





Disc-Aware Ensemble Network for Glaucoma Screening From Fundus Image

Huazhu Fu¹⁰, Jun Cheng¹⁰, Yanwu Xu, Changqing Zhang¹⁰, Damon Wing Kee Wong, Jiang Liu, and Xiaochun Cao¹⁰

Abstract—Glaucoma is a chronic eye disease that leads to irreversible vision loss. Most of the existing automatic screening methods first segment the main structure and subsequently calculate the clinical measurement for the detection and screening of glaucoma. However, these measurement-based methods rely heavily on the segmentation accuracy and ignore various visual features. In this paper, we introduce a deep learning technique to gain additional image-relevant information and screen glaucoma from the fundus image directly. Specifically, a novel discaware ensemble network for automatic glaucoma screening is proposed, which integrates the deep hierarchical context of the global fundus image and the local optic disc region. Four deep streams on different levels and modules are, respectively, considered as global image stream, segmentation-guided network, local disc region stream, and disc polar transformation stream. Finally, the output probabilities of different streams are fused as the final screening result. The experiments on two glaucoma data sets (SCES and new SINDI data sets) show that our method outperforms other state-of-the-art algorithms.

Index Terms—Deep learning, glaucoma screening, optic disc segmentation, neural network.

I. INTRODUCTION

C LAUCOMA is one of the major leading causes of blindness among eye diseases, predicted to affect around 80 million people by 2020 [1]. Unlike other eye diseases such

Manuscript received April 14, 2018; accepted May 7, 2018. Date of publication May 15, 2018; date of current version October 29, 2018. This work was supported in part by the National Key Research and Development Plan under Grant 2016YFB0800603, in part by the Ningbo 3315 Innovation Team under Grant Y61102DL03, in part by the National Natural Science Foundation of China under Grant U1605252, Grant 61650202, Grant 61733007, and Grant 61602337, and in part by the Key Program of the Chinese Academy of Sciences under Grant QYZDB-SSW-JSC003. (Corresponding authors: Jun Cheng; Yanwu Xu.)

- H. Fu and D. W. K. Wong are with the Institute for Infocomm Research, Agency for Science, Technology and Research, Singapore 138632 (e-mail: huazhufu@gmail.com; wkwong@i2r.a-star.edu.sg).
- J. Cheng and J. Liu are with the Cixi Institute of Biomedical Engineering, Chinese Academy of Sciences, Ningbo 315201, China (e-mail: sam.j.cheng@gmail.com; jimmyliu@nimte.ac.cn).
- Y. Xu is with CVTE Research, Guangzhou Shiyuan Electronics Co., Ltd., Guangzhou 510530, China (e-mail: xuyanwu@cvte.com).
- C. Zhang is with the School of Computer Science and Technology, Tianjin University, Tianjin 300072, China (e-mail: zhangchangqing@tju.edu.cn).
- X. Cao is with the State Key Laboratory of Information Security, Institute of Information Engineering, Chinese Academy of Sciences, Beijing 100093, China (e-mail: caoxiaochun@iie.ac.cn).

Color versions of one or more of the figures in this paper are available online at http://ieeexplore.ieee.org.

Digital Object Identifier 10.1109/TMI.2018.2837012

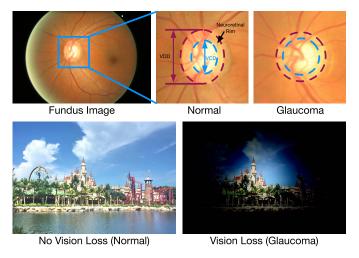


Fig. 1. Top: the whole fundus image and zoom-in normal/glaucoma disc regions, where the vertical cup to disc ratio (CDR) is calculated by the ratio of vertical cup diameter (VCD) to vertical disc diameter (VDD). Bottom: the visual fields with normal and glaucoma cases.

as cataracts and myopia, vision loss from glaucoma cannot be reversed. Early screening is thus essential for early treatment to preserve vision and maintain life quality. However, many glaucoma patients are not aware of their condition [2]. That is why glaucoma is also called the "silent theft of sight", as shown in the bottom row of Fig. 1. Clinically, there are three examinations practiced to screen glaucoma: intraocular pressure (IOP) measurement, function-based visual field test, and optic nerve head (ONH) assessment. IOP is an important risk factor but not specific enough to be an effective detection tool for a great number of glaucoma patients with normal tension. Function-based visual field testing requires specialized perimetric equipment not normally present in primary healthcare clinics. Moreover, the early glaucoma often does not have visual symptoms. ONH assessment is a convenient way to detect glaucoma early, and is currently performed widely by trained glaucoma specialists [3]-[5].

Manual ONH assessment by trained clinicians is time-consuming and costly. Thus, an automatic method is necessity for screening. One popular ONH assessment method is based on the measurement of clinical parameters, such as the vertical cup to disc ratio (CDR) [6], rim to disc area ratio, and disc diameter [7]. Among them, CDR is well accepted and commonly used by clinicians. As shown in the top row of Fig. 1, the CDR is calculated by the ratio of vertical cup

diameter (VCD) to vertical disc diameter (VDD). In general, a larger CDR suggests a higher risk of glaucoma and vice versa. For automatic screening, measurement-based methods have been proposed [8]-[12], which segment the main structure (e.g., optic disc and optic cup) first, and then calculate the clinical measurement values to identify the glaucoma cases. For example, a superpixel-based classifier with various handcrafted visual features is utilized to extract the optic disc and cup regions [10]. The CDR value is then calculated based on the segmented regions. In [13], a CDR assessment using fundus image is proposed, where a sparse dissimilarityconstrained coding approach is employed to consider both the dissimilarity constraint and the sparsity constraint from a set of reference discs with known CDRs. The reconstruction coefficients are used to compute the CDR for the testing disc. Recently, the deep learning based techniques have been also introduced to segment the fundus image. A multi-label deep network is proposed in [12] to segment the optical disc and cup jointly. Then the CDR is accordingly calculated to verify the glaucoma detection. However, these measurement-based methods rely heavily on the segmentation accuracy, which are easily affected by pathological regions and low contrast quality. Moreover, some clinical measurements identify a wider set of image properties, some of which are unrelated to what clinicians seem to recognize as relevant.

In contrast, learning-based methods have been recently proposed to utilize the various visual features from the fundus image and screen the glaucoma disease directly through a learned classifier [14]–[17]. The extracted visual features could explore more image relevant information, and have more representation capacity than clinical measurements. Noronha et al. [16] proposed an automated glaucoma diagnosis method using higher order spectra cumulants extracted from Radon transform applied on digital fundus images. Besides, Acharya et al. [17] proposed a screening method using various features extracted from Gabor transform applied on digital fundus images. Deep learning techniques have been recently demonstrated to yield highly discriminative representations that have aided a considerable number of computer vision applications [18], [19] and medical image analysis areas [20]-[22]. For example, Convolutional Neural Networks (CNNs) learn hierarchical features from raw image data and are demonstrated to achieve a high performance on retinal vessel detection [23], [24]. For glaucoma detection, works in [25] and [26] applied deep learning systems to detect glaucoma from fundus images directly. However, these methods merely focus on optic disc region limited contextual information contained in the patch, and thus ignore the global information from the whole fundus image. The accuracy of optic disc region cropping disturbs the detection performance.

In this paper, we propose a glaucoma screening network for the fundus image with the following aspects:

1) **Disc-aware:** Glaucoma is an eye disease that is often associated with elevated intraocular pressure, which damages the optic nerve (disc). Thus the main clinical symptoms appear in the optic disc region [27], [28].

- Based on this, the proposed network should highlight the contextual information in the optic disc region.
- 2) **Multi-level:** Most of existing deep learning networks focus on the global image directly, and employ pooling the layer to achieve hierarchical representations [18], [20]. A common limitation of deep learning based approaches is the need to downsample the input image to a low resolution (i.e., 224 × 224) in order for the network size to be computationally manageable. However, this downsampling leads to loss of image details that are important for discrimination of subtle pathological changes. In comparison, the local region scale preserves a fine representation, but ignores the global structure information. Thus incorporating both local and global contexts could effectively improve the performance.
- 3) Multi-module: In clinical trials, fine measurements are often used to evaluate disease indicator. The geometric transformation could alter the morphology of the pathological region and enhance the detection changing [29]. A good example is the optic disc and cup segmentation [12], in which polar transformation is employed to balance the region proportion and improve the optic cup segmentation performance. In our work, what we consider is whether the geometry transformation could improve the screening performance.

To address these, we propose a novel Disc-aware Ensemble Network (DENet) for automatic glaucoma screening, which contains four deep streams corresponding to various levels and modules of the fundus image, as shown in Fig. 2. The first one is a global image stream, which represents the global fundus structure on image level, and works as a classification on the fundus image directly. The second one is a segmentationguided network, which detects the disc localization from the whole fundus image and embeds the disc segmentation representation to detect glaucoma at the image level. The third stream is based on the local disc region, which produces the screening probability from the disc region level. The final stream focuses on disc region with polar transformation, which enlarges the disc and cup structure with the geometry operation and improves the screening performance. Finally, all the outputs of these deep streams are combined to obtain the final screening result. Our proposed DENet is an automatic system for the whole fundus image that includes disc detection and glaucoma screening, and uses the hierarchical representations on different levels (global image and disc region) and modules (Cartesian and Polar coordinates). For evaluation, we test our method on two clinical datasets, the SCES dataset and a new collected SINDI dataset. The experiments show that our method outperforms state-of-the-art methods on glaucoma screening. The main contributions of this paper are as follows:

 An ensemble deep network is proposed for glaucoma screening by considering various levels and modules of the fundus image. Our method achieves promising results as it can make further improvement based on the existing networks. Another advantage of our ensemble method is that it can be flexibly embedded within various

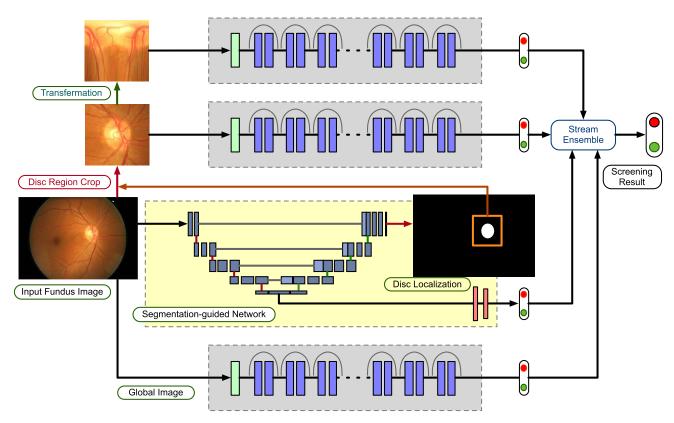


Fig. 2. Architecture of our DENet, which contains four streams: global image stream produces the result based on the global fundus image; the segmentation-guided network localizes the optic disc region and generates a detection output embedded the disc-segmentation representation; disc region stream works on disc region cropped by disc segmentation map from segmentation-guided network; disc polar stream transfers the disc region image into the polar coordinate system. The combination of these four streams is fused as the final glaucoma screening result.

networks, which makes it easier to adapt to different scenarios.

- 2) Beside the ensemble architecture, we also provide a novel Segmentation-guided Network, which localizes the disc region and generates the screening result by embedding the disc-segmentation representation.
- 3) For evaluation of glaucoma screening, we construct a new dataset, called the Singapore Indian Eye Study (SINDI) dataset, which is a population-based study conducted and contains a total of 5783 eye images including 113 glaucomatous eyes and 5670 normal eyes.

II. DISC-AWARE ENSEMBLE NETWORK

Our Disc-aware Ensemble Network (DENet) takes into account two levels of fundus image information, that is the global image and the local disc region, as shown in Fig. 2. The global image level provides the coarse structure representation on the whole fundus image, while the local disc region is utilized to learn a fine representation around the optic disc.

A. Global Fundus Image Level

In our DENet, two streams are employed to learn representations on the global fundus image level. The first stream is a standard classification network by using Residual Network (ResNet) [30]. The ResNet is based on a Convolutional Neural Network and introduces the shortcut connection to handle the vanishing gradient problem in very deep networks,

as shown in Fig. 3 (b). We utilize a ResNet-50 as the backbone model to learn the global representation on the whole fundus image directly, which consists of 5 down-sampling blocks, followed by a global max pooling layer and a fully connected (FC) layer for glaucoma screening. The input image of this stream is resized to 224×224 to enable use of pre-trained model in [30] as initialization for our network.

The second global level stream is the segmentation-guided network, which localizes the optic disc region and produces a detection result based on the disc-segmentation representation. As shown in Fig. 3 (a), the main architecture of the segmentation-guided network is adapted by the U-shape convolutional network (U-Net) in [31], which is an efficient fully convolutional neural network for biomedical image segmentation. Similar to the original U-Net architecture, our method consists of the encoder path (left side) and decoder path (right side). Each encoder path performs convolutional layer with a filter bank to produce a set of encoder feature maps, and the element-wise rectified-linear non-linearity (ReLU) activation function is utilized. The decoder path also utilizes the convolutional layer to output the decoder feature map. The skip connections transfer the corresponding feature map from encoder path and concatenate them to up-sampled decoder feature maps. Finally, a classifier utilizes 1×1 convolutional layer with Sigmoid activation as the pixel-wise classification to produce the disc probability map. Moreover, we extend a new branch from the saddle layer of the U-shape network, where the size scale is the smallest (i.e., 40×40) and the

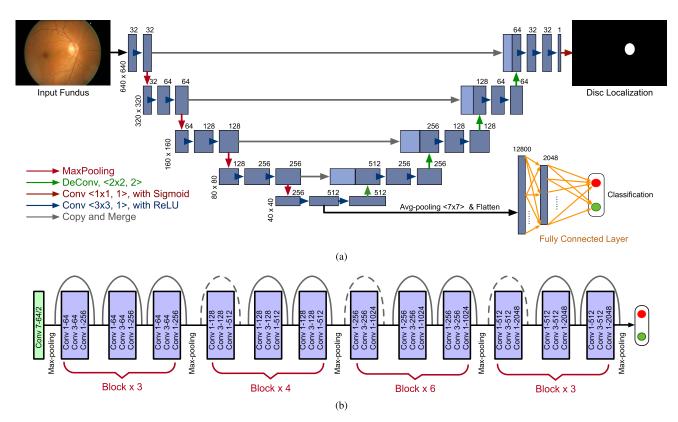


Fig. 3. (a) The detailed architecture of our segmentation-guided network, which is based on U-shape convolutional network to detect optic disc region and guide glaucoma screening on the whole fundus image. (b) Architecture of ResNet [30] with 50 layers. Each convolutional unit (denoted by a blue block) consists of layers for convolution, batch normalization, and ReLU activation.

number of channel is the highest (i.e., 512-D). The extended branch acts as an implicit vector with average pooling and flatten layers. Next it connects two fully connected layers to produce a glaucoma classification probability. This pipeline embeds the segmentation-guided representation through the convolutional filters on decoder path of the U-shape network. The input image of this stream is resized to 640×640 , which guarantees that the image has enough details to localize disc region accurately.

In our global fundus image level networks, two loss functions are employed. The first one is the binary cross entropy loss function for glaucoma detection layer. The other is the Dice coefficient for assessing disc segmentation [32], which is defined as:

$$L_{Dice} = 1 - \frac{2\sum_{i}^{N} p_{i}g_{i}}{\sum_{i}^{N} p_{i}^{2} + \sum_{i}^{N} g_{i}^{2}},$$
(1)

where N is the pixel number, $p_i \in [0, 1]$ and $g_i \in \{0, 1\}$ denote predicted probability and binary ground truth label for disc region, respectively. The Dice coefficient loss function can be differentiated yielding the gradient as:

$$\frac{\partial L_{Dice}}{\partial p_i} = \frac{4p_i \sum_{i}^{N} p_i g_i - 2g_i (\sum_{i}^{N} p_i^2 + \sum_{i}^{N} g_i^2)}{(\sum_{i}^{N} p_i^2 + \sum_{i}^{N} g_i^2)^2}.$$
 (2)

These two losses are efficiently integrated into back-propagation via standard stochastic gradient descent (SGD).

Note that we use two phases for training the segmentationguided model. Firstly, the U-shape segmentation network for disc detection is trained by pixel-level disc training data with Dice coefficient loss. Then the parameters of CNN layers are frozen and the fully connected layers for the classification task are trained by using glaucoma detection training data. We use the separate phases to train our segmentation-guided model instead of the multi-task based single stage training, with the following reasons: 1) Using the disc-segmentation representation on screening could add diversity of the proposed network. 2) The pixel-level label data for disc segmentation is more expensive than image-level label data for glaucoma detection. The separate stage could employ different training datasets and configuration (e.g, different batch sizes and image numbers). 3) The extracted disc region by network influences the follow-up stream, thus the accuracy of disc detection is more important than classification branch.

B. Optic Disc Region Level

The second level in our network is based on the local optic disc region, which is cropped based on the previous segmentation-guided network. The local disc region preserves more detailed information with higher resolution and it is benefited to learn a fine representation. Two local streams are used in our network to learn representations on the local disc region. The first one is a standard classification network based on ResNet [30] on the original local disc region, as shown in Fig.2, while the other stream focuses on the disc polar transformation.

In our method, we apply the pixel-wise polar transformation to transfer the original image to the polar coordinate system.

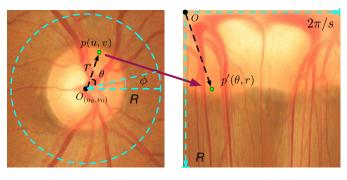


Fig. 4. Illustration of the mapping from Cartesian coordinate system (Left) to the polar coordinate system (Right) by using the polar transformation. The point p(u, v) in Cartesian coordinate corresponds to the point $p'(\theta, r)$ in polar coordinate.

Let p(u, v) denote the point on the original Cartesian plane, and its corresponding point on polar coordinate system is denoted $p'(\theta, r)$, as shown in Fig. 4, where r and θ are the radius and directional angle of the original point p, respectively. Three parameters are utilized to control the polar transformation: the disc center $O(u_o, v_o)$, the polar radius R, and the polar angle ϕ . The polar transform mapping is calculated as follows:

$$\begin{cases} u = u_o + r\cos(\theta + \phi), \\ v = v_o + r\sin(\theta + \phi), \end{cases}$$
 (3)

and the inverse polar transform mapping is as:

$$\begin{cases} r = \sqrt{(u - u_o)^2 + (v - v_o)^2}, \\ \theta = \tan^{-1}(\frac{v - v_o}{u - u_o}) - \phi. \end{cases}$$

The height and width of polar image are set as the polar radius R and discretization $2\pi/s$, where s is the stride. The disc polar transformation has the following advantages:

- Since the disc and cup are structured as near concentric circles shape. The polar transformation could enlarge the cup region by interpolation. The increased cup region displays more details. As shown in Fig. 4, the proportion of cup region is increased and more balanced than that in original fundus image.
- 2) Due to the pixel-wise mapping, the polar transformation is equivariant to the data augmentation on the original fundus image [33]. For example, moving the transformation center $O(u_o, v_o)$ corresponds to the drift translation in polar coordinates. Using different polar radius R is the same as augmenting with various scaling factor. And changing the polar angle ϕ will shift in horizontal coordinate in the image. Thus the data augmentation of deep learning can be done by using the polar transformation with various parameters.

The input disc region of these two local region streams are resized to 224×224 to enable use of pre-trained parameters from ResNet model [30] as initialization. And the binary cross entropy loss function is used for glaucoma detection in the local disc region.

TABLE I
THE DATASETS USED IN OUR EXPERIMENTS

Method	ORIGA dataset	SCES dataset	SINDI dataset
Purpose	Training	Testing	Testing
Ground Truth	Disc + Glaucoma	Glaucoma	Glaucoma
Image Num.	650	1676	5783
Open / Close	482 / 168	1630 / 46	5670 / 113

III. EXPERIMENT

A. Dataset and Evaluation Criteria

In our experiments, we use three glaucoma screening datasets, as listed in Table I. The first one is the ORIGA dataset, which contains 650 fundus images from different eyes, including 168 glaucomatous eyes and 482 normal eyes with manual ground truth of optic disc boundary. The second one is the Singapore Chinese Eye Study (SCES) dataset, which consists of 1676 images with 46 glaucoma cases. Moreover, a new dataset, called the Singapore Indian Eye Study (SINDI) dataset, is also used. The SINDI dataset is a populationbased study conducted also by the Singapore Eye Research Institute, aiming to assess the risk factors of visual impairment in the Singapore Indian community. In this dataset, there is a total of 5783 eye images including 113 glaucomatous eyes and 5670 normal eyes. Only the ORIGA dataset has the manually labelled ground truth of optic disc boundary, thus we use all the 650 images in ORIGA dataset for network training including disc segmentation and glaucoma screening, and employ the SCES and SINDI datasets for testing. The resolution of the fundus image in these three datasets is 3072×2048 , and the size of cropped disc region is 800×800 .

For evaluation, we report the Receiver Operating Characteristic (ROC) curves and the area under ROC curve (AUC). Moreover, we also employ three evaluation criteria to measure performance, including: Sensitivity (Sen), Specificity (Spe), and Balanced Accuracy (BAcc), which are defined as:

$$\operatorname{Sen} = \frac{TP}{TP + FN}$$
, $\operatorname{Spe} = \frac{TN}{TN + FP}$, $\operatorname{BAcc} = \frac{\operatorname{Sen} + \operatorname{Spe}}{2}$,

where TP and TN denote the number of true positives and true negatives, respectively, and FP and FN denote the number of false positives and false negatives, respectively. Tuning the diagnostic thresholds could obtain a series of criteria scores, and then we use the threshold with highest B-Accuracy score as the final threshold to report the performance.

B. Implementation Details

Our whole framework is implemented with Python based on Keras with Tensorflow backend. The four streams in our system are trained separately according to different data augmentation strategies, which dues to three reasons: 1) The disc local streams are based on the disc detection result of global image stream. 2) The separate training stage is convenient to add new stream into the ensemble network. 3) The separate training stage could employ different training datasets and configuration for different stream. For the global image stream and segmentation-guided network, the data augmentation is done on the training set by random rotations

TABLE II
PERFORMANCE COMPARISONS OF THE DIFFERENT METHODS ON DATASETS

	SCES Dataset			SINDI Dataset				
Method	AUC	B-Accuracy	Sensitivity	Specificity	AUC	B-Accuracy	Sensitivity	Specificity
Airpuff IOP	0.6600	0.6452	0.3696	0.9209	0.6233	0.5991	0.3451	0.8531
Wavelet [15]	0.6591	0.6544	0.7174	0.5914	0.6487	0.6262	0.6283	0.6242
Gabor [17]	0.7594	0.7191	0.9130	0.5252	0.7417	0.7117	0.8053	0.6182
GRI [14]	0.8051	0.7629	0.6957	0.8301	0.7892	0.7121	0.5752	0.8490
Superpixel [10]	0.8265	0.7800	0.7391	0.8209	0.7712	0.7360	0.7257	0.7464
DeepCDR [12]	0.8998	0.8157	0.7609	0.8706	0.7929	0.7585	0.7522	0.7647
Image	0.8258	0.8011	0.7826	0.8196	0.7806	0.7170	0.7611	0.6728
Seg	0.8386	0.7659	0.7391	0.7926	0.7250	0.6656	0.6018	0.7295
Disc	0.8592	0.8138	0.8913	0.7362	0.7560	0.7011	0.6018	0.8004
Polar	0.8748	0.8294	0.8913	0.7675	0.7663	0.7205	0.6726	0.7684
Image + Disc	0.8709	0.8228	0.8696	0.7761	0.8005	0.7481	0.7345	0.7617
Image + Seg	0.8832	0.8089	0.8478	0.7699	0.7952	0.7353	0.8319	0.6388
Image + Polar	0.8996	0.8331	0.8913	0.7748	0.7978	0.7437	0.7965	0.6910
Disc + Polar	0.8876	0.8285	0.8913	0.7656	0.7736	0.7204	0.6195	0.8213
Disc + Seg	0.8805	0.8037	0.7609	0.8466	0.7815	0.7173	0.8053	0.6293
Seg + Polar	0.8894	0.8370	0.9348	0.7393	0.7660	0.7121	0.8673	0.5570
Image + Disc + Seg	0.8965	0.8335	0.8043	0.8626	0.8137	0.7559	0.7788	0.7330
Image + Disc + Polar	0.9019	0.8479	0.9130	0.7828	0.8060	0.7470	0.6991	0.7949
Disc + Seg + Polar	0.9066	0.8520	0.9783	0.7258	0.7912	0.7316	0.6726	0.7907
Image + Seg + Polar	0.9146	0.8348	0.8696	0.8000	0.8093	0.7564	0.7788	0.7340
Our DENet	0.9183	0.8429	0.8478	0.8380	0.8173	0.7495	0.7876	0.7115

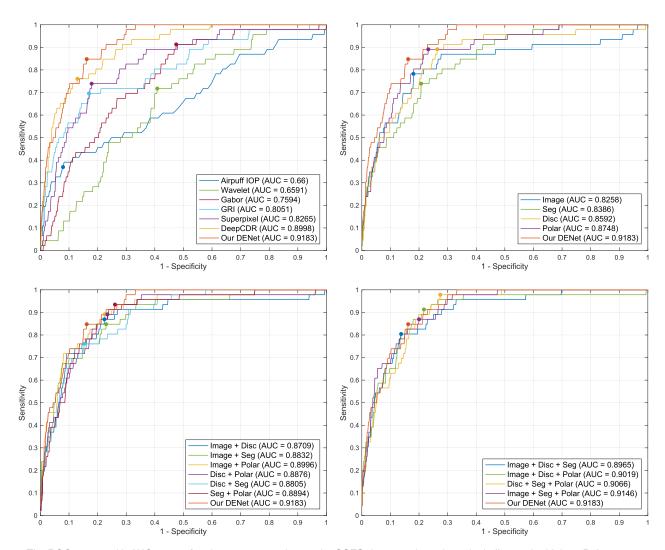


Fig. 5. The ROC curves with AUC scores for glaucoma screening on the SCES dataset, where the point indicates the highest B-Accuracy score.

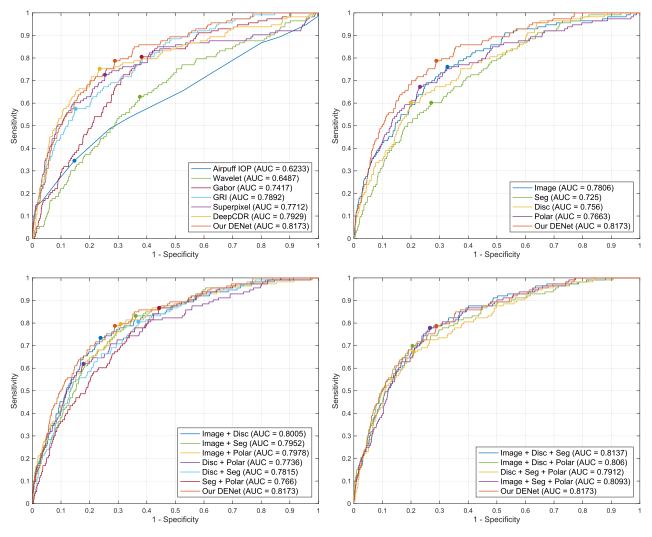


Fig. 6. The ROC curves with AUC scores for glaucoma screening on the SINDI dataset, where the point indicates the highest B-Accuracy score.

(0/90/180/270 degrees) and random flips. For the local disc region scream, the data augmentation is done by random rotations (0/90/180/270 degrees), crop drift $(\pm 20 \text{ pixels})$ and random flips. For the disc polar region, we tune the polar transformation parameters to control the data augmentation, by polar angle $(\phi = 0/90/180/270 \text{ degrees})$, polar center drift $(C_{(u_o,v_o)} \pm 20)$, and polar radius $(R = 400 \times \{0.8,1\})$. During training, we employ stochastic gradient descent (SGD) for optimizing the deep model. We use a gradually decreasing learning rate starting from 0.0001 and a momentum of 0.9. Our ResNet-50 stream employs pre-trained parameters based on ImageNet as initialization, and all the layers are fine-tuned during the training.

C. Glaucoma Screening

We compare our method with five state-of-the-art glaucoma screening baselines: wavelet-based feature method (Wavelet) in [15], Gabor transformation method (Gabor) in [17], Glaucoma risk index method (GRI) in [14], superpixel-based classification method (Superpixel) in [10] and Deep Segmentation method (DeepCDR) in [12]. We additional provide the intraocular pressure (IOP) measurement result as the clinical

baseline. Moreover, for our algorithm, we show not only the final screening result (Our DENet), but also provide results obtained by each stream, such as the global image stream (Image), segmentation-guided network (Seg), local disc region stream (Disc), and local disc polar stream (Polar). The experiment results are reported in Table II and Figs. 5 and 6.

From the glaucoma screening results, the IOP performs poorly with 0.66 AUC on the SECS dataset and 0.6233 AUC on the SINDI dataset. The wavelet-based feature method in [15] utilizes the energy property of the wavelet transformed image, which does not provide enough discriminative capability for glaucoma screening. By contrast the SRI in [14] combines multiple image features (e.g., intensity value, FFT coefficient, and B-spline coefficient), and achieves the higher scores than Wavelet in [15]. The non-deep learning method, Superpixel [10], produces a competitive performance on the SCES dataset (0.8265 AUC) and the SINDI dataset (0.7712 AUC), which is better than IOP. The deep learning method, DeepCDR [12], which jointly segments the optic disc and cup, achieves the higher performances on both datasets. Our DENet obtains the best performances on the SCES dataset (0.9183 AUC) and the SINDI dataset (0.8173 AUC), which has

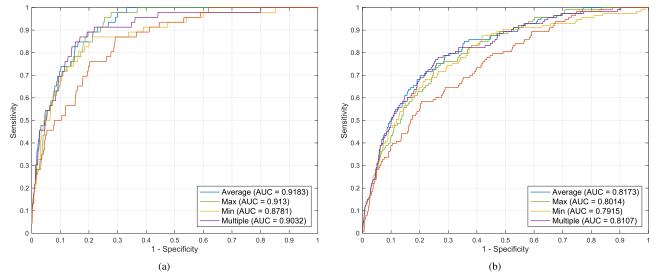


Fig. 7. The performances of the various ensemble operations on the SCES and SINDI datasets. (a) SCES dataset. (b) SINDI dataset.

about 2% improvement on AUC than DeepCDR [12]. It can be seen that without extracting clinical parameters (e.g., CDR), the visual features could be used for glaucoma screening. One possible reason is that the clinical parameters are based on what information clinicians currently observe, while visual features deeply represent a wider set of image properties, some of which may be unrelated with what clinicians defined explicitly. Hence, visual features gain more latent image representations, and more discriminative information than clinical parameters. Moreover, our DENet also outperforms other deep learning based methods. For example, the deep learning method in [25] provides a glaucoma screening system by using CNN feature on the disc region directly, which obtained 0.898 AUC on the SCES dataset. Our method is comparable to that of the deep system [25], and is also able to localize the disc region from the whole fundus image.

D. Discussion

1) Stream Analysis: For each stream of our DENet, the performance on global image level is lower than that on local disc region. It is reasonable as most clinical symptoms are from region close to the optic disc [3], [27], [28]. The ResNet-50 has more CNN layers than our segmentation-guided classification network (50 CNN layers vs 13 CNN layers), which leads the global image stream to get a better performance than segmentation-guided network on the SINDI dataset, and a similar score on the SCES dataset. Another possible reason is that the ResNet loads ImageNet pre-train model as the initialized parameter on training step, while the segmentationguided network only trains on ORIGA dataset with 650 images without any pre-train model. For local disc region, the polar transformation promotes the representation capability of deep feature, and outperforms original disc region on both datasets. Moreover, we also report the different combinations of our streams on the SCES and SINDI datasets. It can be observed that the combinations with three streams have the higher performances than those with two streams. And the combination of all four streams obtains the best scores on both datasets.

TABLE III
THE PERFORMANCES OF DIFFERENT METHODS AT 0.95 SENSITIVITY

	SCES I	Dataset	SINDI Dataset		
Method	B-Accuracy	Specificity	B-Accuracy	Specificity	
Airpuff IOP	0.4932	0.0515	0.4953	0.0437	
Wavelet [15]	0.6065	0.2564	0.5362	0.1166	
Gabor [17]	0.6415	0.3264	0.6002	0.2446	
GRI [14]	0.6148	0.2730	0.6515	0.3473	
Superpixel [10]	0.6681	0.3816	0.5018	0.0478	
DeepCDR [12]	0.7700	0.5834	0.5430	0.1303	
Our DENet	0.8316	0.7067	0.6655	0.3753	

2) Ensemble Analysis: In this experiment, we test different ensemble methods, including average, max, min, and multiple operations. The performances for different ensemble methods on SCES and SINDI datasets are shown in Fig. 7. Most ensemble operations (e.g., average, max and multiple) result in significant improvements, even the min operation is better than the single stream. This means that the ensemble provides an effective way to improve the screening performance. Moreover, the average operation performs best on SCES dataset. Thus, our DENet uses averaging to ensemble the outputs of four streams. Note that we do not use the weighted sum to ensemble the streams, as the average operation is a simple and efficient way without any prior information.

3) Running Time: The training phase of each stream takes about 5 hours on a single NVIDIA Titan X GPU (200 iterations). Since we train each stream individually, they could be done parallelly and offline. In testing, our DENet costs only 0.5 seconds to generate the final result for one fundus image including disc localization and glaucoma screening, which is faster than the existing measurement-based methods, e.g., superpixel method [10] takes 10 seconds, DeepCDR [12] takes 1 second.

4) High-Sensitivity Performance: In real-world, the glaucoma screening system is required to achieve a high sensitivity (not to miss glaucoma case). Thus it would be useful to compare specificities at a certain high sensitivity setting. Table III shows the performances of the different methods at 0.95 Sensitivity,

where our DENet achieves the highest performances on both datasets.

IV. CONCLUSION

In this paper, we have proposed a novel Disc-aware Ensemble Network (DENet) for automatic glaucoma screening, which integrates four deep streams on different levels and modules. The multiple levels and modules are beneficial to incorporate the hierarchical representations, while the disc-aware constraint guarantees contextual information from the optic disc region for glaucoma screening. The experiments on two glaucoma datasets (SCES and new collected SINDI datasets) show that our method outperforms state-of-the-art glaucoma screening algorithms. The work implementation details are available at http://hzfu.github.io/proj_glaucoma_fundus.html.

REFERENCES

- [1] Y.-C. Tham, X. Li, T. Y. Wong, H. A. Quigley, T. Aung, and C.-Y. Cheng, "Global prevalence of glaucoma and projections of glaucoma burden through 2040: A systematic review and meta-analysis," *Ophthalmology*, vol. 121, no. 11, pp. 2081–2090, 2014.
- [2] S. Y. Shen et al., "The prevalence and types of glaucoma in malay people: The Singapore malay eye study," *Invest. Ophthalmol. Vis. Sci.*, vol. 49, no. 9, p. 3846, 2008.
- [3] J. Jonas, W. Budde, and S. Panda-Jonas, "Ophthalmoscopic evaluation of the optic nerve head," *Surv. Ophthalmol.*, vol. 43, no. 4, pp. 293–320, 1999.
- [4] J. E. Morgan, N. J. L. Sheen, R. V. North, Y. Choong, and E. Ansari, "Digital imaging of the optic nerve head: Monoscopic and stereoscopic analysis," *Brit. J. Ophthalmol.*, vol. 89, no. 7, pp. 879–884, 2005.
- [5] H. Fu et al., "Segmentation and quantification for angle-closure glaucoma assessment in anterior segment OCT," IEEE Trans. Med. Imag., vol. 36, no. 9, pp. 1930–1938, Sep. 2017.
- [6] J. B. Jonas, A. Bergua, P. Schmitz–Valckenberg, K. I. Papastathopoulos, and W. M. Budde, "Ranking of optic disc variables for detection of glaucomatous optic nerve damage," *Invest. Ophthalmol. Vis. Sci.*, vol. 41, no. 7, pp. 1764–1773, 2000.
- [7] M. D. Hancox, "Optic disc size, an important consideration in the glaucoma evaluation," *Clin. Eye Vis. Care*, vol. 11, no. 2, pp. 59–62, 1999.
- [8] G. D. Joshi, J. Sivaswamy, and S. R. Krishnadas, "Optic disk and cup segmentation from monocular colour retinal images for glaucoma assessment," *IEEE Trans. Med. Imag.*, vol. 30, no. 6, pp. 1192–1205, Jun. 2011.
- [9] F. Yin et al., "Model-based optic nerve head segmentation on retinal fundus images," in Proc. EMBC, 2011, pp. 2626–2629.
- [10] J. Cheng et al., "Superpixel classification based optic disc and optic cup segmentation for glaucoma screening," *IEEE Trans. Med. Imag.*, vol. 32, no. 6, pp. 1019–1032, Jun. 2013.
- [11] J. Cheng, D. Tao, D. W. K. Wong, and J. Liu, "Quadratic divergence regularized SVM for optic disc segmentation," *Biomed. Opt. Exp.*, vol. 8, no. 5, pp. 2687–2696, 2017.
- [12] H. Fu, J. Cheng, Y. Xu, D. Wong, J. Liu, and X. Cao, "Joint optic disc and cup segmentation based on multi-label deep network and polar transformation," *IEEE Trans. Med. Imag.*, to be published, doi: 10.1109/TMI.2018.2791488.

- [13] J. Cheng, F. Yin, D. W. K. Wong, D. Tao, and J. Liu, "Sparse dissimilarity-constrained coding for glaucoma screening," *IEEE Trans. Biomed. Eng.*, vol. 62, no. 5, pp. 1395–1403, May 2015.
- [14] R. Bock, J. Meier, L. G. Nyul, J. Hornegger, and G. Michelson, "Glaucoma risk index: Automated glaucoma detection from color fundus images," *Med. Image Anal.*, vol. 14, no. 3, pp. 471–481, 2010.
- [15] S. Dua, U. R. Acharya, P. Chowriappa, and S. V. Sree, "Wavelet-based energy features for glaucomatous image classification," *IEEE Trans. Inf. Technol. Biomed.*, vol. 16, no. 1, pp. 80–87, Jan. 2012.
- [16] K. P. Noronha, U. R. Acharya, K. P. Nayak, R. J. Martis, and S. V. Bhandary, "Automated classification of glaucoma stages using higher order cumulant features," *Biomed. Signal Process. Control*, vol. 10, no. 1, pp. 174–183, 2014.
- [17] U. R. Acharya et al., "Decision support system for the glaucoma using Gabor transformation," Biomed. Signal Process. Control, vol. 15, pp. 18–26, Jan. 2015.
- [18] A. Krizhevsky, I. Sutskever, and G. Hinton, "Imagenet classification with deep convolutional neural networks," in *Proc. NIPS*, 2012, pp. 1097–1105.
- [19] Y. LeCun, Y. Bengio, and G. Hinton, "Deep learning," *Nature*, vol. 521, pp. 436–444, May 2015.
- [20] V. Gulshan et al., "Development and validation of a deep learning algorithm for detection of diabetic retinopathy in retinal fundus photographs," J. Amer. Med. Assoc., vol. 304, no. 6, pp. 649–656, 2016.
- [21] A. Esteva et al., "Dermatologist-level classification of skin cancer with deep neural networks," Nature, vol. 542, no. 7639, pp. 115–118, 2017.
- [22] D. S. W. Ting and et al., "Development and validation of a deep learning system for diabetic retinopathy and related eye diseases using retinal images from multiethnic populations with diabetes," JAMA, vol. 318, no. 22, pp. 2211–2223, 2017.
- [23] Q. Li, B. Feng, L. Xie, P. Liang, H. Zhang, and T. Wang, "A cross-modality learning approach for vessel segmentation in retinal images," *IEEE Trans. Med. Imag.*, vol. 35, no. 1, pp. 109–118, Jan. 2016.
- [24] H. Fu, Y. Xu, S. Lin, D. W. K. Wong, and J. Liu, "DeepVessel: Retinal vessel segmentation via deep learning and conditional random field," in *Proc. MICCAI*, 2016, pp. 132–139.
- [25] X. Chen, Y. Xu, S. Yan, D. Wing, T. Wong, and J. Liu, "Automatic feature learning for glaucoma detection based on deep learning," in *Proc. MICCAI*, 2015, pp. 669–677.
- [26] J. I. Orlando, E. Prokofyeva, M. D. Fresno, and M. B. Blaschko, "Convolutional neural network transfer for automated glaucoma identification," *Proc. SPIE*, vol. 10160, p. 101600, Jan. 2016.
- [27] D. F. Garway-Heath and R. A. Hitchings, "Quantitative evaluation of the optic nerve head in early glaucoma," *Brit. J. Ophthalmol.*, vol. 82, no. 4, pp. 352–361, 1998.
- [28] M. C. Leske, S.-Y. Wu, A. Hennis, R. Honkanen, and B. Nemesure, "Risk factors for incident open-angle glaucoma: The barbados eye studies," *Ophthalmology*, vol. 115, no. 1, pp. 85–93, 2008.
- [29] H. Salehinejad, S. Valaee, T. Dowdell, and J. Barfett, "Image augmentation using radial transform for training deep neural networks," in *Proc. ICASSP*, 2018, pp. 1–5.
- [30] K. He, X. Zhang, S. Ren, and J. Sun, "Deep residual learning for image recognition," in *Proc. CVPR*, 2016, pp. 770–778.
- [31] O. Ronneberger, P. Fischer, and T. Brox, "U-Net: Convolutional networks for biomedical image segmentation," in *Proc. MICCAI*, 2015, pp. 234–241.
- [32] W. R. Crum, O. Camara, and D. L. G. Hill, "Generalized overlap measures for evaluation and validation in medical image analysis," *IEEE Trans. Med. Imag.*, vol. 25, no. 11, pp. 1451–1461, Nov. 2006.
- [33] C. Esteves, C. Allen-Blanchette, X. Zhou, and K. Daniilidis, "Polar transformer networks," in *Proc. ICLR*, 2018, pp. 1–14.