Gene Selection using a Hybrid RFE Along with LASSO for Cancer Classification

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Abstract: Gene expression profiling using microarray technology has done with the chip based phenomena. For studying gene expression data are more helpful in knowing various diseases and more useful in finding diseases. Recently in the bioinformatics field, cancer prediction using gene expression data had made the assuring area. Samples having the gene attributes will not surely give the efficient amount of classification. Overcoming these contribution, a strong method is required for selecting the relevant gene features for building the classification model effectively. Basically least absolute shrinkage and selection operator (LASSO) and Recursive feature elimination (RFE) are automatic gene feature selection methods used for classification. Here in our proposed work, we use these two methods as a hybrid one for selecting the features and later it applied into the Support vector machine (SVM) for easy classification. It made best when compared to the existing techniques by their performance measures, were regulated on six publically available cancer datasets. Just out it gives the good awareness in the selection of features.

Keywords: LASSO, Gene selection, RFE, SVM, Cancer Classification.

I. INTRODUCTION

Recently, the microarray technology is used for measuring the abundant amount of genes arranged in a single chip. It helpful in measuring the genes with expression values. These gene expression levels indicate the varying number of gene Ribonucleic Acid (RNA). For analyzing these genes expression is normally difficult. It causes redundancy. Gene expression measure is the transfer of DNA into RNA protein molecule, while using the microarray technology causes the dimensionality issue for measuring the genes [1]. For some classification problems using gene expression profiling it is hard to use the traditional methods straightforward. It leads to the dimensionality issue. So we need to use the feature selection it reduces the redundancy and the dimensionality. Towards it reduces the redundancy level of the gene expression data [2, 3]. The statistical learning methods such as SVM was first charted by Vapnik et.al, in 1960 mainly for classifying the data. SVM classifies the large datasets which will be in the linear or non-linear, by constructing the hyperplane. By separating the data in two subsets among these data closer to the hyperplane are called as support vectors. And draw a margin between the hyperplane where the good separation of data is attained by greatest distance named as functional margin for non-linear kernel methods are also applicable [4]. Cancer classification is made using the genetic programming by the gene expression data [5]. L.Wang et al. and Statnikov A., et al. [6, 7] proposed the classification, using support vector machines done in two phases i.e., gene selection and classification. An experiment has been handled on human acute leukemia’s for cancer classification with the help of gene expression microarrays to finding the class discovery acute myeloid leukemia (AML) and acute lymphoblastic leukemia (ALL) for predicting and finding the classes [8]. Using the three feature selection methods filter, wrapper and embedded are used in classifying the predictive accuracy with the help of the significant features [9]. By using support vector machine, the tissue samples of cancer diseases are validated[10]. RFE-SVM is used for choosing the suited features and later the given datasets are changed to two subsets for further classification. And then the result validation is carried out to get the goodness of the features. Finally, RFE-SVM yields the better results [11]. Two stages of SVM-RFE method are used for feature selection because to avoid inconsistency. The noisy, irrelevant genes are eliminated in the first stage by the pre-filtering process. And the elimination of the single gene is carried in each step is occurred in the next stage. Later classification is done by the linear SVM and this method is competed along with the correlation method [12]. The functional margin that maximizes the hyper-plane for classification using support vector machine [13]. Tibshirani et.al. proposed the method for regularization and the selection of variables for linear regression called LASSO [14]. Gene selection and class prediction by SLR and SLRB, among the high dimensional of genes most of them are most of them are noisy, irrelevant. Later the deficiency of LASSO causes consistency [15]. An automatic gene selection method is carried with Dantzig selector and LASSO technique, this method is more supportive in choosing the significant genes with the linear regression method [16]. Here in our work we have to suggest a newfangled method for selecting features as hybrid recursive feature elimination (RFE) with least absolute shrinkage and selection operator (LASSO) and later classification is done using SVM. This feature selection method yields the better one compare to the other available methods. The surplus of the work is prepared with gene selection method, recommended work, results and discussions append with conclusion and future work.
II. FEATURE SELECTION METHODS

A. RFE

RFE is comes under the wrapper based feature selection method [13], by the sequential backward elimination the waited features are selected, among all the features it removes a single feature. The feature with good ranks are selected and it removes the unimportant feature. For every step the weight vector of coefficients are helps to calculate the feature score in every step of the linear SVM. In SVM – RFE the objective function k will enhance in changing the removal of insignificant genes.

\[ k = \frac{\|w\|^2}{2} \]

B. LASSO

Tibshirani et al. [14] proposed the regression method called LASSO, it performs selection of variables and the regularization of variables for linear regression. It correlates to ordinary least squares (OLS) with residual sum of squares (RSS) that reduces the sum where the constant value is less while compared with the absolute values. And as an additional constraint, it is slightly similar to ridge regression with the sum of squared values as coefficients to the larger constraint. So the simple modification occurred during the LASSO in the variable selection. It set the coefficients values to zero during the variable selection, it automatically punishes the extra features shrunnk to zero entirely.

\[ Lasso = \min \left( \sum_{i=1}^{n} (y_i - \beta_0 - \sum_{j=1}^{p} \beta_j x_{ij})^2 + \lambda \sum_{j=1}^{p} |\beta_j| \right) \]

Where \( \beta_j \) denotes full least square estimates
\( x_{ij} \) denotes predictor variables
\( y_i \) denotes output response
\( \beta_0 \) is usually ‘0’ & not necessary, will be neglected

III. PROPOSED WORK

A. Hybrid RFE Based LASSO

In association with the combination of SVM-RFE and LASSO is the new approach. Better performance gives by SVM-RFE on classification but it gives the poor performance in the repetitions of class labels. In concord with the LASSO it evades the redundancy level. The combination of SVM-RFE with LASSO will attain their good in performance. The figure 1 displays the proposed work carried in this scenario. Using RFE the significant features are selected by calculated with the threshold value and the irrelevant features get eliminated and later the selected features get regularized and the coefficients value are set to zero automatically during the selection of good features by LASSO. Among them, that 70% features are carried for training during classification and the remaining 30% are used for testing under the evaluation process in classification.
Table I: Dataset Description

| Datasets          | Number of samples | Number of Genes | Class       |
|-------------------|-------------------|-----------------|-------------|
| Leukemia[8]       | 72                | 7129            | ALL         |
| Colon[17]         | 62                | 2000            | Tumor       |
| Lung Cancer[18]   | 181               | 12533           | MPM         |
| Breast[19]        | 97                | 24481           | Tumor       |
| Prostate[20]      | 136               | 12600           | Tumor       |
| Liver[21]         | 156               | 1648            | Non-Tumor   |

Table II: Comparative performance measure for RFE and LASSO

| Algorithm          | Precision | Recall | Kappa statistics | Feature Selection accuracy |
|--------------------|-----------|--------|------------------|---------------------------|
| RFE and BBF        | 0.95      | 0.93   | 0.95             | 92.55                     |
| SFS and LASSO      | 0.96      | 0.95   | 0.96             | 94.45                     |
| RFE and LASSO      | 0.97      | 0.96   | 0.97             | 97.61                     |

Table III: Comparative feature selection accuracy for RFE and LASSO

| Algorithm          | Feature Selection accuracy |
|--------------------|----------------------------|
| RFE and BBF        | 88.55                      |
| SFS and LASSO      | 95.45                      |
| RFE and LASSO      | 97.61                      |

Table IV: Classification model performance with different datasets

| Classifier Model   | Leukemia | Colon | Lung Cancer | Breast | Prostate | Liver |
|--------------------|----------|-------|-------------|--------|----------|-------|
| ANN                | 62       | 51    | 65          | 72     | 75       | 63    |
| KNN                | 74       | 63    | 85          | 87     | 62       | 83    |
| SVM                | 92       | 86    | 94          | 95     | 81       | 92    |

IV. RESULTS AND DISCUSSION

Here, we are using six publically available datasets for summarization mentioned in table I. For our approach, the performance metrics is checked using precision, recall, classification accuracy. This evaluation measure, gives the better outcomes in finding the results. Here, precision denotes the positive predictor value. Recall is called as sensitivity in binary classification. Kappa statistics is the performance measure is used to check the interrelationship between the two variables whether it having good agreement or less agreement. And the classification accuracy is measured by the absolute number of the checked features divided by the total number of features. Below equations (1, 2, 3) as,

\[
\text{Precision} = \frac{TP}{TP + FP}
\]

\[
\text{Recall} = \frac{TP}{TP + FN}
\]

\[
\text{Accuracy} = \frac{Total\ observed\ (prediction)}{Total\ entered\ (prediction)}
\]

Fig. 2. Graphical representation for RFE and LASSO

The proposed work is analyzed using R programming language. It is a statistical programming language. Recently, much statistical analysis based on machine learning, deep learning techniques are evaluated using this language. The classification accuracy along with conventional SVM using feature selection method SVM-RFE and LASSO have attained the better results as 97.61%. Likewise, 70% of selected features are used for training with the classifier and remaining 30% is used for testing the classifier model. The results obtained are observed with feature selection methods SVM-RFE and BBF, SFS and LASSO using the conventional SVM. The table 2 denotes the comparative feature selection accuracy for RFE and LASSO. And fig 2. shows the graphical representation for that.

Table- III: Comparative feature selection accuracy for RFE and LASSO

Table- IV: Classification model performance with different datasets

Compared with the existing methods RFE and BBF, SFS and LASSO table 3 shows the feature selection accuracy results of the RFE and LASSO have attained 97.61%. Fig 3. denotes the graphical representation for that. Along with that, the training and testing done with the classifier model such as ANN, KNN and SVM among these SVM will attained the better results for six datasets as laid out in table 4. With their graph representation in figure 4.
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V. CONCLUSION

Cancer based classification using gene expression data is an assuring research recently. Here in our paper, a compound approach using the SVM-RFE and LASSO is used for variable selection. After that, this method combined with SVM for classification. This is the novel approach had applied in the six datasets. And later it checks for the precision, recall and classification accuracy. This work results is compared with other classifier models such as ANN, KNN. By using this approach, it reduces the redundancy level. In future, this approach is compared with the other techniques such as deep learning, reinforcement learning using large dimensional tools such as Hadoop, Spark.

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