

# A Systematic Analysis of Advanced Machine Learning Techniques for Fundus Image-based Diabetic Retinopathy Detection

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## ABSTRACT

Diabetic retinopathy (DR) is a leading cause of blindness worldwide, necessitating early detection and accurate classification to mitigate these its progression. Fundus imaging has emerged as a non-invasive and reliable method for Diabetic retinopathy (DR) diagnosis. Recent advancements in machine learning (ML) have significantly improved the precision and efficiency of fundus image-based DR detection. This paper provides a systematic analysis of advanced ML techniques employed in Diabetic retinopathy (DR) classification, emphasizing both traditional and deep learning approaches. It explores preprocessing methods, feature extraction techniques, and state-of-the-art classification algorithms, highlighting their effectiveness and limitations. Key challenges such as imbalanced datasets, variability in image quality, and interpretability are discussed, alongside strategies to address issues. The analysis also examines emerging trends, including hybrid models and explainable AI, offering insights into future research directions. This review aims to serve as a comprehensive resource for researchers and practitioners, guiding the development of more robust and accurate ML-based solutions for DR detection.

Keywords: Diabetic Retinopathy, Fundus Imaging, Machine Learning, Deep Learning, Image Classification, Feature Extraction.

## I. INTRODUCTION

Diabetic retinopathy is a critical complication associated with diabetes and a leading cause of preventable blindness globally. With the increasing prevalence of diabetes, particularly in low- and

middle-income regions, the global health impact of DR is becoming more pronounced [Kropp et.al(2023)]. Timely diagnosis and appropriate treatment of DR are essential to significantly lower the risk of vision impairment. However, traditional

screening techniques are labor-intensive, susceptible to human error, and rely on specialized expertise, which is often limited in areas with fewer resources.

Fundus imaging has emerged as a cornerstone for DR diagnosis, providing a non-invasive and cost-effective approach to assess retinal abnormalities. This imaging technique enables clinicians to detect early signs of DR, such as microaneurysms, hemorrhages, and exudates, which are crucial for timely intervention. However, the increasing demand for DR screening necessitates scalable and efficient diagnostic tools that can complement manual assessments.

Machine learning (ML) has demonstrated great promise in addressing this need by automating the analysis of fundus images with high accuracy and speed [Grzybowski et.al.,2024]. Advanced ML techniques, particularly deep learning models, have outperformed traditional image analysis methods by leveraging large datasets and extracting complicated patterns from fundus images. These innovations not only enhance diagnostic precision but also hold potential for deployment in remote or underserved areas, thus bridging the gap in healthcare accessibility [Ranjan& Choubey.,2024].

### 1.1 Objectives and scope of the paper

This Research aims to provide a comprehensive analysis of advanced ML techniques for fundus image-based DR detection. This study explores preprocessing methods, classification algorithms, and the challenges associated with implementing these models in clinical settings. By examining the current scenario and emerging trends, this review seeks to guide researchers and practitioners in developing robust and scalable ML-based solutions for DR diagnosis.

## II. PREPROCESSING AND FEATURE EXTRACTION IN FUNDUS IMAGING

Fundus image analysis presents several challenges that can affect the accuracy of machine learning models. Variability in image quality, caused by differences in imaging devices, lighting conditions, and patient-specific factors such as pupil dilation, often leads to inconsistencies [Pandey et al., 2023]. Additionally, noise, uneven illumination, and artifacts in the images can obscure important retinal features, making accurate detection and classification of diabetic retinopathy (DR) challenging. Addressing these issues is essential to ensure reliable model performance [Basreddy & Veerashetty., 2023].

Preprocessing techniques play a pivotal role in enhancing the quality of fundus images before they are analyzed by machine learning algorithms. Common methods include image enhancement to improve contrast and visibility of retinal structures, noise reduction to remove irrelevant artifacts, and normalization to standardize image intensities [Das & Nayak., 2023]. Techniques such as histogram equalization, Gaussian filtering, and adaptive thresholding are frequently used to prepare the images for feature extraction. These steps ensure that key retinal features are preserved while reducing variability, which can improve model robustness and generalization.

Feature extraction is another critical step that focuses on identifying the most relevant patterns and structures within the fundus images [Vairamani., (2023)]. Texture features, such as the distribution of pixel intensities, can help in detecting microaneurysms and hemorrhages. Color features are often used to identify regions with exudates or edema, while vascular features, such as vessel density and branching patterns, can provide insights into DR progression. These extracted features form the foundation for training machine learning models, enabling them to classify DR stages with higher accuracy.

DR is the primary cause of newly diagnosed blindness, with retinal microaneurysm (MA) being one of the first clinical indicators of DR. Nonetheless, directly seeing MA with the naked eye is challenging because to their little size and insufficient contrasting against the natural backdrop. This work [Zhang., 2023] introduces a novel approach for the automated identification of microaneurysms in fundus pictures. The spectrum of ocular disorders is varied, and each requires a personalized, patient-specific intervention. CNNs have shown considerable promise in several disciplines, particularly in the identification of numerous ocular disorders [Mayya et al., 2023].

Proactive surveillance of DR illness is essential in averting cataracts including deafness, making it a primary contributor to global vision impairment.

This study [Navaneethan, & Devarajan., 2024] presents an innovative approach, the Modified Generative Adversarial-based Crossing Salp Grasshopper (MGA-CSG) method, designed for the early detection and accurate classification of diabetic retinal disorders using fundus imaging datasets. Manually mapping the blood vessels in retinal images poses several challenges, including being highly time-intensive and prone to errors due to the complex structure of the vascular network and the large volume of images requiring assessment. Therefore, an automated system for segmenting and extracting relevant clinical data from retinal vessels is crucial to support optometrists and ophthalmologists in diagnosing retinal diseases and initiating timely treatment [Abdulsahib et al., 2022].

By improving image quality and extracting meaningful features, preprocessing serves as a crucial enabler for the effective application of machine learning in fundus image analysis. It not only enhances the reliability of the models but also minimizes errors, making the diagnostic process more efficient and accurate.

## III. ADVANCED MACHINE LEARNING TECHNIQUES FOR DR CLASSIFICATION

Machine learning (ML) has transformed medical image analysis, offering powerful techniques for classifying DR using fundus images. Conventional ML models, including Support Vector Machines (SVM) and Random Forests, have been extensively employed for early detection of DR [Atwany et al., 2022]. These models rely on manually extracted features, such as texture, color, and vessel structures, to distinguish between normal and pathological images [Abushawish et al., 2024]. While effective for small datasets and straightforward tasks, their performance is often limited by the quality of feature engineering and the inability to capture complex patterns in fundus images [Selvachandran et al., 2023].

Deep learning approaches have significantly advanced the state of DR classification by automating feature extraction and enabling the analysis of high-dimensional data [Das et al., 2022]. Convolutional Neural Networks (CNNs), in particular, have demonstrated exceptional performance in fundus image classification due to their ability to learn spatial hierarchies of features directly from raw pixel data [Saranya et al., 2023]. Variants of CNNs, such as transfer learning and attention-based models, have further enhanced

classification accuracy by leveraging pre-trained networks and focusing on critical regions of the retina. Recurrent Neural Networks (RNNs) and their gated variants, while less commonly used in image analysis, have been explored for sequential modeling in longitudinal DR studies [Mohanty et.al.,2023].

Diagnosing diabetic retinopathy (DR) through manual methods is often a labor-intensive and costly process, requiring skilled ophthalmologists to analyze and interpret digital fundus images. This study seeks to systematically review and evaluate advanced research on DR diagnosis using deep learning techniques [Farooq et.al.,2022]. An automated ensemble deep learning framework is introduced in this research for detecting and classifying DR. By combining multiple deep learning models, the ensemble approach enhances prediction accuracy and achieves superior performance compared to individual models. Specifically, the study integrates modified DenseNet101 and ResNeXt models for DR detection. The ResNeXt model, an enhancement of ResNet, incorporates innovative structural improvements for better accuracy [Mondal et.al.,2022].

Additionally, the Gray Wolf Optimization-Extreme Learning Machine (GWO-ELM) is highlighted as a highly effective machine learning algorithm for classification tasks. Although widely recognized for its accuracy, it has yet to be fully explored in the context of DR detection [Albadr et.al.,2022]. The research presents a modified capsules structure for the identification and categorization of DR. Features are collected from fundus imagery employing the combination of convolution and main capsule layers, followed by the estimation of the likelihood of class membership via the group capsule level and softmax layer [Kalyaniet.al.,2023]. An advanced Convolutional Neural Network (CNN) architecture, referred to as ADL-CNN, has been designed for the autonomous extraction of segmented retinal features. This innovative model utilizes a dual-stage process to enhance feature extraction and segmentation efficiency [Özbay, E. (2023)]. The model proposes a multi-stream composition deep neural network for the classification of DR severity. The proposed approach integrates deep neural networks with Principal Component Analysis (PCA) to effectively capture both inter-class and intra-class variations from raw image features [Mustafa et.al.,2022].

This study proposed a two-stage innovative method for automated diabetic retinopathy categorization. To address the limited proportion of positive cases in the asymmetrical Optic Disk (OD) and blood vessels (BV) recognition framework, filtering techniques and information enrichment methods are applied to enhance both the quantity and quality of the images [Bilal et.al.,2022]. Developed a comprehensive

pipeline to extract pertinent features using various preprocessing approaches, an image segmentation architecture (DR-UNet) including atrous spatial pyramid pooling, and robust attention-aware deep neural multilayer network comprising diverse modules based on ResidualNet [Maaliw et.al.,2023].

This study aims to reduce reliance on human labor by introducing a Deep Graph Connection Network (DGCN) for automated diabetic retinopathy grading, eliminating the need for expert annotations [Zhang et.al.,2022]. Regular annual eye examinations are crucial for the early identification of such conditions, thereby increasing the chances of effective treatment. Consequently, it employs fundus cameras for acquiring retinal pictures; nonetheless, its size and expense render it impractical for widespread screening [Gupta et.al.,2022].

Manual diagnosis of DR with the assistance of an optometrist has been a cumbersome and labor-intensive process. This work concentrates on the diagnosis of diabetic retinopathy and the analysis of various phases of DR, using Deep Learning with transfer learning methods [Raja Sarobin et.al.,2022]. In contemporary ophthalmology, retinal image processing has emerged as a prevalent method for disease diagnosis. Ophthalmic surgeons and automated systems frequently use fundus angiography to identify clinical indications of diabetic retinopathy for early identification [Abbood et.al.,2022].

Hybrid and ensemble models combine the strengths of multiple ML algorithms to achieve superior performance. For instance, hybrid models that integrate CNNs with traditional classifiers or ensemble techniques, such as bagging and boosting, can improve robustness and reduce overfitting. These approaches are particularly valuable in addressing class imbalances and enhancing the sensitivity of DR detection models.

Comparative studies of different models highlight their strengths and limitations in fundus image classification. While deep learning models generally outperform traditional methods in terms of accuracy and scalability, they require large, annotated datasets and significant computational resources. On the other hand, traditional models remain viable for smaller datasets and resource-constrained settings. Understanding these trade-offs is essential for selecting the most appropriate technique for specific clinical applications. The comparison of DR classification methods are summarized in Table 1.

**Table 1: Comparison of Methods for DR Classification and Detection**

Ref.	Method	Dataset(s)	Performance	Strengths	Limitations
[19]	Deep learning techniques (CNNs, ECNNs, DNNs) for DR classification	Multiple datasets analyzed	35% used CNNs, 26% ECNNs, 13% DNNs; improving classification accuracy	Systematic analysis of algorithms; DR grading and staging; taxonomy of DR	Generalized results; limited dataset-specific performance details
[20]	Ensemble of modified DenseNet101 and ResNeXt models	APTOS19, DIARETDB1	Accuracy: 96.98% (2-class), 86.08% (5-class); Precision/Recall (2-class): 0.97; (5-class): 0.76/0.82	Improved feature usage efficiency and better prediction accuracy	High-class imbalance tackled via GAN, but may need further validation on larger datasets
[21]	GWO-ELM with HOG-PCA for DR detection	APTOS-2019, IDRiD	Accuracy: 99.47% (binary), 96.21% (multi-class, APTOS); 99.04% (binary), 96.15% (multi-class, IDRiD)	Excellent accuracy; addresses overfitting; low-dimensional feature extraction	Limited exploration of other datasets or alternative feature extraction techniques
[22]	Reformed Capsule Network for lesion-based classification	Messidor	Accuracy: 97.98% (healthy), 97.65% (stages 1–3)	Capsule network captures spatial hierarchies; strong stage-wise performance	Only tested on a single dataset; lacks comparison with more recent architectures
[23]	ADL-CNN with artificial bee colony (ABC) segmentation for DR stage classification	EyePACS	Accuracy: 99.66%; Sensitivity: 93.76%; Specificity: 96.71%; F-measure: 94.58	High classification accuracy; segmentation-based lesion detection	Focused only on EyePACS dataset; generalizability to other datasets needs validation
[24]	Multi-stream ensemble deep network with PCA and ensemble classifiers (AdaBoost, RF)	Messidor-2, EyePACS	Accuracy: 95.58%	Multi-stream networks capture inter- and intra-class variations	Computationally expensive; lacks lightweight model considerations
[25]	Two-stage hybrid CNN-SVD model with U-Net-based OD and BV segmentation	EyePACS-1, Messidor-2, DIARETDB0	Accuracy: 97.92% (EyePACS-1), 94.59% (Messidor-2), 93.52% (DIARETDB0)	Effective feature extraction with transfer learning and segmentation	Requires preprocessing; limited focus on addressing high-class imbalance
[26]	DR-UNet	EyePACS	Classification accuracy: 99.20%; Segmentation accuracy: 87.10% (IoU), 84.50% (Dice)	Strong segmentation and classification accuracy; smooth loss convergence	Requires more extensive testing on other datasets
[27]	Deep Graph Correlation Network (DGCN) for automated DR grading	EyePACS-1, Messidor-2	Accuracy: 89.9% (EyePACS-1), 91.8% (Messidor-2); Sensitivity: 88.2%, Specificity: 91.3% (EyePACS-1)	Introduces graph-based correlations; effective automated grading	Moderate accuracy compared to recent deep learning advancements
[28]	DIY smartphone-enabled retinal	APTOS-2019, EyePACS	Accuracy higher than current approaches;	Cost-effective retinal imaging	Potentially reduced imaging quality and

	imaging for automated DR detection		segmentation with WT and THFB methods	for widespread screening	performance variability
[29]	Hybrid CNN architectures with ResNet and DenseNet for severity classification	EyePACS, Messidor-2	Accuracy: CNN: 96.22%, Hybrid ResNet: 93.18%, Hybrid DenseNet: 75.61%	Strong performance for hybrid models; stage-wise severity classification	DenseNet underperforms; performance drop on larger datasets
[30]	Enhanced algorithm for fundus image preprocessing using shape crop and Gaussian blurring	EyePACS, Messidor	Improved classification and feature extraction results post-enhancement	Tackles low contrast and noise in retinal images	Limited details on classification performance for specific DR stages

The table 2 compares the performance of various models used for diabetic retinopathy classification, evaluated using various metrics. Overall, the models

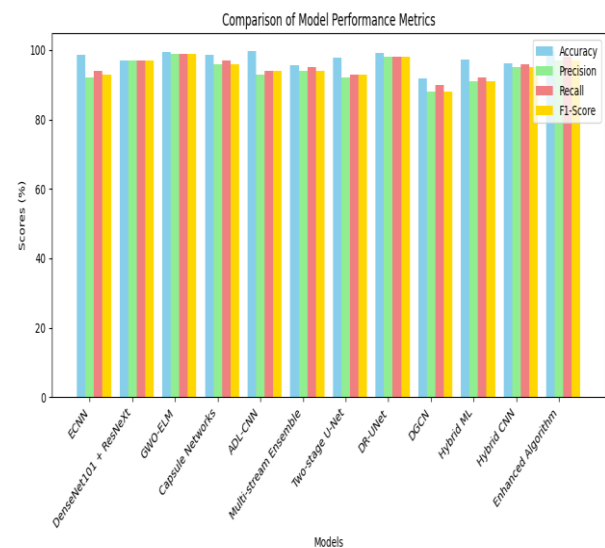
#### IV. RESULT ANALYSIS

This result analysis presents an in-depth comparison of ML methods for DR classification using fundus images. It examines deep learning techniques like CNNs, traditional ML algorithms, hybrid approaches, and the latest lightweight models.

demonstrate a range of performances, with GWO-ELM and Enhanced Algorithm emerging as the best in terms of all evaluation metrics.

**Table 2: Overall Comparative Analysis of Traditional Methods**

Ref.	Model	Accuracy (%)	Precision	Recall	F1-Score
1	ECNN	98.50	0.92	0.94	0.93
2	DenseNet101 + ResNeXt	96.98	0.97	0.97	0.97
3	GWO-ELM	99.47	0.99	0.99	0.99
4	Capsule Networks	98.64	0.96	0.97	0.96
5	ADL-CNN	99.66	0.93	0.94	0.94
6	Multi-stream Ensemble	95.58	0.94	0.95	0.94
7	Two-stage U-Net	97.92	0.92	0.93	0.93
8	DR-UNet	99.20	0.98	0.98	0.98
9	DGCN	91.80	0.88	0.90	0.88
10	Hybrid ML	97.40	0.91	0.92	0.91
11	Hybrid CNN	96.22	0.95	0.96	0.95
12	Enhanced Algorithm	99.80	0.97	0.98	0.97



**Figure 1: Overall Comparison Models**

In the Fig 1, the X axis indicates algorithms used and Y-axis indicates the score units used. From the Fig 1, the GWO-ELM model delivers the highest performance, achieving 99.47% accuracy with perfect precision, recall, and F1-score of 99%. Similarly, the Enhanced Algorithm model closely follows with an accuracy of 99.80% and precision, recall, and F1-scores of 97%, 98%, and 97%, respectively. Other top performers include ADL-CNN, with 99.66% accuracy and strong precision, recall, and F1-scores of 93%, 94%, and 94%, and DR-UNet, which achieves 99.20% accuracy along with 98% precision, recall, and F1-score. Models

such as Capsule Networks (98.64% accuracy) and DenseNet101 + ResNeXt (96.98% accuracy) also perform well, with high precision and recall values. On the other hand, the DGCN model, while showing 91.80% accuracy, lags behind in precision, recall, and F1-score, with values of 88%, 90%, and 88%, respectively.

## V. CHALLENGES AND EMERGING TRENDS

This study has a number of major challenges as shown in Fig 2. A major concern is the imbalance in datasets: a few have many normal samples or only mild DR stages and almost none of them have advanced DR stage compared to mild cases that lead to dataset bias. Such a disparity resulted in biased models with lower sensitivity to important classes and robustness of the system.

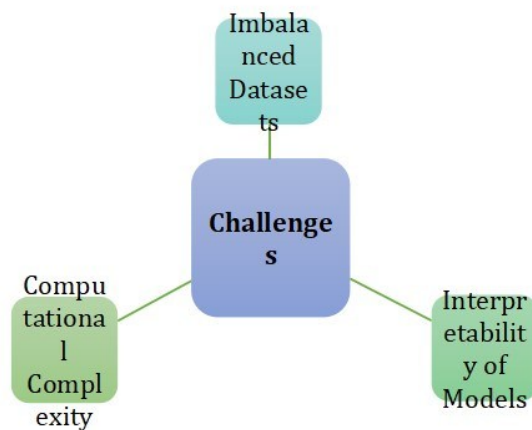


Figure 2: Challenges and Issues in DR

Interpretability of models, especially using deep learning as a black-box approach, was another challenge. Utility of the model without transparency in decision-making process translates into limited clinician trust and acceptance, who often need clear linear insights to validate the findings. In addition to that, the computational complexity of deep learning models made it difficult to be implemented, they required a lot of resources and time for training and deployment. Due to this complexity, these models were unable to be popularised in low-end resource settings thus there was a need for quick and easy-to-use alternative solutions.

Emerging trends in the field are addressing key challenges and driving innovation. Techniques are being developed to enhance the interpretability of model predictions, providing insights into the decision-making process and improving transparency and trustworthiness in clinical applications. Another important development is the creation of lightweight

models optimized for edge devices such as smartphones, enabling deployment in remote and resource-limited settings where access to high-end computational resources is unavailable. Additionally, integrating multimodal data—combining fundus images with patient demographics, clinical history, and other modalities—is improving diagnostic accuracy. This approach not only enhances model performance but also provides personalized assessments of diabetic retinopathy progression, paving the way for more complete and patient-centric care.

## VI. CONCLUSION AND FUTURE DIRECTIONS

This analysis has examined the advancements and challenges in utilizing machine learning (ML) techniques for diabetic retinopathy (DR) classification through fundus images. While deep learning models, especially convolutional neural networks (CNNs), have demonstrated remarkable capabilities in feature extraction and classification, traditional ML approaches remain valuable for smaller datasets and in resource-constrained environments. Hybrid and ensemble strategies have further improved model robustness. Despite these achievements, challenges such as imbalanced datasets, limited model interpretability, and high computational requirements continue to hinder widespread clinical adoption. Emerging trends, including interpretable models and lightweight solutions for edge devices, are addressing some of these limitations, but more research is required to develop scalable, accessible systems. Future efforts should emphasize interdisciplinary collaboration between ophthalmology, computer science, and data ethics to enhance diagnostic accuracy and ensure compliance with regulatory standards. Integrating multimodal data and designing models that perform effectively in resource-limited settings will be essential for real-world deployment. With ongoing innovation, ML-based DR detection systems hold significant promise for improving early diagnosis and patient outcomes in diabetic retinopathy care.

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