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## Research Article

# **Identification of Diabetic Retinopathy through Machine Learning**

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A cross-sectional study of patients with suspected diabetic retinopathy (DR) who had an ophthalmological examination and a retinal scan is the focus of this research. Specialized retinal images were analyzed and classified using OPF and RBM models (restricted Boltzmann machines). Classification of retinographs was based on the presence or absence of disease-related retinopathy (DR). The RBM and OPF models extracted 500 and 1000 characteristics from the images for disease classification after the system training phase for the recognition of retinopathy and normality patterns. There were a total of fifteen different experiment series, each with a repetition rate of 30 cycles. The study included 73 diabetics (a total of 122 eyes), with 50.7% of them being men and 49.3% being women. The population was on the older side, at 59.7 years old on average. The RBM-1000 had the highest overall diagnostic accuracy (89.47) of any of the devices evaluated. The RBM-500 had a better autodetection system for DR signals in fundus images than the competition (100% sensitivity). In terms of specificity, RBM-1000 and OPF-1000 correctly identified all of the images that lacked DR signs. In particular, the RBM model of machine learning automatic disease detection performed well in terms of diagnostic accuracy, sensitivity, and application in diabetic retinopathy screening.

## 1. Introduction

Diabetes mellitus (DM) comprises a group of metabolic disorders that share the phenotype of hyperglycemia, caused by an absolute or relative deficit in insulin production or action. Chronic hyperglycemia is associated with damage, dysfunction, or failure of organs and tissues, including retina, kidney, nervous system, heart, and blood vessels [1]. Diabetic retinopathy is present in both type 1 and type 2 diabetes, especially in patients with long-term disease and poor glycemic control [2]. The pathogenesis is multifactorial, being related to the metabolic effects of chronic hyperglycemia. Among the main factors involved in the genesis of diabetic retinopathy are the biochemical (earlier), hemodynamic, and endocrine factors, which interact with each other, leading to nonenzymatic protein glycation, oxidative stress, accumulation of polyols, and activation of protein kinase C [3]. The need for the early detection of DR has encouraged the search for more effective alternative screening methods. Automatic retinal image analysis models

are options that aim at reducing physicians' workload and providing a practical and cost-effective method [4]. The automatic image analysis software, in general, uses artificial intelligence based on machine learning (machine learning), and some more advanced ones, on deep learning (deep learning), a recent advance in artificial neural networks, which allows to improve the classification of images gross [5].

Given the current scenario of research on alternative screening methods for diabetic retinopathy, we proposed to carry out this study, considering that the automatic models of image analysis, optimum-path forest (OPF) and restricted Boltzmann machines (RBM), can be used together, combining the practicality and low operational cost of OPF with the reduction of the computational load that the RBM allows, in situations that require many input variables for image recognition, as in diabetic retinopathy. Studies using OPF and RBM to identify diabetic retinopathy are rare, and the personal and social burden of blindness caused by diabetic retinopathy encourages the search for new forms of

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screening that have similar accuracy to the gold standard, which is the analysis of photograph by the specialist.

## 2. Material and Methods

- 2.1. Type of Study. This study is a transverse study.
- 2.2. Study Location. The study was conducted at the Department of Ophthalmology, Otorhinolaryngology and Head and Neck Surgery, University of Baghdad, Baghdad, Iraq.
- 2.3. Ethical Analysis. The research project was approved by the Research Ethics Committee of the Faculty of Medicine of University of Baghdad, Baghdad, Iraq.
- 2.4. Study Population. Participants who signed an informed consent form were evaluated for inclusion and exclusion criteria.
- 2.5. Inclusion Criteria. Individuals over 18 years of age with a diagnosis of type I or II diabetes mellitus and suspected diabetic retinopathy, regardless of gender and race, were included. We considered as suspicion of retinopathy diabetic patients with prolonged evolution time (>5 years for type I diabetes and >10 years for type II) and/or complaints related to low vision.
- 2.6. Exclusion Criteria. The following were excluded from the study:
  - (i) Diabetic individuals with media opacities that prevented the fundus from being performed
  - (ii) Diabetic individuals with causes of low vision other than diabetic retinopathy
  - (iii) Diabetic individuals undergoing previous treatments for diabetic retinopathy (laser, antiangiogenic injections, vitrectomy)
- 2.7. Ophthalmological Evaluation. All patients underwent an eye examination with the following:
  - (i) Measurement of visual acuity
  - (ii) Applanation tonometry
  - (iii) Biomicroscopy with retinal assessment
  - (iv) Indirect binocular ophthalmoscopy
  - (v) Color photograph
  - (vi) Retinography analysis was performed by a masked retina specialist for ophthalmological evaluation data; when any sign of diabetic retinopathy was detected, the classification used was that of the WHO guidelines

Based on this classification, the specialist separated the photos into four classes:

- (i) Absence of diabetic retinopathy
- (ii) Mild diabetic retinopathy
- (iii) Moderate/severe diabetic retinopathy
- (iv) Proliferative diabetic retinopathy
- 2.8. Techniques Used in Ophthalmological Evaluation
- 2.8.1. Visual Acuity. Verification of visual acuity was performed with better optical correction when used by the patient, using an optotype table from the Early Treatment Diabetic Retinopathy Study (ETDRS) [5]. To analyze the level of visual impairment (low vision and blindness) of the patients, the WHO parameters were used [6].
- 2.8.2. Biomicroscopy. Biomicroscopic examination was performed using slit lamps from the FMB retinal outpatient clinic, evaluating the conjunctiva, cornea, anterior chamber, iris, pupil, lens, vitreous, and retina, with the aid of a 78-diopter lens (Volk Optical Inc., Mentor, USA).
- 2.8.3. Applanation Tonometry. The measurement of intraocular pressure was performed with the patient in the sitting position, after topical anaesthesia with 1 drop of eye drops prepared with 5 mg of 0.5% proxymetacaine hydrochloride (Merck, Germany) and 15 drops of sodium fluorescein eye drops (Sigma-Aldrich), using a Goldmann-type applanation tonometer, model R-900 (Haag-Streit AG., Bern, Switzerland), coupled to a lamp crack.
- 2.8.4. Pupillary Dilation. For the biomicroscopy of the lens, vitreous, and retina, as well as for fundus examination, the individuals underwent pupillary dilation with three instillations, spaced five minutes apart, of a drop of 1% tropicamide eye drops (Mydriacyl® 1%, Sigma-Aldrich).
- 2.8.5. Fundoscopy. Fundus examination was performed with an indirect binocular ophthalmoscope, model H-OI (Neitz Instruments Co. Ltd., Tokyo, Japan), and a 20-diopter lens (Nikon Corporation, Tokyo, Japan), with the patient in the supine position and the pupil dilated.
- 2.8.6. Retinography. Retinography was performed using a Visucam retinograph (Carl Zeiss Meditec, Inc., Dublin, USA), with the recording of the retina and optic disc.
- 2.9. Evaluation of Images by Machine Learning: The Evaluation of the Images Was Carried Out at the Department of Computing, Faculty of Sciences
- 2.9.1. Optimum-Path Forest (OPF)
  - (i) Training: before the OPF was applied to the patients included in the study, a training set of the system was assembled, using photographic images of other diabetic patients with and without diabetic retinopathy, taken by the same Visucam retinal imager,

and also analyzed by a retina specialist (137 photos). The training enabled the OPF algorithm to recognize the patterns of diabetic retinopathy and normality. The procedures were as follows.

- (ii) Assembly of the OPF training set: referring to the photographic characteristics of each patient. In this set, the results of the specialist's diagnosis were included, informing whether the patient had the disease or not.
- (iii) OPF training: making use of the training set, in order to recognize disease patterns.
- (iv) After training: application of the OPF in the new batch of fundus images (of the patients included in the study) and comparing the result generated by the system with the specialist's diagnosis (gold standard).

2.9.2. Restricted Boltzmann Machine (RBM) and Deep Belief Networks (DBNs). The extraction of features used in the OPF was done by RBM/DBN [6, 7].

1 Experimental Setup. Before performing the classification, the dataset was preprocessed as follows: all images were resized to  $200 \times 200$  pixels, and the RGB images (red, green, blue) were converted to gray scale with pixel intensities in the interval [0; 1].

Table 1 describes six different models employed to solve the classification problem. Such models vary in the number of hidden layers and hidden neurons. Also, it is important to note that the "Norm." (normalized) setting employs a naive (native) RBM architecture as it employs a different preprocessing approach, which was only effective for a hidden layer (single RBM).

Considering a sample image  $x_i \varepsilon X$ , where X represents the entire dataset, the aforementioned preprocessing for the "Norm." was a simple "statistical trick" to improve the quality of the gray scale of  $x_i$ , which corresponds to the following formula:

$$\widetilde{x}_i = \frac{\left(x_i - \overline{x}_i\right)}{\sigma(x_i)},\tag{1}$$

where  $\tilde{x}_i$  represents the normalized image,  $\bar{x}_i$  represents the average pixel values of the image  $x_i$ , and  $\sigma(x_i)$  represents the standard deviations of  $x_i$  pixels. Then, the entire dataset was normalized using a Gaussian distribution with a mean of zero and a unity standard deviation. This fact allowed us to make use of the well-known Gaussian–Bernoulli RBM, a variant of the RBM that samples the visible values of a Gaussian distribution, given the latent binary space. It is important to explain that only one model was chosen to use visible Gaussian units, since preliminary experiments with more than one hidden layer did not show good convergence. After the end of the RBM learning process, the datasets composed of resources extracted from the hidden layer units fed the supervised learning algorithm (OPF). Four models were considered to extract features and classify the images:

TABLE 1: Different configurations.

Model	DBN templates
M1	i:1000:1000:4
M2	i:2000:2000:4
M3	<i>i</i> :500:500:4
M4	<i>i</i> :500:500:500:4
M5	<i>i</i> :1000:1000:1000:4
M6	<i>i</i> :2000:2000:2000:4
Standard	i:1000:4

DBN: deep belief network and i: number of neurons in the input layer.

- (i) OPF-500: where 500 image features were extracted by the RBM and classified by the OPF
- (ii) OPF-1000, where 1000 image features were extracted by the RBM and classified by the OPF
- (iii) RBM-500: where 500 image features were extracted and classified by RBM
- (iv) RBM-1000: where 1000 image features were extracted and classified by RBM

2.10. Statistical Analysis. To analyze the results, the Wilcoxon test was used to compare whether the position measures of two samples are equal in the case where the samples are dependent. Sensitivity and specificity values were calculated for the algorithms used to evaluate the images. The hypothesis that the probability of error was less than 5% (p < 0.05) was assumed as true.

## 3. Results

3.1. General Characterization of the Sample. A total of 73 patients (122 eyes) were evaluated, 50.7% men and 49.3% women. Age ranged from 36 to 79 years, with an average of 59.7 years. Most patients classified themselves as "white" (86%). Seventy patients had type 2 DM, and only three had type 1. The insulin treatment regimen occurred in 53.4% of the patients. Among the diseases associated with DM, the main one was systemic arterial hypertension (70%). The association with smoking and alcoholism was small, corresponding to 12.3% and 4.3% of patients, respectively (Table 2). The main complaint of patients was poor vision, both near (93.1%) and far (91.8%), even among users of optical correction (63.1%). Table 3 shows the level of visual impairment in the 122 eyes studied. As for ocular history, the three most mentioned were cataract (10.9%), glaucoma (6.8%), and previous surgeries (9.6%).

3.2. Ophthalmological Evaluation. The specialist's evaluation of 122 retinographies revealed the absence of diabetic retinopathy (DR) in 17 eyes (Figure 1) and the presence of DR in 105 eyes (86%), classified as follows:

- (i) Mild diabetic retinopathy: 19 (18.1%)
- (ii) Moderate/severe diabetic retinopathy: 77 (73.3%)
- (iii) Proliferative diabetic retinopathy: 9 (8.6%) (Figure 2)

Table 2: Characteristics of the evaluation of the 73 patients (122 eyes) studied.

Characteristics of the initial assessment	73 patients (122 eyes)
Middle ages)	59.7 years
Gender (male/female)	37 (50.7%)/36 (49.3%)
Diabetes (type I/type II)	3 (4.1%)/70 (95.9%)
Insulin dependents	39 (53.4%)
Arterial hypertension	70.00%
Smoking	12.30%
Elitism	4.30%

TABLE 3: Distribution of visual impairment in 122 eyes studied.

Visual commitment* Absent	Visual acuity** ≥0.3 and <1	Number of eyes 96	Percentage 78.60
Moderate low vision	≥0.1 and <0.3	16	13.20
Low vision serious	≥0.05 to <0.1	0	0
Blindness	< 0.05	10	8.20
Total		122	100

<sup>\*</sup>As per the World Health Organization (WHO) [6]; \*\*VA with the best optical correction.



FIGURE 1: Retinography of the left eye of a patient without DR.



FIGURE 2: Retinography of the right eye of a patient with proliferative DR.

3.3. Image Evaluation. The set of images evaluated contained 259 photographic images (137 from the training set, used for learning the system, and 122 new ones, from 73 patients included in the study, used in the tests).

Fifteen sets of experiments were performed with a repetition of 30 cycles in each set, varying the ratio of training/test sets at random, in each repetition of each set (80% for training and 20% for tests). Thus, all samples participated in the tests (in each execution different groups of 20% were used for training, totaling 100% of the samples). After the execution of the 15 series, the results presented in Table 4 were obtained.

Tables 5 and 6 show the average performance of the 15 series of experiments with RBM-500 and RBM-1000, considering four classes (no DR, mild DR, moderate/severe DR, and proliferative DR). Table 7 shows the average performance of the 15 experiment series with OPF-500 and OPF-1000

Considering only the presence or absence of the disease (diabetic retinopathy), Tables 8 and 9 show the performance of RBM (500 and 1000) and OPF (500 and 1000).

Sensitivity measured the model's ability to correctly identify individuals who had diabetic retinopathy, whereas specificity measured those who did not (Table 9). Sensitivity values for the RBM were high, especially for the RBM-500 (100%) versus 95.7% for the RBM-1000 (Table 8). The OPF-500 had higher sensitivity values (95.5%) than the OPF-1000 (86, 4%). As for specificity, both RBM-1000 and OPF-1000 presented 100%; that is, they correctly identified all images that did not have DR. This capacity was lower in the RBM-500 (64.3%) and OPF-500 (87.5%) (Table 10).

## 4. Discussion

Early screening is one of the pillars of the prevention of blindness in diabetics, since retinopathy can be asymptomatic in the beginning and vision loss can only be noticed when the disease is advanced (51). The results of two large studies on diabetic retinopathy, the Diabetic Retinopathy Study (DRS) and the Early Treatment Diabetic Retinopathy Study (ETDRS), showed that early and effective treatment can prevent severe vision loss and reduce the risk of blindness from the disease in about 50% (52–54).

Machine learning automatic disease detection models provide a quick and practical means of screening for diabetic patients still undiagnosed with retinopathy, especially those living in remote communities without access to specialized services (41).

This study used two image classifiers with different characteristics, the OPF (optimum-path forest) and the RBM (restricted Boltzmann machine). The metrics used to assess the classifiers in this research were accuracy, sensitivity, and specificity.

The results obtained showed that the RBM, which used 1000 characteristics extracted from the analyzed photographic images, had the best overall performance in terms of diagnostic accuracy (89.5%); that is, it was the model that had the greatest ability to correctly differentiate images with and without DR signals after 15 sets of experiments.

Table 4: Precision results for optimum-path forest (OPF) (500 and 1000) and restricted Boltzmann machine (RBM) (500 and 1000).

Repetitions	OPF-500	OPF-1000	RBM-500	RBM-1000
0	0.7667	0.7667	0.868	0.868
1	0.8	0.8	0.789	0.895
2	0.7667	0.8	0.737	0.921
3	0.8	0.8	0.789	0.895
4	0.8333	0.8	0.711	0.921
5	0.7333	0.7333	0.763	0.868
6	0.8	0.7667	0.711	0.921
7	0.7667	0.7667	0.763	0.895
8	0.7667	0.7333	0.763	0.842
9	0.7667	0.7667	0.763	0.895
10	0.8	0.8	0.789	0.895
11	0.8667	0.7667	0.737	0.947
12	0.7667	0.7333	0.711	0.895
13	0.8	0.7667	0.737	0.895
14	0.6667	0.7667	0.816	0.868
Average	78	77.11	76.31	89.47
Standard deviation	4.51	2.48	4.31	2.64

Values in bold indicate the best results (mean and standard deviation) by the Wilcoxon test, with 5% significance.

Table 5: Average performance of the 15 series of experiments with RBM 500, considering four classes (no DR, light DR, mod/severe, and proliferative).

RBM	I 500	Machine learning			Hit %	
Specialist		Without DR		With DR		1110 /0
орес	iuiist	Absence	Light	Mod/record	Proliferative	
Without DR	Absence	9	0	5	0	9/14 (64.2%)
	Light	0	2	2	0	2/4 50%
With DR	Mod/record	0	2	11	0	11/13 85%
	Proliferative	0	0	0	7	7/7 100%

RD: diabetic retinopathy, mod: moderate, and RBM: restricted Boltzmann machine.

Table 6: Average performance of the 15 series of experiments with RBM-1000, considering four classes (no DR, light DR, mod/severe, and proliferative).

RBM	-1000	Machine learning			Hit %	
Spac	ialist	Without DR		With DR		1111 /0
Spec	idiist	Absence	Light	Mod/record	Proliferative	
Without DR	Absence	14	0	0	0	14/14 (100%)
	Light	0	3	1	0	3/4 (75%)
With DR	Mod/record	1	1	10	0	10/12 (83.3%)
	Proliferative	0	0	0	77 7	7/7 (100%)

As for sensitivity, the four models analyzed (RBM-500 and RBM-1000 and OPF-500 and OPF-1000) presented performance above 80%, considered ideal for imaging screening tests (55). The RBM 500 achieved 100% accuracy when the RD was present, followed by the RBM-1000 (95.7%). The OPF-500 had a sensitivity of 95.5%, whereas the OPF-1000 had a sensitivity of 86.4%.

Regarding specificity, the RBM-1000 and OPF-1000 classifiers showed the best results (100% correctness when

there was no disease). The OPF-500 had the lowest performance (87.5%), but still above 80%, whereas the RBM-500 had the worst performance of the four models, with 64.3%.

Results similar to those of the present study were found in a systematic review of 2019 [8], which evaluated deep learning detection models for DR image classification. The sensitivity values of eight studies evaluated ranged from 80 to 100% and specificity values from 84 to 90%, indicating the

Table 7: Average performance of the 15 series of experiments with OPF-500 and OPF-1000, considering four classes (no DR, light DR, mod/severe, and proliferative).

OPF-500		Machine learning			Hit %	
Snec	cialist	Without DR		With DR		1110 /0
<b>З</b> рес	lalist	Absence	Light	Mod/record	Proliferative	
Without DR	Absence	7	0	1	0	07/08 (87.5%)
	Light	0	2	2	0	02/04 (50%)
With DR	Mod/record	1	2	10	1	10/14 (71.4%)
	Proliferative	0	0	0	4	4/4 (100%)
OPF-1000						
Without DR	Absence	8	0	0	0	8/8 (100%)
	Light	0	2	2	0	2/4 (50%)
With DR	Mod/Record	3	1	9	1	9/14 (64.2%)
	Proliferative	0	0	0	4	4/4 (100%)

RD: diabetic retinopathy, mod: moderate, and OPF: optimum-path forest.

Table 8: Average performance of the 15 series of experiments with RBM-500 and RBM-1000 (considering disease versus nondisease).

RBM-500	Machine l	Machine learning	
Specialist	Without DR	With DR	
Without DR	9	5	8/14 (64.3%)
With DR	0	24	24/24 (100%)
RBM-1000			
Without DR	14	0	14/14 (100%)
With DR	1	22	22/23 (95.7%)

RBM: restricted Boltzmann machine.

Table 9: Average performance of the 15 series of experiments with OPF-500 and OPF-1000 (considering disease versus nondisease).

OPF-500	Machine l	Machine learning	
Specialist	Without DR	With DR	
Without DR	7	1	7/8 (87.5%)
With DR	1	21	21/22 (95.5%)
OPF-1000			_
Without DR	8	0	8/8 (100%)
With DR	3	18	19/22 (86.4%)

potential of using automatic methods of artificial intelligence as screening tools [9].

Regarding the performance of the RBM and OPF algorithms in differentiating the severity of diabetic retinopathy, the results of the present study showed better performance of the classifiers in identifying signs of proliferative retinopathy (100% correctness in the four models) and worse performance in detecting initial or mild, with no obvious signs of the disease. These results do not compromise the use of algorithms in DR screening, since the focus of current international screening programs is the early identification of the so-called "referential diabetic retinopathy," which presents signs of advanced disease and indication for immediate treatment [10].

Following this line of conduct, Nagasawa et al. [11] used a deep learning model to identify only the untreated

Table 10: Sensitivity and specificity values of the RBM (500 and 1000) and OPF (500 and 1000) in identifying disease versus nondisease and their respective 95% confidence intervals.

	Sensitivity (%)	95% CI	Specificity (%)	95% CI
RBM-500	100	82.8%-100%	64.30	35.6%-86.0%
RBM-1000	95.70	76%-99.8%	100	76.2%-100%
OPF-500	95.50	75.1%-97.8%	87.50	46.7%-99.3%
OPF-1000	86.40	64%-96.4%	100	59.8%-100%

The sensitivity of RBM-500 is 100%, and the sensitivity of OPF-500 is 95.50%, which are indicated in bold.

proliferative form, due to the importance of indicating immediate treatment, and obtained a sensitivity of 94.7% and a specificity of 97.2% [12]. Other studies using artificial intelligence medical devices have shown good results, especially in the moderate and severe forms of DR and with the identification of specific signs of these forms, such as the presence of exudates [13].

The present study had some limitations. The number of retinographs analyzed was lower than planned due to technical problems with the retinograph, which led to the loss of a portion of the images already stored. There was also variation in the number of images evaluated in each class, but the imbalance was artificially compensated by the generation of synthetic images by the algorithms. Furthermore, it was not possible to use an external image bank to train the classifier system within the research schedule, which would help in the external validation process of the analyzed algorithms. According to Gulshan et al., access to a large database of high-quality international images facilitates the validation of deep learning models, as they contain data from populations with different ethnic characteristics [11].

A relevant issue in Iraq, due to the high miscegenation of the population, is the wide variation in retinal color, a detail that becomes positive as it provides more information to the algorithms and facilitates the differentiation between the natural components of the fundus and the signs of DR [10, 12]. In the present study, 96% of the participants declared themselves "white," which would minimize the effect

of this factor. However, self-declaration does not exclude the possibility of miscegenation. Most validated deep learning systems used public external image banks that mostly make available images of a single ethnicity [11]. To verify the influence of the background color factor, Li et al. studied four ethnic groups with diabetic retinopathy (Chinese, Malay, Caucasian Australian, and Indigenous Australian) with different degrees of physiological retinal pigmentation and concluded that this factor did not affect the performance of the tested deep learning algorithm. According to the authors, this result could be extrapolated to most populations in the world [5]. Although the results obtained in this study are promising, there are still challenges to be overcome before the use of automatic disease detection models through machine learning becomes a reality in clinical practice. One of them is the capture of good quality retinal images, which enable the correct classification of DR by algorithms. Difficulties are manifold: inadequate pupillary dilation, opacity of the eye's transparent media, problems with contrast and focus, poor technique, or inexperience of the retina camera operator. Imaging rates with low quality and without classification conditions reach 25% in some studies [4, 14]. However, scientific evidence has shown that artificial intelligence itself may solve this problem in the near future. A study achieved good results by adapting a deep neural network to correct the quality of retinal images through reprocessing [6]. The authors suggest that the proposed system has the potential to be implemented in the healthcare network. Another important issue that must be considered is how AI models can fit into the healthcare network. For their use to be effective in preventing blindness by DR, a restructuring involving implementation costs, management logistics, health policies, and acceptance by health professionals and patients would be necessary.

The algorithms would be incorporated into the computer system of the primary care point of patients with suspected DR, which would be equipped with a traditional retinograph or a portable retinal image capture device. Another option would be the development of mobile applications with the algorithms, which could be made available to health professionals far from specialized retina centers. With the system in place, automated screening would be easy and fast and the results known at the time of screening. A pilot study carried out in Australia compared the use of an automated screening model, where a deep learning algorithm provided real-time reporting of results, with a manual model in which retinal images were transferred to a specialized classification center and the results transferred to the patient only two weeks after the evaluation. Findings showed that 96% of participants said they were satisfied or very satisfied with the automated screening model and about 80% reported preferring the automated screening model over the manual model, suggesting a good level of acceptance of this system

Machine learning identification of diabetic retinopathy represents an important technology, offering great potential to improve the efficiency and accessibility of DR screening programs, particularly in developing countries where the number and location of specialist physicians are poor and patient identification needing faster care to avoid a major and debilitating visual loss is imperative.

The novelty of the present study was the comparison of two different image analysis models (OPF and RBM) for the automatic detection of DR, through accuracy, sensitivity, and specificity, and the demonstration that the RBM performed better than the OPF, especially in sensitivity, which is important for clinical diagnosis. The present study also demonstrated that the combination of algorithms can be a strategy to improve the performance of DR screening by artificial intelligence.

For future research, it is proposed to use a larger database and the generation of synthetic images to improve the training process, using deep neural networks such as GANs (generative adversarial nets), already used in some studies with retinal vessels [5, 8], which are capable of generating as many synthetic images as the user requires, with good quality. The development of a mobile app is also being planned as the next step in this line of research.

#### 5. Conclusions

Given the results obtained and under the experimental conditions of this study, we can conclude that

- (a) The RBM-1000 performed best in terms of diagnostic accuracy (89.47  $\pm$  2.64) in the overall analysis
- (b) The RBM-500 was superior in the automatic detection of signs of diabetic retinopathy on photographic images (100% sensitivity)
- (c) As for specificity, both RBM-1000 and OPF-1000 presented 100%; that is, they correctly identified all images without diabetic retinopathy
- (d) Machine learning automatic disease detection models, especially the RBM, showed good diagnostic accuracy and high sensitivity and have potential for use in screening for diabetic retinopathy

## **Data Availability**

The data used to support the findings of this study are included within the article.

## **Disclosure**

This study was performed as a part of the Employment of Institutions.

## **Conflicts of Interest**

The authors declare that they have no conflicts of interest regarding the publication of this article.

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## References

- [1] Q. Zou, K. Qu, Y. Luo, D. Yin, Y. Ju, and H. Tang, "Predicting diabetes mellitus with machine learning techniques," *Frontiers in Genetics*, vol. 9, 2018.
- [2] T. J. MacGillivray, J. R. Cameron, Q. Zhang et al., "Suitability of UK biobank retinal images for automatic analysis of morphometric properties of the vasculature," *PLoS One*, vol. 10, no. 5, Article ID e0127914, 2015.
- [3] P. Costa, A. Galdran, M. I. Meyer et al., "End-to-End adversarial retinal image synthesis," *IEEE Transactions on Medical Imaging*, vol. 37, no. 3, pp. 781–791, 2018.
- [4] P. H. Scanlon, C. Foy, R. Malhotra, and S. J. Aldington, "The influence of age, duration of diabetes, cataract, and pupil size on image quality in digital photographic retinal screening," *Diabetes Care*, vol. 28, no. 10, pp. 2448–2453, 2005.
- [5] P. H. Scanlon, R. Malhotra, G. Thomas et al., "The effectiveness of screening for diabetic retinopathy by digital imaging photography and technician ophthalmoscopy," *Diabetic Medicine*, vol. 20, no. 6, pp. 467–474, 2003.
- [6] V. Gulshan, L. Peng, M. Coram et al., "Development and validation of a deep learning algorithm for detection of diabetic retinopathy in retinal fundus photographs," *Journal of* the American Medical Association, vol. 316, no. 22, pp. 2402–2410, 2016.
- [7] U. Alam, O. Asghar, S. Azmi, and R. A. Malik, "General aspects of diabetes mellitus," *Diabetes and the Nervous System*, vol. 126, pp. 211–222, 2014.
- [8] P. Khojasteh, L. A. Passos Júnior, T. Carvalho et al., "Exudate detection in fundus images using deeply learnable features," *Computers in Biology and Medicine*, vol. 104, no. 1, pp. 62–69, 2018.
- [9] S. Sivanandam and S. N. Deepa, Introduction to Neural Networks Using Matlab 6.0, p. 656, Tata McGraw-Hill Education, New York, NY, USA, 2006.
- [10] R. O. Duda, P. E. Hart, and D. G. Stork, *Pattern Classification*, John Wiley & Sons, Hoboken, NJ, USA, 2nd edition, 2001.
- [11] T. Nagasawa, H. Tabuchi, H. Masumoto et al., "Accuracy of ultrawide-field fundus ophthalmoscopy-assisted deep learning of detecting treatment-naïve proliferative diabetic retinopathy," *International Ophthalmology*, vol. 39, no. 10, pp. 2153–2159, 2019.
- [12] C. P. Wilkinson, F. L. Ferris 3rd, R. E. Klein et al., "Global Diabetic Retinopathy Project Group. Proposed international diabetic retinopathy and diabetic macular edema disease severity scales," *Ophthalmology*, vol. 110, no. 9, pp. 1677–1682, 2003
- [13] A. Tufail, C. Rudisill, C. Egan et al., "Automated diabetic retinopathy image assessment software: diagnostic accuracy and cost-effectiveness compared with human graders," *Ophthalmology*, vol. 124, no. 3, pp. 343–351, 2017.
- [14] Diabetic Retinopathy Study Research Group, "Grading diabetic retinopathy from stereoscopic color fundus photographs. An extension of the modified Airlie House classification. ETDRS report number 10. Early treatment diabetic retinopathy Study research group," Ophthalmology, vol. 98, no. 5 Suppl, pp. 786–806, 1991.