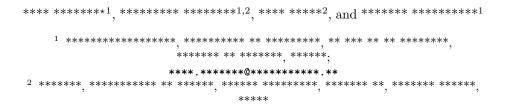
# An optimization approach to segment breast lesions in ultra-sound images using clinically validated visual cues



Abstract. This paper addresses the problem of automatic classification of Spectral Domain OCT (SD-OCT) data for automatic identification of patients with Diabetic Macula Edema (DME) versus normal subjects. Our method is based on Local Binary Patterns (LBP) features to describe the texture of Optical Coherence Tomography (OCT) images and we compare different LBP features extraction approaches to compute a single signature for the whole OCT volume. Experimental results with two datasets of respectively 32 and 30 OCT volumes show that ...(TB BE WRITTEN BASED ON RESULTS)

Moreover, the experiments show that the proposed method achieves better classification performances than other recent published works.

**Keywords:** Diabetic Macula Edema, Optical Coherence Tomography, acsdme, OCT, LBP.

# 1 Introduction

Eye diseases such as Diabetic Retinopathy (DR) and Diabetic Macula Edema (DME) are the most common causes of irreversible vision loss in individuals with diabetes. Just in United States alone, health care and associated costs related to eye diseases are estimated at almost \$500 M [13]. Moreover, the prevalent cases of DR are expected to grow exponentially affecting over 300 M people worldwide by 2025 [18]. Early detection and treatment of DR and DME play a major role to prevent adverse effects such as blindness. Indeed, the detection and diagnosis of retinal diseases are based on the detection of vascular abnormalities or lesions in the retina.

In past decades, Computer Aided Diagnosis (CAD) systems devoted to ophthalmology, have been developed focusing on the automatic analysis of fundus

images [1, 16]. However, the use of fundus photography is limited to the detection of signs which are correlated with retinal thickening such as hard and soft exudates, hemorrhages or micro-aneurysms. However, DME is characterized as an increase in retinal thickness within 1 disk diameter of the fovea center with or without hard exudates and sometimes associated with cysts [7]. Therefore, fundus photography cannot always identify the clinical signs of DME; for example cysts, which are not visible in the retinal surface. In addition, it does not provide any quantitative measurements of retina thickness or information about cross-sectional retinal morphology.

Recently, Optical Coherence Tomography (OCT) has been widely used as a valuable diagnosis tool for DME detection. OCT is based on optical reflectivity and produces cross-sectional and three-dimensional images of the central retina, thus allowing quantitative retinal thickness and structure measurements. The new generation of OCT imaging, namely Spectral Domain OCT (SD-OCT) offers higher resolution and faster image acquisition over conventional time domain OCT. SD-OCT can produce 27,000 to 40,000 A-scans/seconds with an axial resolution ranging from  $3.5\,\mu\mathrm{m}$  to  $6\,\mu\mathrm{m}$  [4].

Many of the previous works on OCT image analysis have focused on the problem of retinal layers segmentation, which is a necessary step for retinal thickness measurements [5, 8]. Few works have addressed the specific problem of DME and its associated features detection from OCT images. Quellec et al. proposed a method for the identification of fluid-filled regions in SD-OCT images of the macula based on texture features extracted in the pre-segmented retinal layers [11].

The authors in [15] proposed a classification method to distinguish DME, Age-related Macular Degeneration (AMD) and normal SD-OCT volumes. The OCT images are pre-processed by reducing the speckle noise by enhancing the sparsity in a transform-domain and flattening the retinal curvature to reduce the inter-patient variations. Then, Histogram of Oriented Gradients (HOG) are extracted for each slice of a volume and a linear Support Vector Machines (SVM) is used for classification. On a dataset of 45 patients equally subdivided into the three aforementioned classes, this method leads to a correct classification rate of 100%, 100% and 86.67% for normal, DME and AMD patients, respectively.

Venhuizen et al. also proposed a method for OCT images classification using the Bag-of-Words (BoW) models [17]. The method starts with the detection and selection of keypoints in each individual B-scan by keeping the most salient points corresponding to the top 3% of the vertical gradient values. Then, a texton of size  $9 \times 9$  pixels is extracted around each keypoint, and Principal Component Analysis (PCA) is applied to reduce the dimension of every texton to get a feature vector of size 9. All extracted feature vectors are used to create a codebook using k-means clustering, and the obtained codebook from the training is used to represent each OCT volume as a feature vector occurrence histogram. Finally, this histogram is used as feature vector to train a Random Forest (RF) with a maximum of 100 trees. The method was used to classify OCT volumes

between AMD and normal cases and achieved an Area Under the Curve (AUC) of 0.984 with a dataset of 384 OCT volumes.

The most similar work to ours is the work of Liu et al. who proposed a method for macular pathology detection in OCT images using Local Binary Patterns (LBP) and gradient information as attributes [9]. The method starts by aligning and flattening the images, then a 3-level multi-scale spatial pyramid is created and edge and LBP histograms are extracted in each block at every level of the pyramid. All obtained histograms are concatenated into a global descriptor whose dimensions are reduced using PCA. Finally a SVM is used as classifier. The method achieved good results in detection OCT scan containing different pathology such as DME or AMD, with an AUC of 0.93 using a dataset of 326 OCT scans.

In this paper, we propose a method for automatic identification of patients with DME versus normal subjects by classifying the OCT volumes. Our method is based on LBP features to describe the texture of OCT images and dictionary learning using the BoW models [14]. However, our method do not rely on keypoints detection as opposed to the work of Venhuizen *et al.* who also employed the BoW models [17]. We rather divide the images into local patches and extract a dense set of LBP descriptors. We also use the entire OCT volume and extract 3D-LBP features to describe the volume, which is different from the work of Liu *et al.* who classified only the foveal scan for each patient [9].

This paper is organized as follows. Section ?? describes the features extraction methodology and the classification approach based on the BoW models. Experiments and results are discussed in Sect. ?? and Sect. ??, respectively. Conclusions and avenue for future directions are drawn in Sect. ??

#### 2 Materials and Methods

The proposed method, as well as, its experimental set-up for OCT volume classification are outlined in Fig. 1. The methodology is formulated as a standard classification procedure. First, the OCT volumes are pre-processed as presented in details in Sect. 2.1. The mapping stage is used to determine a discrete set of elements (or structures) which is used for representing the OCT volume. The feature extraction and representation are presented in depth in Sect. 2.2 and Sect. 2.3. A RF classifier has been selected to perform the classification of the OCT volume [2].



Fig. 1. Machine learning classification basic scheme

#### 2.1 Image pre-processing

OCT images are known to be affected by a speckle noise [12]. Subsequently, Non-Local Means (NL-means) [3] filtering has been previously successfully used in Ultra-Sound (US) images to filter similar noise [6] and is used in our framework to denoise each B-scan (i.e. y axis) of the OCT volumes (see in Fig. 2(a)). NL-means filtering offers the advantage to use all the possible self-predictions that the image can provide rather than local or frequency filters such as Gaussian, anisotropic or Wiener filters [3]. The different parameters were empirically tested and fixed such that the patch size, the search window and the filtering parameter h were set to  $(15 \times 15)$ ,  $(35 \times 35)$  and 0.4, respectively. An example of filtering using NL-means filter on OCT image is depicted in Fig 2(b) and Fig. 2(c).

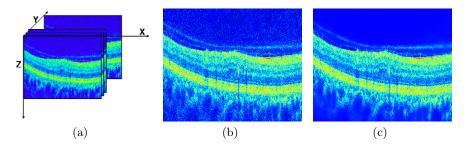


Fig. 2. OCT: (a) Organization of the OCT data - (b) Original image - (c) NL-means filtering.

#### 2.2 Features extraction

LBP is a texture descriptor based on the signs of the differences of a central pixel with respect to its neighboring pixels [10]. These differences are encoded in terms of binary patterns as in Eq. 1:

$$LBP_{P,R} = \sum_{p=0}^{P-1} s(g_p - g_c) 2^p , \qquad s(\cdot) = \begin{cases} 1 & \text{if } (g_p - g_c) \ge 0 \\ 0 & \text{otherwise} \end{cases}.$$

where  $g_c$ ,  $g_p$  are the intensities of the central pixel and a given neighbor pixel, respectively. P is the number of sampling points in the circle of radius R.

Ojala et al. further extend the original LBP formulation to achieve rotation invariance at the expense of limiting the texture description to the notion of circular "uniformity" [10]. Volume encoding is later proposed by Zhao et al. by computing LBP descriptors in each orthogonal planes, so called LBP from Three Orthogonal Planes (LBP-TOP).

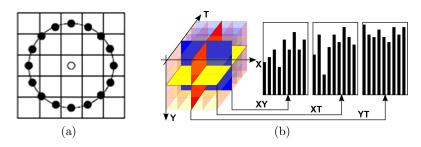


Fig. 3. The different LBP descriptors: (a) LBP - (b) LBP-TOP.

#### 2.3 Feature representation

Texture is defined as the area property related to spatial repetition of structures, statistical properties of the area or both. Two strategies here are employed.

Low-level representation The texture descriptor of an OCT volume is defined as the concatenation of the LBP histogram. Furthermore, the size of this entire feature vector is defined according to the mapping strategy chosen (see Fig. 1).

**High-level representation** According to the chosen mapping strategy, the low-level representation can lead to a high dimensional feature space leading. High-level representation simplifies this high dimensional feature space into a more discriminant lower space. PCA and BoW among other methods, are used for this purpose [14]. Although PCA maps the data according to their variance, BoW models represent the features by creating a visual dictionary, or "codebook", from the set of low-level features. The set of low-level features is clustered using k-means in order to create the codebook with k defining the number of visual words. After creating the codebook, each of the training example is represented as a histogram of size k obtained by calculating the frequency of occurrences of each of the k words in the features extracted from the training example.

Note that in the 2D case, each slice is divided into patches of size  $N \times N$  and we extract 2D-LBP from each patch, while in the 3D case, the volume is divided into  $N \times N \times N$  patches and 3D-LBP histograms are computed. In our experiments in Section 3, we set N=7, and vary the size of the codebook K in the range  $\{2,4,8,16,32,64,100\}$ .

# 3 Experiments and Validation

#### 3.1 Data sets

**SERI** - data sets were acquired by Singapore Eye Research Institute (SERI), using CIRRUS TM (Carl Zeiss Meditec, Inc., Dublin, CA) SD-OCT device. The data sets consist of 32 OCT volumes (16 DME and 16 normal cases). Each

volume contains 128 B-sane with dimension of  $512 \times 1024$  pixels. All SD-OCT images are read and assessed by trained graders and identifies as normal or DME cases based on evaluation of retinal thickening, hard exudates, intraretinal cystoid space formation and subretinal fluid.

**Duke** - data sets published by Srinivasan et al. [15] were acquired in Institutional Review Board-approved protocols using Spectralis SD-OCT (Heidelberg Engineering Inc., Heidelberg, Germany) imaging at Duke University, Harvard University and the University of Michigan. This data sets consist of 45 OCT volumes (15 AMD, 15 DME and 15 normal). In this study we only consider a subset of the original data containing 15 DME and 15 normal OCT volumes.

#### 3.2 Validation

For evaluation purposes, the results have been cross-validated, by splitting the data in training and testing using a Leave-One-Patient Out (LOPO) strategy. In this manner for each round a pair DME, normal has been selected to be used as the round test set, while the rest of the data sets have been used as a training. Doing the cross validation in this manner, has the limitation that despite the fact that the results are robust due to the cross validation, no results variance can be reported. However, and despite this limitation, LOPO has been choose due to the reduced amount of OCT volumes available.

#### 3.3 Experiment

The SERI data sets are provided in complete OCT volumes by  $512\times1024\times128$  dimensions. Using this data sets, first the three low-level features such as LBP, LBP+PCA and LBP-TOP are extracted. The rotation invariant uniform (riu2) descriptors are calculated with the P number of 8, 16 and 24 for the radius if 1, 2 and 3 respectively. The features are classified using RF with 100 tress. Table 1 shows the relative results for 8riu2, 16riu2, 24riu2 and their combination 8riu2 + 16riu2 + 24riu2. The results are presented in terms of Sensitivity (SE) and Specificity (SP) percentages.

The second experiment is carried out using high-level features and BoW approach, on SERI data sets. The first high-level feature LBP+BoW is obtained by applying BoW with 32 visual-words on the previously low-level LBP features (applied on each B-scan). The second and third high-level descriptors are obtained using a dense approach by applying the Sliding Window (SW) of size  $(7\times7)$  on each B-scan and SW of size  $(7\times7\times7)$  to the whole volume respectively. LBP+BoW+SW represent the second high-level feature where the 2D-LBP features are extracted for each sliding window on each B-scan and the visual-words are selected from the pool, consisting of their histograms. The third high-level feature, LBP-TOP+BoW+SW, is defined using LBP-TOP. By using the sliding window the 3D-LBP features are extracted for each patch. Same as previous experiment with low-level features, the descriptors are calculated with the P-

number of 8, 16 and 24 for the radius if 1, 2 and 3 respectively. The obtained results of this experiment are illustrated in Tab. 2.

In order to compare our proposed framework the third experiment is carried out using the subsection of Duke data sets [15]. The OCT volumes provided by this data sets are of different volume size, cropped and denoised by the method of authors choice. Subsequently only the second experiment with high-level features and low-level LBP-TOP features comply with these requirements. The number of visual-words and the size of SW for 2D and 3D features are the same than the previous experiment. The 2D and 3D LBP features are extracted with P number of 8, 16 and 24 for the radius if 1, 2 and 3 respectively. The obtained results for this experiment are shown in Tab. 3.

# 3.4 Results

 $\textbf{Table 1.} \ \textbf{Obtained results with LBP}, \textbf{LBP+PCA} \ \textbf{and LBP-TOP} \ \textbf{features and RF} \ \textbf{with} \ 100 \ \textbf{trees on SERI dataset}$ 

Features	LBP		LBP+PCA		LBP-TOP	
	SE	SP	SE	SP	SE	SP
$ 8^{riu2} $ $ 16^{riu2} $ $ 24^{riu2} $	43.75 37.50 50.00	43.75 50.00 62.50	50.00 68.75 56.25	68.75 56.25 37.50	56.25 <b>87.50</b> 68.75	62.50 <b>75.00</b> 68.75
$\{8, 16, 24\}^{riu2}$	37.50	56.25	68.75	68.75	81.25	$\bf 81.25$

**Table 2.** Obtained results for LBP+BoW, LBP+BoW+SW, and LBP-TOP+BoW+SW features with K=32 and RF with 100 trees using SERI dataset.

Features	LBP+BoW		LBP+	LBP+BoW+SW		LBP-TOP+BoW+SW		
	SE	SP	SE	SP	SE	SP		
$8^{riu2}$ $16^{riu2}$ $24^{riu2}$	57.50	81.25 68.75 50.00	81.25	75.00		68.75 37.50		

# 4 Conclusions

# References

 Abramoff, M.D., Garvin, M.K., Sonka, M.: Retinal image analysis: a review. IEEE Review Biomed. Eng. 3, 169–208 (2010)

**Table 3.** Obtained results for LBP+BoW+SW, and LBP-TOP+BoW+SW, LBP-TOP features on Duke data sets. Number of RF trees is set to 100 and K = 32.

Features	LBP+BoW+SW		LBP-7	ΓΟP+BoW+SW	LBP-TOP	
	SE	SP	SE	SP	SE	SP
$8^{riu2}$	80.00	86.67	80.00	86.67	80.00	93.33
$16^{riu2}$	86.67	100.00	86.67	86.67	73.33	86.67
$24^{riu2}$	93.33	86.67	60.00	80.00	73.33	86.67

**Table 4.** Comparing the proposed method by [17] on SERI and Duke data sets with the **our two** best proposed methods. K = 32 and RF is trained using 100 tress

Data sets	SERI		Duke	
	SE	SP	SE	SP
[17] $\{LBP+BoW+SW\}, 16^{riu2}$ $\{LBP-TOP\}, 16^{riu2}$	81.25	58.82 75.00 75.00	71.42 86.67 73.33	100.00

- 2. Breiman, L.: Random forests. Machine learning 45(1), 5-32 (2001)
- 3. Buades, A., Coll, B., Morel, J.M.: A non-local algorithm for image denoising. In: Computer Vision and Pattern Recognition, 2005. CVPR 2005. IEEE Computer Society Conference on. vol. 2, pp. 60–65. IEEE (2005)
- 4. Chen, T.C., Cense, B., Pierce, M.C., Nassif, N., Park, B.H., Yun, S.H., White, B.R., Bouma, B.E., Tearney, G.J., de Boer, J.F.: Spectral domain optical coherence tomography: ultra-high speed, ultra-high resolution ophtalmic imaging. Arch. Ophtalmol. 123(12), 1715–1720 (2005)
- 5. Chiu, S.J., Li, X.T., Nicholas, P., Toth, C.A., Izatt, J.A., Farsiu, S.: Automatic segmentation of seven retinal layers in sd-oct images congruent with expert manual segmentation. Optic Express 18(18), 19413–19428 (2010)
- Coupe, P., Hellier, P., Kervrann, C., Barillot, C.: Nonlocal means-based speckle filtering for ultrasound images. IEEE Trans Image Process 18(10), 2221–2229 (Oct 2009)
- 7. Early Treatment Diabetic Retinopathy Study Group: Photocoagulation for diabetic macular edema: early treatment diabetic retinopathy study report no 1. Arch. Ophtalmol. 103(12), 1796–1806 (1985)
- 8. Kafieh, R., Rabbani, H., Abramoff, M.D., Sonka, M.: Intra-retinal layer segmentation of 3d optical coherence tomography using coarse grained diffusion map. Medical Image Analysis 17, 907–928 (2013)
- Liu, Y.Y., Chen, M., Ishikawa, H., Wollstein, G., Schuman, J.S., M., R.J.: Automated macular pathology diagnosis in retinal oct images using multi-scale spatial pyramid and local binary patterns in texture and shape encoding. Medical Image Analysis 15, 748–759 (2011)
- Ojala, T., Pietikäinen, M., Mäenpää, T.: Multiresolution gray-scale and rotation invariant texture classification with local binary patterns. Pattern Analysis and Machine Intelligence, IEEE Transactions on 24(7), 971–987 (2002)

- Quellec, G., Lee, K., Dolejsi, M., Garvin, M.K., Abramoff, M.D., Sonka, M.: Threedimensional analysis of retinal layer texture: identification of fluid-filled regions in sd-oct of the macula. IEEE Trans. on Medical Imaging 29, 1321–1330 (2010)
- 12. Schmitt, J.M., Xiang, S., Yung, K.M.: Speckle in optical coherence tomography. Journal of biomedical optics 4(1), 95–105 (1999)
- Sharma, S., Oliver-Hernandez, A., Liu, W., Walt, J.: The impact of diabetic retinopathy on health-related quality of life. Curr. Opin. Ophtalmol. 16, 155–159 (2005)
- Sivic, J., Zisserman, A.: Video google: a text retrieva approach to object matching in videos. In: IEEE ICCV. pp. 1470–1477 (2003)
- Srinivasan, P.P., Kim, L.A., Metttu, P.S., Cousins, S.W., Comer, G.M., Izatt, J.A., Farsiu, S.: Fully automated detection of diabetic macular edema and dry agerelated macular degeneration from optical coherence tomography images. Biomedical Optical Express 5(10), 3568–3577 (2014)
- 16. Trucco, E., Ruggeri, A., Karnowski, T., Giancardo, L., Chaum, E., Hubschman, J., al Diri, B., Cheung, C., Wong, D., Abramoff, M., Lim, G., Kumar, D., Burlina, P., Bressler, N.M., Jelinek, H.F., Meriaudeau, F., Quellec, G., MacGillivray, T., Dhillon, B.: Validation retinal fundus image analysis algorithms: issues and proposal. Investigative Ophthalmology & Visual Science 54(5), 3546–3569 (2013)
- Venhuizen, F.G., van Ginneken, B., Bloemen, B., van Grisven, M.J.P.P., Philipsen, R., C., H., Theelen, T., Sanchez, C.I.: Automated age-related macular degeneration classification in oct using unsupervised feature learning. In: SPIE Medical Imaging. vol. 9414, p. 941411 (2015)
- Wild, S., Roglic, G., Green, A., Sicree, R., King, H.: Global prevalence of diabetes estimates for the year 2000 and projections for 2030. Diabetes Care 27(5), 1047– 1053 (2004)