Simplified Labeling Process for Medical Image Segmentation

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Abstract. Image segmentation plays a crucial role in many medical imaging applications by automatically locating the regions of interest. Typically supervised learning based segmentation methods require a large set of accurately labeled training data. However, the labeling process is tedious, time consuming and sometimes not necessary. We propose a robust logistic regression algorithm to handle label outliers such that doctors do not need to waste time on precisely labeling images for training set. To validate its effectiveness and efficiency, we conduct carefully designed experiments on cervigram image segmentation while there exist label outliers. Experimental results show that the proposed robust logistic regression algorithms achieve superior performance compared to previous methods, which validates the benefits of the proposed algorithms.

1 Introduction

To assist doctors locate pathologies, automatic segmentation of different regions of medical images is very useful. Supervised learning based segmentation method, which use manually segmented training data as references, has superior performance. Those methods perform well as long as the feature space sufficiently distinguishes each label. They are relatively computationally efficient and not sensitive to parameters. Many popular learning methods have been applied to solve challenging medical problems, such as, but not limited to support vector machine (SVM) [1], neural network [2,3], conditional random field (CRF) [4], logistic regression (LR) [5].

One disadvantage of the learning-based segmentation is the requirement of manual interaction for obtaining training data. Manual segmentation by different people are subjective due to the lack of standard when performing the manual segmentation. In reality, the segmentation training data may not perfectly labeled. However, all the previous methods ignored the imperfection of training data. In such cases the training set is misleading, the guidance given by the labels may not be reliable. Consequently learning results may not totally reliable. On the other hand, the precisely labeling process is time consuming,

laborious and herpaps not necessary. In this paper, we take the advantage of learning methods handling label outliers and propose a simplified lableing process. The proposed simplified labeling process could significantly reduce doctors time and effort in labeling the training data. And the proposed learning methods would handle the labeling errors introduced by the new labeling process. While learning in the idealized setting has been thoroughly studied, learning in the presence of labeling errors has not been well explored. And nothing has been done for medical image segmentation with labeling error.

According to the practical consideration mentioned above, we are the first to systematically study logistic regression and sparse logistic regression method with outliers and apply it on cervigram image segmentation. Logistic regression (LR) is a classical method for classification and has been widely used in many applications. Sparse logistic regression, which is the logistic regression with l_1 -norm regularization, is a very effective method for classification on large scale data with high dimension. Both methods could be adopted in image segmentation.

In this paper, we propose a robust sparse logistic regression method to handle the classification problem with label outliers. In our method, a latent variable is introduced for the true correct labels, and then we estimate the probability of labels being flipped. Conjugate gradient method and Lassplore algorithm [6] are used respectively to minimize the loss function under the noise assumption. After optimization, the probability of the flipped labels is updated using the new classification estimation. The algorithm can quickly converge after several iterations.

To demonstrate the effectiveness of the proposed algorithm, we apply the proposed method on the task of automatically segmenting the biomarker Ace-toWhite (AW) regions in an archive of 60,000 images of the uterine cervix [7]. The most important observation in a cervigram image is the AW region, which is caused by whitening of potentially malignant regions of the cervix epitheliuem, following application of acetic acid to the cervix surface. Since the texture, size and location of AW regions have been shown to correlate with the pathologic grade of disease severity, accurate identification and segmentation of AW regions in cervigrams have significant implications for diagnosis and grading of cervical lesions. Carefully designed experiments on the cervigram images with label outliers demonstrate the superior performance of the proposed method and validate the benefits of the proposed algorithm in practical applications.

2 Related Work

In this section, we briefly review logistic regression and sparse logistic regression.

Logistic regression is a conditional probability of the label y , given a sample $x \in \mathbb{R}^n$, and a weight vector $w \in \mathbb{R}^n$:

$$p(y = 1 \mid x, w) = \sigma(w^T x) = \frac{1}{1 + e^{-w^T x}}. \quad (1)$$

Given a set of training examples $D = \{(x_1, \hat{y}_1), \dots, (x_i, \hat{y}_i), \dots, (x_m, \hat{y}_m)\}$, we want to learn a classifier $y = f(x)$. Here $x_i \in \mathbb{R}^n$ denotes a sample, and $\hat{y}_i$ is the corresponding observed class label. The likelihood function associated with these m samples is defined as

$$\begin{aligned} f(w) &= \log \prod_{i=1}^m p(\hat{y}_i | x_i, w) \\ &= \sum_{i=1}^m \frac{\hat{y}_i + 1}{2} \log(p(\hat{y}_i = 1 | x_i, w)) + \frac{1 - \hat{y}_i}{2} \log(p(\hat{y}_i = -1 | x_i, w)), \end{aligned}$$

which is a smooth and convex function. We can determine w by minimizing the logistic loss:

$$\hat{w} = \underset{w}{\operatorname{argmin}} f(w). \quad (2)$$

This is a smooth convex optimization problem. Regularization is usually used to avoid the overfitting problem. The l_2 -norm regularization leads to a smooth and differentiable unconstrained convex optimization problem. Standard optimization algorithm such as Newton method and conjugate gradient method can be applied for solving such optimization problem [8,9].

The l_1 -norm regularization is used to obtain a sparse model. However, the l_1 -norm regularization term is non-differentiable. There are many algorithms proposed in the past for solving the l_1 regularized logistic regression [10,11]. Liu et al. proposed the Lassplore algorithm for solving large scale sparse logistic regression. They formulate the problem as the l_1 -ball constrained logistical regression formulation, in which the objective function is continuously differentiable, and the problem domain set is closed and convex [6].

3 Proposed Algorithm

We propose a new simplified labeling process. Doctors would label a bounding box around the region of interest rather than precisely label the boundaries. Please see Figure 2 (a,b,c) as illustration of the proposed labeling process. Figure 2(a) is the image for segmentation. As we can see from (b), the groundtruth, the boundary of the region of interest is nontrivial and complicated. Our labeling process is illustration in (c), which would roughly include the region of interest in a bounding box. The simplified labeling process would significantly save time and effort of doctors in labeling the training data.

Considering the new labeling process, which introduces some labeling noise, none of the previous work has been able to handle label outliers. In this paper, we propose a robust logistic regression method to handle the classification problem on large scale data with label outliers.

3.1 Robust Logistic Regression

If there exists some label noise, Eq. 2 is no longer valid. We introduce a latent variable y_i , which represents the true label. The probability of the observed label $p(\hat{y}_i | x_i, w)$ is written as the following:

Table 1. Probabilistic relationship between observed label and true label

		Observed Label	
		-1	1
True Label	-1	γ_{00}	γ_{01}
	1	γ_{10}	γ_{11}

$$\begin{aligned}
S_i^k &\doteq p(\hat{y}_i = k \mid x_i, w) \\
&= \sum_{j=0}^K p(\hat{y}_i = k \mid y_i = j) p(y_i = j \mid x_i, w).
\end{aligned} \tag{3}$$

Here $p(\hat{y}_i = k \mid y_i = j)$ represents the probability that the label has flipped from the true label j to the observed label k . The probability relationship between observed label and true label is represented in Table 1. By expanding the terms in Eq. 3 we find that,

$$\begin{aligned}
S_i^0 &= p(\hat{y}_i = -1 \mid x_i, w) \\
&= p(\hat{y}_i = -1 \mid y_i = -1) p(y_i = -1 \mid x_i, w) + p(\hat{y}_i = -1 \mid y_i = 1) p(y_i = 1 \mid x_i, w) \\
&= \gamma_{00}(1 - \sigma(w^T x)) + \gamma_{10}\sigma(w^T x) \\
S_i^1 &= p(\hat{y}_i = 1 \mid x_i, w) \\
&= p(\hat{y}_i = 1 \mid y_i = -1) p(y_i = -1 \mid x_i, w) + p(\hat{y}_i = 1 \mid y_i = 1) p(y_i = 1 \mid x_i, w) \\
&= \gamma_{01}(1 - \sigma(w^T x)) + \gamma_{11}\sigma(w^T x).
\end{aligned} \tag{4}$$

By substituteing Eq. 4 into Eq. 2, we find:

$$f(w) = \sum_{i=1}^m \frac{\hat{y}_i + 1}{2} \log(S_i^1) + \frac{1 - \hat{y}_i}{2} \log(S_i^0). \tag{5}$$

This formulation remains smooth and differentiable. It can be solved using standard optimization algorithm such as conjugate gradient method [9,8].

3.2 Robust Sparse Logistic Regression

To handle outliers in sparse logistic regression, we need to optimize Eq. 5 subject to the l_1 -ball constrain. We use the Lassplore algorithm to solve the sparse logistic regression problem [6]. The Lassplore algorithm is a first-order black-box method that evaluates the function value and the gradient at each iteration. The gradient of Eq. 5 is the following:

$$\nabla_w f(w) = \sum_{i=1}^m \frac{\hat{y}_i + 1}{2S_i^1} \nabla_w S_i^1 + \frac{1 - \hat{y}_i}{2S_i^0} \nabla_w S_i^0, \tag{6}$$

where $\nabla_w S_n^0$ and $\nabla_w S_n^1$ are:

$$\begin{aligned}\nabla_w S_i^0 &= (\gamma_{10} - \gamma_{00})\sigma(w^T x_i)(1 - \sigma(w^T x_i)) \cdot x_i, \\ \nabla_w S_i^1 &= (\gamma_{11} - \gamma_{01})\sigma(w^T x_i)(1 - \sigma(w^T x_i)) \cdot x_i,\end{aligned}\quad (7)$$

$\sigma(w^T x_n)$ is the conditional probability of $\hat{y}_i$ given a sample x_i .

An initial estimation of the label flipping probability can be given to start the optimization. After solving the Lassplore algorithm, label flipping probability is updated as the following:

$$\begin{aligned}e_i &= \sigma(w^T x_i), \\ \gamma_{00} &= \frac{(e_i < 0.5) \& (\hat{y}_i = -1)}{e_i < 0.5}, \gamma_{01} = 1 - \gamma_{00}, \\ \gamma_{11} &= \frac{(e_i > 0.5) \& (\hat{y}_i = 1)}{e_i > 0.5}, \gamma_{10} = 1 - \gamma_{11},\end{aligned}\quad (8)$$

To conclude, an iterative framework is proposed to update the label-flipping probability. First given an initialization of the label-flipping probability, the Lassplore method is used to minimize the loss function. Then, the label-flipping probability is estimated again using the new classification results. Using this scheme we iteratively update the label-flipping probability and solve the optimization problem.

4 Experimental Results and Discussion

We first use a synthetic dataset to illustrate the effectiveness of our algorithm. The examples are sampled from two multivariate normal distributions differing in mean and covariance. Here we focus on the non-uniform flipping of labels, where

Fig. 1. (a) shows the samples with true labels and use Logistic Regression (LR) for separation. (b) shows the samples with observed labels, where about 20% labels from negative class are flipped to the positive class. Robust algorithms push the classification line to the true boundary line on true labels.

Table 2. Trained on the corrupted training labels, tested on true labels. The proposed methods show better performance.

Method	Sensitivity	Specificity	DSC
LR	0.9307 ± 0.0207	0.9987 ± 0.0027	0.9570 ± 0.0158
rLR	0.9920 ± 0.0040	0.9687 ± 0.0031	0.9809 ± 0.0013
SLR	0.9153 ± 0.0248	0.9987 ± 0.0027	0.9508 ± 0.0158
rSLR	0.9840 ± 0.0033	0.9807 ± 0.0020	0.9833 ± 0.0015

Fig. 2. (d) uses the groundtruth and LR in training for comparison. The simplified labeling data, as shown in (c) are used for training for robust algorithm. (f) and (h) include lots of negative points since we use some of the negative points in training. Use the robust algorithms, the result expelled many negative points.

Table 3. Testing errors on segmenting the cervigram data

Method	Sensitivity	Specificity	DSC
LR using groundtruth	0.8351	0.8820	0.8157
LR	0.9486	0.5660	0.6950
rLR	0.8175	0.8825	0.8058
SLR	0.9354	0.5931	0.7012
rSLR	0.7121	0.8986	0.7517

there are negative data being labeled as positive ones. This setting is to imitating the proposed labeling process in medical images. All the data are training on the corrupted data and then tested on the true labels. We quantitatively evaluate the classification results in Table 2. Compared to logistic regression (LR) and sparse logistic regression (SLR) methods, both robust logistic regression (rLR) and robust sparse logistic regression (rSLR) perform better on sensitivity but worse on specificity. Dice similarity coefficient(DSC), which is defined as $(2 * \text{true positive}) / (2 * \text{true positive} + \text{false positive} + \text{false negative})$, measures the consistency of results and groundtruth. DSC is a more comprehensive measurement of the coincidence of the classification result with the groundtruth. The proposed methods perform better on this measurement.

We apply the proposed algorithm to classify pixels in optical cervigram images into two classes, namely normal and abnormal tissues while not given accurate segmentation training labels. The experiment is designed as the following: We have experts precisely labeled the segmentation, which is considered as groundtruth. We also have experts roughly labeled the segmentation, using our proposed simplified labeling method. The roughly labeled data is used for training, and then we test our segmentation results against the precisely labeled segmentation. Our proposed labeling process and segmentation results would achieve competitive results compared to using the the groundtruth for training. In our experiments, HSV feature is chosen for classification because it achieves the best performance.

Table 3 shows the results of different classifiers measured by sensitivity, specificity and DSC. Our method consistently achieves significantly better performance in terms of DSC. The results from our method also compares favorably with other state-of-the-art methods in this application [12].

5 Conclusion

This paper proposed a novel method on handling labeling noise. The proposed method has been demonstrated on cervigram segmentation problem. Using a bounding box around the region of interest as labels other than accurate labels, we can still get comparable results. In the future, we want to extend our work using group sparsity [13].

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