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# Voxelwise Detection of Cerebral Microbleed in CADASIL Patients by Naive Bayesian Classifier

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Abstract—It is important to detect cerebral microbleed voxels from the brain image of cerebral autosomal-dominant arteriopathy with subcortical infarcts and Leukoencephalopathy (CADASIL) patients. Methods developed by other researchers before have a high variability of intra-observer and inter-observer. In our study, we collect our dataset from the 20 brain volumetric images, 10 for CADASIL patients and 10 for healthy controls. And we used naive baysian classifier to get the results. We use cross validation to improve the performance of naive Baysian classifier. The results show that the average sensitivity is  $74.53\pm0.96\%$ , the average specificity is  $74.51\pm1.05\%$ , and the average accuracy is  $74.52\pm1.00\%$ .

Keywords—CADASIL; voxel; naive Baysian classifier; cross validation

#### I. INTRODUCTION

The cerebral autosomal-dominant arteriopathy with subcortical infarcts and Leukoencephalopathy (CADASIL) syndrome is the most common hereditary stroke diseases which can never be ignored.

The subclinical sign of cerebral microbleed (CMB) [1] of CADASIL patients can be detected years prior to clinical manifestation by magnetic resonance imaging (MRI). Susceptibility weighted imaging (SWI) is a 3D flow-compensated T2\* imaging technique. Compared to standard MRI protocol, the SWI scans the patient at high resolution (less than 1 mm). It uses the phase image to enhance the contrast, providing 2 to 6 times increased sensitivity in CMB detection.

Manual detection results of CMB location of CADASIL patients were regarded as the ground truth. However, it takes too much time to detect with a high variability of intra-observer and inter-observer. A large amount of advanced methods based on computer vision and digital image processing were proposed to help manual CMB evaluation. <u>Hou (2017) [2]</u> used leaky rectified linear unit (ReLU). <u>Chen (2016) [3]</u> utilized sparse deep neural network. <u>Hou (2018) [4]</u> employed autoencoder. Jiang (2017) [5] used rank-based average pooling. Lu (2017) [6] presented to use deep convolutional neural network.

Though methods mentioned above are very important attempts in the past, they are not acceptable for 2 reasons: (1) The complexity of these methods are very high. (2) Their detection do not reach voxel-wise resolution.

Our method used naive Bayesian theory to perform the classification task. For general classification tasks, if we have

enough training data and the features are well designed, then the accuracy will be high enough to accept. It estimates the posterior probability by priori probability collected for training. And the mathematical foundations for Bayesian theory provides solid explain ability without much complexity. Besides, our purpose it to identify CMB at voxel resolution. In all, our methods is a new computer vision [7-10] and image processing [11-14] approaches.

#### II. METHODOLOGY

Naive Bayesian classifiers are classifiers based on statistical analysis [15]. They can predict class membership probabilities i.e. the probability that the sample with given values of its features belongs to a specific category [16]. Naive Bayesian classifiers assume that the effect of an attribute value on a given category is independent of the values of the other attributes. This assumption is called class conditional independence. It is made to simplify the computations involved and, in this sense, is considered "naive."

Naïve Bayesian classifier can present a comparable performance to state-of-the-art classifiers, such as support vector machine [17-21], shallow neural network [22-27], etc.

Let X be a sample which is described by measurements made on a set of n attributes. Let H be some hypothesis such as that the data sample X belongs to a specified category C. For classification problems, we want to determine P(H|X), the probability that the hypothesis H holds given the observed sample X. In other words, we are looking for the probability that sample X belongs to category C, given that we know the attribute description of X.

P(H|X) is the posterior probability of H conditioned on X. P(H) is the prior probability. Similarly, P(X|H) is the priori probability of X conditioned on H, and P(X) is the prior probability of X. Bayes' theorem is useful in that it provides a way to calculate the posterior probability, P(H|X), from P(H), P(X|H), and P(X). Bayes' theorem is [28]:

$$P(H|X) = \frac{P(X|H) * P(H)}{P(X)}.$$
 (1)

Let *D* be a training set of samples and their associated class labels. each sample is represented by an *n*-dimensional attribute vector,  $X = (x_1, x_2, ..., x_n)$ , depicting *n* measurements made on the sample from *n* attributes, respectively, A<sub>1</sub>, A<sub>2</sub>, ..., A<sub>n</sub>.



Suppose that there are *m* categories,  $C_1$ ,  $C_2$ , ...,  $C_m$ . Given a sample *X*, the classifier will predict that *X* belongs to the category having the highest posterior probability [29], conditioned on *X*. That is, the naive Bayesian classifier predicts that sample *X* belongs to the class  $C_i$  if and only if

$$P(C_i|X) > P(C_j|X) \text{ for } 1 \le j \le m, j \ne i.$$
(2)

By Bayes' theorem, we have

$$P(C_i|X) = \frac{P(X|C_i) * P(C_i)}{P(X)}.$$
 (3)

Since P(X) is a constant for all categories, only  $P(X|C_i)*P(C_i)$ needs to be determined. Note that the class prior probabilities may be estimated by

$$P(C_i) = \frac{|C_{i,D}|}{|D|} \tag{4}$$

where  $|C_{i,D}|$  is the number of training samples of class  $C_i$  in D and |D| is the total number of training samples of D.

Given data sets with many attributes, it would be extremely computationally expensive to compute  $P(X|C_i)$ . To reduce computation in evaluating  $P(X|C_i)$ , the naive assumption of class-conditional independence is made. This presumes that the attributes' values are conditionally independent of one another, given the class label of the sample (i.e., that there are no dependence relationships among the attributes). Thus, we can obtain  $P(X|C_i)$  by following formula:

$$P(X|C_{i}) = \prod_{k=1}^{n} P(x_{k}|C_{i}).$$
(5)

To predict the class label of X,  $P(X|C_i)*P(C_i)$  is evaluated for each class  $C_i$ . The classifier predicts that the class label of sample X is the class  $C_i$  by formula (2) mentioned above.



FIGURE I. ILLUSTRATION OF NAÏVE BAYESIAN CLASSIFIER

Figure 1 shows the illustration of naïve Bayesian classifier. We did not use deep learning methods, because this dataset is relatively small, and the convolution operation [30-35] is difficult to handle on an 7x7-size input.

## III. MATERIAL AND PREPROCESSING

Figure 2 shows a portion of generated dataset. It can be regarded as a matrix with width of 50 and height of 50. Each row of this matrix represents a sample, the first 49 columns represent input data and the 50-th column represents the target. Input data were generated by vectorizing sliding neighborhood with size of 7\*7.



FIGURE II. GENERATED DATASET (COLUMN 1-49 ARE INPUTS, COLUMN 50 IS THE TARGET)

All the pixels in the 7x7 neighbor were regarded as features, and submitted to the naïve Bayesian classifier. We collected in total 20 subjects, and obtained 69,356 CMB voxels, and 124,063,981 non-CMB voxels. For class imbalance problem,

we selected randomly 69,327 non-CMB voxels from those 124,063,981 samples. Now we have a dataset as listed in Table 1.

#### TABLEI. OUR DATASET

Туре	Number of samples
CMB	69,365
Non-CMB	69,327
Total	138,692

A 10-fold cross validation was used to segment the dataset into 10 folds, and report the out-of-sample error. For avoiding the effect of randomness, we run the 10-fold cross validation ten times, and report the average and standard deviation of sensitivity, specificity, and accuracy. The definition of those three measures are depicted by following three equations:

Sensitivity 
$$= \frac{TP}{TP+FN}$$
 (6)

Sepcifity 
$$= \frac{TN}{FP+TN}$$
 (7)

$$Accuracy = \frac{TP + TN}{TP + FN + FP + TN}$$
(8)

where TP *represents* true positive, TN represents true negative, FP represents false positive, and FN represents false negative.

Figure 3 shows the illustration of one run of cross validation. Figure 3(a) shows the index, where the x-axis is the combination of both CMB and non-CMB voxels, in total 138,692. Figure 3(b) shows the legend



# **IV. RESULTS**

The sensitivities, specificities, and accuracies of this 10x10fold cross validation were shown below in Table 2, The average sensitivity is  $74.53\pm0.96\%$ , the average specificity is  $74.51\pm1.05\%$ , and the average accuracy is  $74.52\pm1.00\%$ . Figure 4 shows the box plot of our algorithm, from which we can observe the three measures can similar results, indicating the effectiveness of our technique to handle the imbalance problem.

TABLEII. PERFORMANCE OF OUR NBC METHOD

Sen	F1	F2	F3	F4	F5	F6	F7	F8	F9	F10	Total
R1	77.4	74.3	75.9	76.3	78.7	76.3	75.5	75.3	75.1	79.0	76.4
	5	1	1	8	1	3	7	5	2	9	2
R2	70.9	77.3	75.5	75.8	76.1	69.4	76.3	75.4	73.4	72.8	74.3
	2	5	8	9	3	5	8	8	0	3	4
R3	73.9	77.2	70.4	74.6	71.1	75.3	73.8	73.4	71.2	74.8	73.6
	4	5	3	7	8	7	5	6	1	0	2
R4	73.8	73.1	75.6	74.5	73.1	72.0	73.8	70.7	72.4	71.2	73.0
	5	8	3	1	4	5	3	5	7	8	7
R5	75.4	74.2	71.6	76.9	75.0	74.3	76.4	74.5	75.1	75.3	74.9
	3	5	1	4	8	5	3	6	2	0	1
R6	74.4	75.3	73.0	74.8	71.8	75.3	75.8	77.2	76.3	75.5	74.9
	1	2	9	6	1	1	4	9	0	5	8
R7	75.6	76.6	70.8	76.9	73.5	74.2	78.2	74.5	78.2	72.1	75.1
-	2	4	8	5	9	1	3	4	9	6	1
R8	71.6	73.8	74.1	72.2	74.4	72.4	76.2	71.8	73.6	75.0	73.5
DO	/	6	/	5	8	/	/	1	6	0	6
R9	73.2	/5.0	73.0	17.4	11.2	13.2	74.9	73.8	72.6	//.8	/4.8
D 1	3	1	7 1	1	2	0	2	3	8	4	4
КI 0	70.0	13.9	/0.1	13.3	13.0	73.9	6	/0.0	70.5	74.5	74.4
Spa	J E1	2 E2	1 E2	5	0 E5	2 E6	E7	7	FO	2 E10	Total
Spc	75.2	Γ2 76.6	T5 75.0	Γ4 74	T5 70.7	T0	Γ/ 7( )	T0 74.0	T9	F10	76.0
KI	15.5	/0.0 5	/5.8	/0.0 5	/8./	//.4	/0.2 2	74.8	/4.0	/0.4 5	/0.2 0
Dγ	5 75 3	5 74.6	4 75 /	5 726	74.0	74.0	5 747	75.2	4753	5 70 1	0 7/3
K2	8	1	9.4	2	9	3	3	1	5	1	74.5 A
R3	74.0	72.5	743	- 73 7	73.0	69.8	71.2	78.2	71.9	74.6	733
K5	5	9	4	0	1	5	4	8	8	1	7
R4	757	74.8	71.0	70.2	753	74.0	67.8	72.2	737	72 5	727
	2	7	3	0	2	1	9	8	6	7	7
R5	-76.4	77.4	72.2	74.6	-76.7	76.8	72.9	74.7	72.4	77.6	75.2
	3	4	1	3	9	9	4	0	8	0	1
R6	74.2	73.8	73.7	75.8	73.1	77.6	74.7	75.0	74.2	78.3	75.0
	7	2	7	8	0	3	9	8	0	2	8
R7	73.6	76.1	75.0	75.8	76.6	77.3	70.5	75.8	75.2	73.7	75.0
	9	7	6	7	3	7	9	0	1	6	1

R8	71.6	73.5	75.8	73.9	73.4	72.2	70.7	74.7	74.7	76.4	73.7
	0	3	5	8	0	3	5	7	3	7	3
R9	76.2	75.0	70.8	76.4	73.0	77.3	73.9	76.4	74.8	78.0	75.2
	7	5	5	0	1	4	7	2	4	9	2
R1	71.0	73.6	74.2	74.8	73.5	72.5	75.9	76.6	74.8	73.5	74.0
0	2	6	7	4	9	2	0	7	7	6	9
Ac	F1	F2	F3	F4	F5	F6	F7	F8	F9	F10	Total
с											
R1	76.4	75.4	75.8	76.5	78.7	76.8	75.9	75.1	74.8	77.7	76.3
	0	8	8	2	3	6	0	1	8	7	5
R2	73.1	75.9	75.5	74.2	75.5	72.1	75.5	75.3	74.3	71.4	74.3
	5	8	4	6	6	9	6	4	7	7	4
R3	73.9	74.9	72.3	74.1	72.1	72.6	72.5	75.8	71.6	74.7	73.4
	9	2	9	9	0	2	4	7	0	1	9
R4	74.7	74.0	73.3	72.3	74.2	73.0	70.8	71.5	73.1	71.9	72.9
	9	3	3	5	3	3	6	1	1	3	2
R5	75.9	75.8	71.9	75.7	75.9	75.6	74.6	74.6	73.8	76.4	75.0
	3	5	1	8	3	2	8	3	0	5	6
R6	74.3	74.5	73.4	75.3	72.4	76.4	75.3	76.1	75.2	76.9	75.0
	4	7	3	7	5	7	1	8	5	4	3
R7	74.6	76.4	72.9	76.4	75.1	75.7	74.4	75.1	76.7	72.9	75.0
	6	1	7	1	1	9	1	7	5	6	6
R8	71.6	73.7	75.0	73.1	73.9	72.3	73.5	73.2	74.1	75.7	73.6
	3	0	1	1	4	5	1	9	9	4	5
R9	74.7	75.0	71.9	76.9	75.1	75.2	74.4	75.1	73.7	77.9	75.0
	5	3	6	0	2	7	4	2	6	7	3
R1	70.8	73.7	75.1	74.0	73.7	74.2	74.0	76.7	75.7	74.0	74.2
0	3	9	9	9	3	2	8	8	1	4	5



FIGURE IV. BOX PLOT OF OUR ALGORITHM

In the next experiment, we compared NBC with decision tree using ID3 algorithm [<u>36</u>]. The results were shown in Figure 5. Here ID3 obtained a lower performance, with an overall

accuracy of 72.19%. We can observe that our native Bayesian classifier can get 2.33% more performance in terms of accuracy.



FIGURE V. COMPARISON BETWEEN OUR METHOD WITH ID3 ALGORITHM

## V. CONCLUSION

In this study, we proposed a new voxelwise CMB detection system using naive baysian classifier for CADASIL patients. The results showed better result than ID3.

In the future, we will collect more brain images and test other advanced techniques for classification and image preprocessing, to see if we can achieve better accuracy which can meet the requirement of real applications.

We shall test using optimization algorithm [37, 38] to improve the performance of our NBC classifier. In addition, we shall collect more data, and try deep learning methods.

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