Lung Cancer Prognosis System using Data Mining Techniques

Yomna Omar, Abdullah Tasleem, Michel Pasquier and Assim Sagahyroon

American University of Sharjah, Department of Computer Science & Engineering
PO Box 26666, Sharjah, U.A.E.

Abstract: This paper describes a Lung Cancer Prognosis System (LCPS) that aims at providing oncologists with an accurate estimate of the health status of their patients. The proposed system is born from two observations: First, lots of efforts are still required in healthcare to improve productivity, accuracy, etc. by providing ad hoc computer-based solutions; second, while increasing popular, AI and data mining tools cannot be used without significant training and expertise. LCPS thus aims at providing the former by integrating the latter into a user-friendly tool, supplementing the knowledge of the expert oncologist with information about their patients, and leading to improved patient care and treatments. LCPS can accept a variety of lung cancer datasets and employs several data mining algorithms to uncover relationships between observed health signs and probable outcomes, and provides oncologists with various statistical results including predictions about their patients’ medical future. Furthermore, LCPS makes it easy to manage patients’ records, allows them view their profiles and any information as deemed suitable by their doctor, including prognosis and other comments. Lastly, while the current application is currently limited to lung cancer treatment, it can be considered a prototype that can be adapted to other diseases.

1 INTRODUCTION

According to scientific experts, cancer remains one of the leading causes of death in adults and the “number one disease killer of children”, and half of them are preventable (American Cancer Society, 2017). Lung cancer has the highest rate of deaths, despite the fact that its prevention and treatment in its early stages is simple and hassle-free. The problem is that early symptoms are often mistaken for cough or flu, thus making early-stage lung cancer difficult to diagnose. Since the initial symptoms of lung cancers can take years to develop, most patients discover their illness at an advanced stage where cancer cells are widespread, making it much harder to cure (American Cancer Society, 2016). Lung cancer is also one of the top five fatal diseases in the UAE (Khaleej Times, 2016) and, according to Dr. Ali Al Dameh, Oncology Consultant from Tawam John Hopkins Hospital, “the disease is in dire need of being addressed with a solid plan of action” (WAM, 2015). That goal includes the two main types of lung cancer i.e., non-small cell lung cancer and small cell lung cancer.

In order to decrease preventable cases of cancer-related deaths and improve the overall health system, a more versatile and powerful technology is needed. Precisely AI and data mining techniques can process health data to better forecast people’s future health conditions and potential risks. But advanced AI tools cannot be used without significant training and expertise, which medical practitioners do not possess. Therefore, we developed a prototype of a Lung Cancer Prognosis System (LCPS) that aims at making those techniques accessible in a user-friendly manner in order to augment the knowledge of the expert oncologist and improve patient care and treatments. The proposed system is trained beforehand on a hospital dataset characteristic of the target population and can subsequently provide better information and prognosis specific to each lung cancer patient based on their vital health signs and test results. LCPS employs several statistical and data mining tools and presents their results in a visual, intuitive manner, for the benefit of both the physicians and their patients.

2 EXISTING SOLUTIONS

GenieMD combines IBM’s Watson with research from Harvard Medical School to provide users with accurate self-triage. It records medical history and can connect to sensors for real-time vitals tracking...
(HMW, 2013), but cannot upload other datasets. WebMD allows users to search through a medical database to read articles, learn about diagnosis, and contact medical doctors (WebMD, 2017). However, this service does not process actual patient data. Sehaty is an app that allows Dubai residents to view their records at the Dubai Health Authority, including prescriptions, appointments, test results, medical advice, and so on (Khaleej Times, 2014). Similarly, My Medical allows users to create a record of their medical history, enter vital sign readings from health trackers, and send the data to a doctor. While tailored for the UAE, these apps do not provide any kind of diagnosis or prognosis tools.

Lungscreen uses a short lifestyle questionnaire to assess whether the person has lung cancer symptoms or not (Mesko, 2015), and gives a prediction for the next 10-20 years. There is no clear way to assess prediction accuracy, and it does not use actual patient data. The Lung Cancer Foundation App is a mobile application that helps with the self-diagnosis of lung cancer and provides educational videos and information (OFWW, 2011). Finally, Cancer Spotter aims to encourage earlier diagnosis by getting people to learn and understand the symptoms of cancer, and to seek help from a doctor as soon as possible (Park, 2001). While it is simple and promotes cancer awareness, it relies on self-assessment and has no diagnosis or prognosis.

In sum, most available tools provide lung-cancer related information and some allow self-assessment, but very few offer any form of medically validated, automated diagnosis or prognosis, based on actual lung cancer data and the patient’s vital signs, let alone tailored to a specific country or population.

3 PROPOSED SOLUTION: LCPS

Based on the above assessment, we have developed a software system to provide users with a consistent and accurate health diagnosis at low cost, therefore facilitating the prognosis of fatal lung cancer. We plan to make use of large databases of lung cancer patients from hospitals all around the world and apply AI and data mining tools to derive from these datasets diagnosis and prognosis models that can be subsequently applied to new patients. Another goal is to integrate all computational techniques and automate the process as much as possible to offer a tool that is user-friendly for both doctors and their patients. Initially, our system was designed to use a patient database from the Tawam Hospital in UAE, then it was extended to handle other datasets. In this paper, the use cases and results presented employ lung cancer datasets publicly available on the Web.

LCPS is a web-based app that uses JSF for functionality and HTML/CSS for webpages and UI. This first version integrates three machine learning algorithms from WEKA. It provides oncologists and patients with different access rights. Patients can view their data, health predictions, and comments from the oncologist. Doctors manage patients, can run various tests on the dataset and new patients data. The LCPS prototype described in this paper provides doctors with a number of public datasets for demonstration and training purposes: one is used to predict patient life expectancy, another to predict whether or not the patient has a cancer gene. LCPS allows doctors to load and edit datasets and patient records, to view statistical indicators, and to apply various algorithms on patient data. It automatically analyses a dataset to determine the the most suitable machine learning algorithm and presents the results to the physician through many tables and plots.

4 TECHNICAL APPROACH

LCPS is a 3-tiered web application, with a back-end SQL database; an application layer; and a front-end user interface. LCPS will make use of several data mining and machine learning algorithms that have been trained with patient data to reach conclusions about the patient’s health within a certain level of accuracy. LCPS can also use other datasets to allow flexibility in predicting several lung cancer markers.

By creating several XHTML pages that are styled by Cascading Style Sheets (CSS), the web app’s main functions lie in JSF code with baking beans to implement UI elements, display options, and access the database. Pages that contain graphs use the Google Charts API via embedded JavaScript code, more specifically jQuery. The app’s database access mainly occurs in baking beans using SQL statements to insert, update, and delete data. WEKA offers multiple ways to integrate their machine learning algorithms using Java. For our purposes, we initially focused on three algorithms: J48 decision tree, Naïve Bayes (NB), and K-Nearest Neighbor (KNN). Further algorithms will be added later on.

5 SYSTEM ARCHITECTURE

LCPS architecture consists of three layers for presentation, application, and data. This helps
developers accommodate future expansions such as different subcomponents in the data layer (Oracle database, MySQL, Firebase) or even in the presentation layer (Java GUI, C# GUI, Python GUI). Importantly, it minimizes dependencies between components, enhances security (as the user has no direct access to the database) and promotes scalability which is a key aspect in a cloud deployed app. Lastly, it allows the three tiers to be coded and tested in parallel, and therefore reduces development and maintenance costs over the long term.

![Figure 1: LCPS system multi-layered architecture.](image)

The presentation layer consists of either command line or GUI, which are used to get user inputs and provide information and feedback. The presentation layer consists of a web based interface programmed using HTML and CSS. The application layer consists of the decision logic i.e., the preprocessing and machine learning algorithms from WEKA that will mine the dataset selected by the oncologist and evaluate new patient data accordingly. The application uses the SQL database on Apache Derby to store its data. The tables and schema can be managed via the interface, or programatically. The node architecture can be displayed through the use of a deployment diagram, which captures the system hardware’s topology. Each layer is deployed within a different hardware component and the presentation layer is mainly deployed via the web interface.

Since one distinguishing feature of LCPS is to provide automated data analysis and prognosis, the choice of data mining component has been subject to extensive study (Witten, 2016). Moreover, the need to offer a user-friendly tool that hides technical complexity and presents results in an intuitive, visual manner, means favoring models based on rules or decision trees for instance, and providing adequate visualizations via tables and plots. As such many algorithms are available, as well as several data mining platforms, that we experimented with.

6 DATA MINING TOOLS

We started with KEEL (Knowledge Extraction based on Evolutionary Learning), an open source data mining framework that is increasingly being used in healthcare. KEEL includes many algorithms for regression, classification, clustering, and more. It allows creating experiments using multiple datasets and algorithms, independently scripted from the user interface. However, we didn’t find the UI user-friendly enough, and KEEL lacks good visualization techniques (Rangra, 2014).

KNIME (Konstanz Information Miner) is another open source tool based on data analytics, reporting and integrating platform, and has been used extensively in pharmaceutical research. It has a modular design and offers interactive execution and data pipelines. It covers all major areas of data analysis but has limited error measurement methods and does not support techniques such as wrapper methods for descriptor selection (Rangra, 2014).

Eventually we selected the open source WEKA framework, which has all the desired data processing and machine learning algorithms and is widely used for research and development. It has a great user interface and provides excellent visualization tools to help understand the models (Witten, 2016).

7 SYSTEM FUNCTIONALITY

The user interface was initially modeled using Adobe Dreamweaver through the use of HTML and CSS. Once tested and approved by users, the design was transferred to the development environment, where core functions were added in. LCPS employs several web interface mechanisms, as depicted in the figures hereafter, to enhance usability.

Upon login and redirection to their home page, an oncologist can view all his/her respective patients, further analyze their medical profile, add comments, deciding for each whether it should be visible to the patient or not. The oncologist can also compare the patient’s health data against a preloaded dataset or one of his/her choosing. The doctor can also edit existing datasets and view previous versions. When the physician compares his/her patient’s data against a chosen dataset using selected algorithms, he/she will be provided with a prediction of the patient’s health status. The physician can also compare results from different algorithms in order to gain a better understanding of the data and the various models.

When a cancer patient logs into LCPS, he/she is redirected to his/her main page where he/she can view
all the tests conducted on him/her by his/her respective oncologist. The patient can also view any comments left by the physician on his/her health status, as well as data plots and results as included.

8 DATA MINING ALGORITHMS

WEKA features hundreds of algorithms for data processing, feature selection, clustering, finding association rules, classification, etc. (Witten, 2016). LCPS integrates selected methods for removing outliers and irrelevant features, and predicting the patients' health status using classification rules, decision trees, instance based learning, probabilistic approach, and later regression trees. Since choosing the “best” predictive algorithm for a given dataset requires a lot of computations, automation is needed. This in turns requires making assumptions, which is another reason why the current system supports databases for lung cancer only.

Including decision trees in LCPS was an obvious choice, and currently the J48 algorithm is used (Witten, 2016). The decision tree generated shows the most relevant input variables for prediction, that are determined using information gain. As an illustrative example, we chose a dataset obtained by Harvard University (Zhu, 2007), that contains 189 instances with all numeric values except for the class attribute (ADEN, SQUA, COID, SCLS, NORMAL). It includes different specimens of lung cancer using various gene selection tests and classified as tumors. Accuracy results and Confusion Matrix are shown and explained to the doctor. The latter table further describes the performance of the classification model. In this example, the user is told that 131 out of the 189 instances were correctly classified as adenocarcinoma (ADEN), 18 as squamous cell lung carcinomas (SQUA), 20 as pulmonary carcinoids (COID), 4 as small-cell lung carcinomas (SCLC) and that 16 instances were normal lung patients. Other values outside the diagonal show exactly how some instances were incorrectly classified. The data can be examined by the oncologist, if desired, to try and analyze the cause of the error, which could be due to some data entry mistake, some noise in measurements, or simply some borderline case that was too difficult for the classifier to figure out. Furthermore, the Detailed Accuracy by Class shows that the TP rate of true positives (correctly classified instances) has a high average of 0.931 while the FP rate of false positives (falsely classified instances) had a very low average of 0.058. Such results are pointed out to the doctor so he/she can appreciate diagnosis or prognosis accuracy, possibly examine the data to further his/her understanding. In this example, J48 achieved a 93% accuracy and new patients can be evaluated with high confidence. The decision tree in Fig. 4 is shown to the doctor.

Another popular classifier is Naïve Bayes (NB), which realizes probabilistic inference based on assumptions of conditional independence between predictors (Witten, 2016). One of the reasons we chose to include it is that, despite its simplicity, it often outperforms more sophisticated classifiers. Another is that it can be easily explained to the doctor. Naive Bayes is essentially a quick method for composition of statistical predictive models, that analyses the subjection between attribute values and classes to derive a conditional probability.
class probabilities then gives an indication of which result is most likely. Using the same lung cancer dataset, we can observe a loss in the TP rate and a gain in the FP rate as shown in Fig. 5.

Figure 5: Classifier output of Naïve Bayes algorithm.

Instance-based learning is a form of lazy learning where the model memorizes data instances and classification relies on measuring proximity (Witten, 2016). To address the issue of noisy data, it is recommended to examine multiple nearest neighbors, hence LCPS integrates the classic K-Nearest Neighbor algorithm. Results using the same lung cancer dataset are shown in Fig. 6 for the case of a single neighbor (K=1). However, LCPS will automatically try different values of K and retain the optimal one. Explanations are given to the oncologist who wishes to understand the significance of the results. For instance, if the number of correctly classified instances increases with K, it means that the dataset is noisy, which in turn may require examination. In this case results show that the dataset is noise-free.

LCPS can display only the best results (achieved by the best algorithm), or show all results, as well as a detailed comparison, in tabular format. In this case, it happens that J48 performs best on the sample lung cancer dataset. There might be various reasons for this, such as for instance, the fact that irrelevant gene predictors have been pruned out, or because it can process both nominal and numerical attributes better than Naïve Bayes and KNN, or because decision trees handle missing values well. It is important to note that there is no classification algorithm that can perform best for all available datasets. Hence LCPS will apply all selected methods and build models anew for each dataset.

Figure 6: Classifier output of the KNN algorithm (K=1).

9 DATASETS AND RESULTS

Since LCPS can handle generic lung cancer datasets, we chose two databases available online for testing and illustration purposes. The Genes database used to predict lung cancer tumors based on the patient’s genes was cleaned and used at Harvard University, hence we knew it was mostly error free and would give reliable results, making for a good tutorial. The database consists of 12600 attributes and 203 instances, all numerical values except for the class. In 2001, lung carcinoma was the leading cause of cancer
death hence this dataset was created and exploited to define distinct subclasses of lung adenocarcinoma (Bhattacharjee, 2001). The dataset can be used by doctors so long as they have a gene expression profile, and LCPS will determine if the patient has the genes to develop either cancer type.

<table>
<thead>
<tr>
<th>Regression Variable</th>
<th>Log Normal</th>
<th>Weibull</th>
<th>Log F (p = 0)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>$\hat{\beta}$</td>
<td>S.E(\hat{\beta})</td>
<td>$\hat{\beta}$</td>
</tr>
<tr>
<td>No prior therapy</td>
<td></td>
<td></td>
<td>(q = 0.43)</td>
</tr>
<tr>
<td>(97 patients)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Performance status</td>
<td>0.030</td>
<td>0.006</td>
<td>0.022</td>
</tr>
<tr>
<td>Squamous vs. large</td>
<td>-0.085</td>
<td>0.34</td>
<td>0.175</td>
</tr>
<tr>
<td>Small vs. large</td>
<td>-0.762</td>
<td>0.31</td>
<td>-0.521</td>
</tr>
<tr>
<td>Adeno. vs. large</td>
<td>-0.804</td>
<td>0.34</td>
<td>-0.640</td>
</tr>
<tr>
<td>Prior therapy</td>
<td></td>
<td></td>
<td>(q = 1.05)</td>
</tr>
<tr>
<td>(40 patients)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Performance status</td>
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<td>0.010</td>
<td>0.054</td>
</tr>
<tr>
<td>Squamous vs. large</td>
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<td>0.46</td>
<td>0.428</td>
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<tr>
<td>Small vs. large</td>
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<td>0.49</td>
<td>-0.944</td>
</tr>
<tr>
<td>Adeno. vs. large</td>
<td>-0.694</td>
<td>0.61</td>
<td>-0.787</td>
</tr>
</tbody>
</table>

The second dataset included for testing is the Veteran’s Administration Lung Cancer Trial dataset used by Venables and Ripley in 2002 to predict the life expectancy of a patient with specific lung cancer tumors (Venables, 2002). It was created using questionnaires from doctors and patients. Initially, a Generalized F-regression model was used to analyze it, as per Fig. 7. Data collection and comparison of statistical models was first done manually, then the Cox model was applied to establish the baseline hazards, as the dataset had several covariates.

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The dataset obtained needed cleaning prior use: outliers were removed and nonsensical or missing values were replaced with mean values, irrelevant attributes were discarded. In the end the dataset had seven attributes: Treatment (standard or test), Age (in years), Karnofsky Score (0-100), Time since diagnosis (in months); Cell type (one of four types); Prior therapy; Survival time (in days, later discretized into 3 bins). The Karnofsky score is one of the methods used to quantify a patient’s general health and physical activities, ranging from 0 (dead) to 100 (perfect health). Results are illustrated in the screenshots of the LCPS user interface hereafter.

As mentioned the decision logic layer of our system heavily relies on WEKA and its excellent Java API, which offers many classes and methods, specifically: to store all data instances, to preprocess the data (discretize, sparse to non-sparse, and attribute selection), to predict the class outcome (J48, Naïve Bayes, KNN), to evaluate accuracy (statistical results, TP/FP, ROC). To integrate WEKA in our system we had to use beans to get information from our XHTML pages and process the data using the provided libraries.

Once logged in, a doctor can select a patient and go to the Analysis page where he/she can upload any lung cancer related dataset to the system, as depicted in Fig. 9. Since a consistent, generic dataset format is required, the doctor is able to view and download the data template. Converters are available as well that help with the formatting. Next, the doctor can view all uploaded datasets, run data mining tools as desired, examine results via tables, plots, and related explanations. If technically inclined, the doctor can customize a number of filters. While discretization...
happens automatically if needed, one can specify which method and how many bins to use. As for attribute selection, PCA is automatically be applied but one can select other methods or even manually specify which attributes to use for classification and which to discard e.g., based on expert knowledge.

From this page, the doctor can modify the database or view statistical information about the dataset, as depicted in the sample screens of Fig. 10. First, the analysis page shows the accuracy of correct classifications using the highest scoring algorithm. LCPS can also display statistics about how the algorithms perform and how they differ in their analysis and predictive model.

Next is shown a table listing the various correctly and incorrectly classified instances in separate classes. Next, the ROC curves and bar charts show the true positives (TP) and false positives (FP) and total of correct and incorrect values. Moreover, after examining these graphs and results, the doctor can still select any available algorithm, examine results in further details, play “what if” scenarios by changing some data and applying the algorithm anew. We aim to implement automated explanatory mechanism later on. Lastly, Fig. 11 shows the prediction page for a fictitious patient using the life expectancy dataset. In this case, the result reported is a bracket of 0–333 days to live. This can be regarded by the doctor as fatal, and will be added to the patient’s record for later use. The oncologist can always inspect the result and calculation details.

LCPS heavily uses three types of visual tools to help oncologists understand and appreciate the generated results. Scatter plots as especially useful to help identify anomalies in the dataset, which the doctor can then decide to edit manually, or to show how attribute values are distributed so the doctor can determine if a patient profile is typical or not. Pie charts show how many instances are correctly classified in each class along with the total incorrectly classified overall. Others show incorrect classifications in each class. Pie charts are separate
for each predictive algorithm and can be used for direct comparison as well. If the doctor wishes to see if a patient falls into a specific class, he can examine the details of each algorithm and the result values. Line charts are also used often. For example, the TP vs. FP rate line graph shows the ROC area which can be used to identify visually how accurate each prediction is. Explanations are given as well: in this case, the more convex the ROC curve the better, whereas a straight line would denote a random (hence useless) classifier. The ROC curves are shown separately for each algorithm, so that the doctor can easily see how each algorithm performs. Lastly, Column charts are best used to depict and compare visually results such all the different TP and FP rate for each algorithm and the overall correct and incorrect classifications.

10 CONCLUSIONS

As mentioned earlier, cancer is one of the leading causes of death in the UAE. In the emirate of Abu Dhabi alone, 1,729 new cases were detected in 2012, with 28 per cent among Emiratis and 72 per cent among expatriate residents. This made cancer the third leading cause of death in Abu Dhabi, accounting for 13% of all fatalities (HAAD, 2016). This death toll will keep on increasing if we do not devise a method to find cancer at an early, curable stage. The UAE has recently launched an initiative to cut down the number cancer cases by 18% by 2021 (HAAD, 2016), hence we hope that a system such as LCPS will be a highly contributing factor.

The early detection of any type of cancer is of paramount importance to pave the way for successful cancer treatment. Unfortunately, most cancers are only detected once they reach an advanced and incurable stage. Individuals who are victims of many types of cancer do not know about it until it's too late. Many people do not even go to the doctor to get themselves checked for several reasons which can include, affordability, fear, travelling cost or even time. Everyone is so absorbed in their work that they disregard the possibility of having cancer, even after the symptoms start to manifest. Hence, this was one of the driving factors for us to make such an application. To be able to measure how much of a societal impact it can have we need to evaluate its impact on both general users and doctors. Therefore, by focusing on just UAE-based users, our proposed application is predicted to have a significant impact on the country as a whole. By focusing on the patient-specific aspect of LCPS, it becomes more apparent why having a tailored prediction, that is specific to the patient’s health, is important. It gives the oncologist and patient room to forecast and define a targeted treatment process, helping in better patient care and improving the patient’s chances of survival and remission. Thus, having such a system will decrease lung cancer death cases significantly, especially since there are no similar systems in the market.

REFERENCES

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