historic stage a Unstaged
- sex Female
- spanish hispanic origin Cuban
- spanish\_hispanic\_origin\_Dominican Republic
- spanish hispanic origin Mexican
- spanish\_hispanic\_origin\_Non-Spanish/Non-hispanic
- spanish\_hispanic\_origin\_Other specified Spanish/Hispanic origin (excludes Dominican Repuclic)
- $\bullet \ {\rm spanish\_hispanic\_origin\_Puerto} \ {\rm Rican}$
- spanish hispanic origin South or Central American (except Brazil)
- spanish hispanic origin Spanish surname only
- spanish hispanic origin Spanish, NOS; Hispanic, NOS; Latino, NOS
- spanish hispanic origin Uknown whether Spanish/Hispanic or not

- year of birth
- year of diagnosis
- month
- newtarget

#### A.3 Breast Cancer Feature Selection

The feature set used as input into both the Random Forest and Neural Network models, after the transformation described in Section is given below and also available in full detail in the file NewPatientBreastML.html.

- cs tumor size
- elevation
- grade moderately differentiated
- grade\_poorly differentiated
- grade\_ndifferentiated; anaplastic
- grade well differentiated
- $\bullet$  histology\_recode\_broad\_groupings\_adenomas and adenocarcinomas
- histology recode broad groupings adnexal and skin appendage neoplasms
- histology recode broad groupings basal cell neoplasms
- histology recode broad groupings complex epithelial neoplasms
- histology recode broad groupings cystic, mucinous and serous neoplasms
- $\bullet~histology\_recode\_broad\_groupings\_ductal~and~lobular~neoplasms$
- $\bullet \ histology\_recode\_broad\_groupings\_epithelial\ neoplasms,\ NOS$
- histology recode broad groupings nerve sheath tumors
- histology\_recode\_broad\_groupings\_unspecified neoplasms
- lat
- laterality Bilateral involvement, lateral origin unknown; stated to be single primary
- laterality Paired site, but no information concerning laterality; midline tumor
- laterality Right: origin of primary
- lng
- marital stats at dx Divorced
- marital\_stats\_at\_dx\_Married (inclding common law)
- marital stats at dx Separated
- marital\_stats\_at\_dx\_Single (never married)
- marital stats at dx Unknown
- marital\_stats\_at\_dx\_Unmarried or domestic partner
- marital stats at dx Widowed
- month of diagnosis Apr
- month of diagnosis Aug
- $\bullet$  month\_of\_diagnosis\_Dec
- month of diagnosis Feb
- month of diagnosis Jan
- month of diagnosis Jul

- month of diagnosis Jun
- $\bullet$  month\_of\_diagnosis\_Mar
- month\_of\_diagnosis\_May
- month of diagnosis Nov
- month of diagnosis Oct
- month of diagnosis Sep
- race ethnicity Amerian Indian, Aletian, Alaskan Native or Eskimo
- race ethnicity Asian Indian
- race ethnicity Black
- race ethnicity Chinese
- $\bullet \ \ race\_ethnicity\_Japanese \\$
- race ethnicity Melanesian
- race ethnicity Other
- race\_ethnicity\_Other Asian
- race\_ethnicity\_Pacific Islander
- race ethnicity Thai
- race ethnicity Unknown
- race ethnicity Vietnamese
- $\bullet$  race\_ethnicity\_White
- seer\_historic\_stage\_a\_Distant
- seer historic stage a In sit
- seer historic stage a Localized
- $\bullet$  seer\_historic\_stage\_a\_Unstaged
- sex Female
- spanish\_hispanic\_origin\_Cuban
- spanish hispanic origin Mexican
- spanish hispanic origin Non-Spanish/Non-hispanic
- spanish\_hispanic\_origin\_Other specified Spanish/Hispanic origin (excldes Dominican Republic)
- spanish hispanic origin Spanish surname only
- spanish hispanic origin Spanish, NOS; Hispanic, NOS; Latino, NOS
- year of birth
- year of diagnosis
- month
- newtarget

# B Model Architecture and Python Code

# C GitHub Repositories

Please always give a title also for appendices.

## Acknowledgments

This is the most common positions for acknowledgments. A macro is available to maintain the same layout and spelling of the heading.

**Note added.** This is also a good position for notes added after the paper has been written.

#### References

- [1] Ben Kuhn. Decision trees for survival analysis. http://www.benkuhn.net/survival-trees, year="2016 (accessed 14 Jan 2016)",.
- [2] Michael Bowles. Machine Learning in Python: Essential Techniques for Predictive Analysis. Wiley, 2015.
- [3] United States Census Bureau. 2010 fips code files for counties geography u.s. census bureau. https://www.census.gov/geo/reference/codes/cou.html, 2016 (accessed 18 Jan 2016).
- [4] Cam Davidson-Pilon. Quickstart lifelines 0.8.0.1 documentation. http://lifelines.readthedocs.org/en/latest/Quickstart.html, 2016 (accessed 14 Jan 2016).
- [5] Google Developers. The google maps elevation api | google maps elevation api | google developers. https://developers.google.com/maps/documentation/elevation/intro?hl=en, 2016 (accessed 18 Jan 2016).
- [6] Google Developers. The google maps geocoding api | google maps geocoding api | google developers. https://developers.google.com/maps/documentation/geocoding/intro, 2016 (accessed 18 Jan 2016).
- [7] Allen Downey. Think Stats. O'Reilly Media, 2014.
- [8] IOBS. Supplemental material | paperdata. https://github.com/doolingdavid/PAPERDATA.git, 2016 (accessed 18 Jan 2016.
- [9] Rabbert Klein. Black holes and their relation to hiding eggs. Theoretical Easter Physics, 2010. (to appear).
- [10] Johann A. Makowsky, Saharon Shelah, and Jonathan Stavi.  $\Delta$ -logics and generalized quantifiers. *Annals of Mathematical Logic*, 10:155–192, 1976.
- [11] National Cancer Institute, the Surveillance, Epidemiology, and End Results Program. About the SEER Program SEER. http://seer.cancer.gov/about, 2016 (accessed 14 Jan 2016).
- [12] National Cancer Institute, the Surveillance, Epidemiology, and End Results Program. Documentation for ASCII Text Data Files SEER Datasets. http://seer.cancer.gov/data/documentation.html, 2016 (accessed 15 Jan 2016).
- [13] Sebastian Raschka. Python Machine Learning Essentials. Packt Publishing, 2015.
- [14] Hyunjung Shin and Yonghyun Nam. A coupling approach of a predictor and a descriptor for breast cancer prognosis. BMC MEDICAL GENOMICS, 7(1), MAY 8 2014. 3rd Annual Translational Bioinformatics Conference (TBC) / ISCB-Asia, Seoul, SOUTH KOREA, OCT 02-04, 2013.

[15] Hamed Majidi Zolbanin, Dursun Delen, and Amir Hassan Zadeh. Predicting overall survivability in comorbidity of cancers: A data mining approach. DECISION SUPPORT SYSTEMS, 74:150–161, JUN 2015.

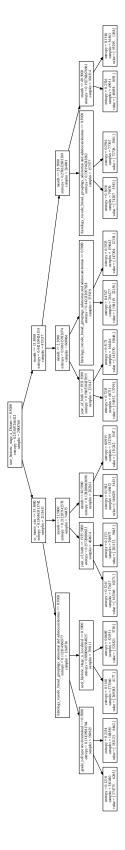


Figure 4. The top levels of a decision tree trained on the Lung Cancer training data.

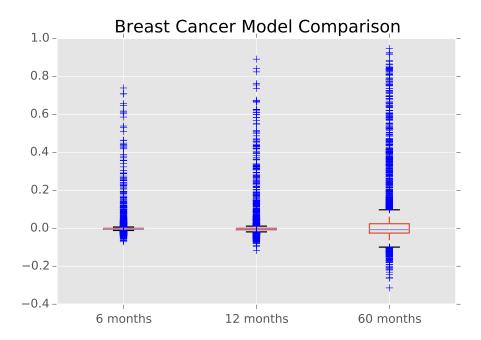
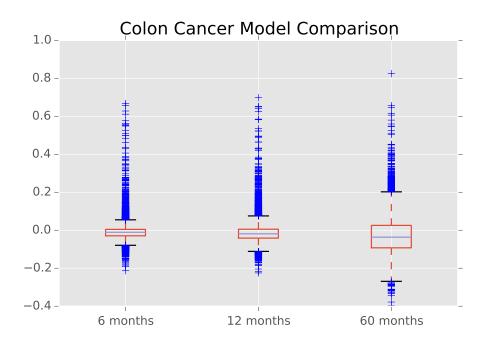


Figure 5. Box plots showing the distributions of the signed difference between the MLP model's prediction for the probability of surviving 6 months and the Random Forest model's prediction of the same quantity for breast cancer. The plot shows the same quantity for the 12 and 60 months classifiers. It is apparent from the figures that the outliers are due to the neural network models predicting higher survival probablitlies than the random forest for some few cases. These differences were evaluated for the 3300 test patients in the breast cancer data.



**Figure 6**. Box plots showing the distributions of the signed difference between the MLP model's prediction for the probability of surviving 6 months and the Random Forest model's prediction of the same quantity for colon cancer. The plot shows the same quantity for the 12 and 60 months classifiers. It is apparent from the figures that the outliers are due to the neural network models predicting higher survival probablitlies than the random forest for some few cases. These differences were evaluated for the 5654 test patients in the colon cancer data.

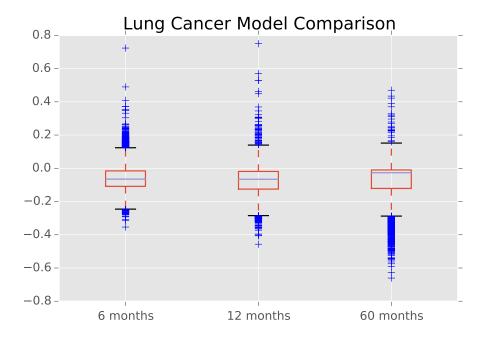


Figure 7. Box plots showing the distributions of the signed difference between the MLP model's prediction for the probability of surviving 6 months and the Random Forest model's prediction of the same quantity for lung cancer. The plot shows the same quantity for the 12 and 60 months classifiers. These differences were evaluated for the 5654 test patients in the colon cancer data. The Interquartile Ranges for lung cancer are visibly larger than those for breast cancer and colon cancer shown in fig 5 and fig 6.