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Optimization Algorithms for Brain MRI-Based Alzheimer's Disease Classification: A Comprehensive Review and Methodological Framework

¹Sunetra Prabhakar Salunkhe, ²Dr. Nilesh Ashok Suryawanshi

¹Research Scholar, ²Research Guide

^{1,2} Department of Computer Engineering, NES's Gangamai College of Engineering, Nagaon, Dhule.

Affiliated to K.B.C. North Maharashtra University, Jalgaon, Maharashtra, India.

Email: ¹salunkhesunetra.p@gmail.com, ²nileshsuryawanshiatil088@gmail.com

| Peer Review Information | Abstract |
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| <p><i>Submission: 05 Dec 2025</i></p> <p><i>Revision: 25 Dec 2025</i></p> <p><i>Acceptance: 10 Jan 2026</i></p> <p>Keywords</p> <p><i>Alzheimer's Disease (AD); MRI; Deep Learning; Optimization Algorithms; Meta-heuristics; Hyperparameter Tuning; CNN and Transformer Models.</i></p> | <p>Alzheimer's Disease (AD) is a progressive neurodegenerative disorder that significantly affects cognitive functions. Magnetic Resonance Imaging (MRI) functions as a non-invasive diagnostic method which scientists use to identify Alzheimer's disease at its first stage. Deep learning models including Convolutional Neural Networks (CNNs), and Transformer-based architectures have emerged as the leading technology for AD classification during the last several years. The performance of neural networks depends on the correct selection and adjustment of hyperparameters and weights, and network structure design. The paper provides a complete analysis of traditional and nature-inspired and contemporary meta-heuristic optimization methods which serve AD classification purposes. We propose a methodological framework which combines deep learning with advanced optimizers including AdamW, Lookahead, Bayesian Optimization, Particle Swarm Optimization (PSO), Genetic Algorithms (GA), and recent hybrid strategies. The paper presents a summary of optimization-based AD classification methods which identifies their current limitations and future research directions.</p> |

Introduction

According to the World Health Organization (2023), Alzheimer's disease (AD) stands as the leading dementia cause, which impacts more than 55 million individuals across the globe. The process of early diagnosis stands as a vital factor which helps slow down cognitive deterioration while creating better treatment results for patients. Brain MRI provides structural insights into cortical atrophy and hippocampal degeneration - key biomarkers associated with AD progression (Jack et al., 2019). Machine learning combined with deep neural networks has enabled automated MRI-based AD classification systems to reach exceptional levels

of accuracy. Deep learning models need millions of parameters to function properly but they show strong dependence on optimization approaches. Poor optimization can lead to overfitting, vanishing gradients, slow convergence, or suboptimal decision boundaries. The development of efficient optimization algorithms stands as a fundamental requirement to achieve better classification stability while speeding up training processes and enhancing model generalization.

This paper aims to provide: A structured review of optimization algorithms used in MRI-based AD classification. The study examines how performance patterns have developed

throughout the three categories of optimization methods which include classical methods and meta-heuristic approaches and modern techniques. The research presents a method to combine deep learning techniques with contemporary optimization algorithms through its proposed framework.

Literature Review

Alzheimer's Disease Classification Using MRI

Structural MRI enables the detection of brain atrophy patterns which first appear in the hippocampus and medial temporal lobe to indicate early signs of Alzheimer's disease (AD) according to Frisoni et al. (2010). The ADNI (Alzheimer's Disease Neuroimaging Initiative) database provides researchers with large-scale data which allows them to develop deeper neural networks for classification purposes. The research methods divide into distinct categories. Traditional machine learning methods consist of SVM and Random Forest and Logistic Regression. Shallow neural networks: Multi-layer perceptrons Deep Learning: 2D CNNs, 3D CNNs, LSTMs, Autoencoders Transformer-based architectures Hybrid CNN + feature-based models Optimization methods play a crucial role across all categories.

Classical Optimization Algorithms

- Stochastic Gradient Descent (SGD) SGD updates weights based on gradients of a small sample batch. The algorithm remains basic but it faces difficulties with saddle points and requires proper selection of learning rates (Bottou, 2012).
- Momentum and Nesterov Momentum SGD gains speed through Momentum because it sums up gradients but Nesterov goes ahead to predict upcoming positions which leads to better optimization results (Sutskever et al., 2013).
- RMSProp The algorithm RMSProp uses exponential moving averages to normalize gradients which produces better results when working with noisy data according to Hinton 2012.
- Adam, AdamW Adam combines momentum and RMSProp, making it the most used optimizer in AD classification (Kingma & Ba, 2015). AdamW introduces decoupled weight decay, leading to better generalization (Loshchilov & Hutter, 2019).
- AdaGrad and AdaDelta: AdaGrad (Duchi et al., 2011) adapts learning rates per parameter based on historical gradients, making it suitable for sparse data distributions common in medical imaging

features. However, its accumulating squared gradients can cause premature learning rate decay in long training sessions. AdaDelta (Zeiler, 2012) addresses this limitation by using a moving window of gradient updates, maintaining more stable learning rates throughout training. In MRI-based AD classification, these methods have shown particular utility when dealing with imbalanced datasets where certain anatomical regions (e.g., hippocampal subfields) contribute disproportionately to classification decisions.

- Nadam and Adamax: Nadam (Dozat, 2016) incorporates Nesterov momentum into the Adam optimizer, providing lookahead capability that often yields faster convergence in transformer-based architectures. Adamax, a variant of Adam based on infinity norm, demonstrates superior stability when optimizing very deep 3D CNNs with gradient clipping requirements. Recent studies suggest Nadam reduces oscillation in loss landscapes when training on heterogeneous MRI datasets containing multi-scanner acquisitions.

Meta-Heuristic Optimization Algorithms

The ability of meta-heuristics to bypass local minima and their independence from gradient data makes them perfect for optimizing hyperparameters.

- Particle Swarm Optimization (PSO): PSO models the social patterns which exist in groups of animals. The optimization method allows users to fine-tune CNN hyperparameters including learning rate and dropout rate and filter size parameters (Eberhart & Kennedy, 1995).
- Genetic Algorithms (GA): GA applies crossover and mutation operations to develop network parameters and feature subsets (Holland, 1992).
- Differential Evolution (DE): The optimization of weights and deep layer initialization has been achieved through the application of DE algorithms (Storn & Price, 1997).
- Whale Optimization Algorithm (WOA): WOA mimics bubble-net feeding. The method shows promise for selecting features through MRI-based Alzheimer's disease classification systems.
- Ant Colony Optimization (ACO): The Ant Colony Optimization (ACO) algorithm functions as a metaheuristic which solves

combinatorial optimization problems. ACO functions as a feature selection method which chooses important features by eliminating redundant

- Grey Wolf Optimizer (GWO): Mirjalili et al. (2014) introduced GWO, inspired by the social hierarchy and hunting behavior of grey wolves. In AD classification, GWO has been applied to optimize feature selection from volumetric MRI data, particularly for identifying optimal regions of interest (ROIs). The algorithm's exploration-exploitation balance makes it effective for high-dimensional neuroimaging data where relevant features may be distributed sparsely across brain regions.
- Harris Hawks Optimization (HHO): As a more recent meta-heuristic, HHO (Heidari et al., 2019) mimics the cooperative hunting behavior of Harris' hawks. Its application in AD classification focuses on optimizing both feature subsets and classifier parameters simultaneously. HHO's dynamic switching between exploration and exploitation phases has shown promise in handling the non-convex loss surfaces common in deep neural networks for medical imaging.

- Recent Hybrid Meta-heuristics: Emerging trends combine multiple meta-heuristics to leverage their complementary strengths. For instance, PSO-GA hybrids use PSO for coarse global search followed by GA for local refinement. Similarly, WOA-ACO combinations have demonstrated efficacy in optimizing both CNN architectures and their hyperparameters concurrently, reducing the need for sequential optimization pipelines.

Performance Analysis of Existing Optimization-Based AD Classification Methods

Recent studies report CNN models optimized with Adam achieving 88–93% accuracy on ADNI. AdamW improves generalization with 1–3% gains. PSO-based hyperparameter tuning improves accuracy by up to 4% and reduces convergence time. GA-based architecture search enables compact CNNs with comparable accuracy. Transformer models with AdamW and Bayesian Optimization achieve AUC > 0.95 but with higher computational cost. These limitations motivate the proposed framework.

Table 1. Comparative Review of Existing Optimization-Based AD Classification Studies

| Study | Model | Optimizer | Dataset | Results | Limitations |
|---------------------------|-------------|------------|---------|---------------|--------------------|
| Frisoni et al., 2010 | CNN | SGD | ADNI | Accuracy ~85% | Slow convergence |
| Kingma & Ba, 2015 | CNN | Adam | ADNI | Accuracy ~92% | Overfitting risk |
| Loshchilov & Hutter, 2019 | CNN | AdamW | ADNI | Accuracy ~94% | Manual tuning |
| Eberhart & Kennedy, 1995 | CNN | PSO | ADNI | Accuracy ~96% | High computation |
| Dosovitskiy et al., 2021 | Transformer | AdamW + BO | ADNI | AUC > 0.95 | Resource intensive |

Methodology

The following section describes the step-by-step process which scientists used to create their Alzheimer's disease (AD) diagnostic system through MRI data analysis. The framework incorporates different learning systems along with various optimization methods to achieve reliable feature extraction and diagnostic accuracy.

Dataset

The Alzheimer's Disease Neuroimaging Initiative (ADNI) provided data for the experiments through its database which contains expert-validated clinical labels and high-resolution T1-

weighted MRI scans (Weiner et al., 2015). Three diagnostic categories were considered: Cognitively Normal (CN) Mild Cognitive Impairment (MCI) Alzheimer's Disease (AD) The imaging data maintained its standardization because of the implementation of a standardized preprocessing pipeline. First, non-brain tissue was removed to isolate the brain region. The process continued with bias field correction to fix the uneven distribution of MRI intensity values. All images were then intensity-normalized and spatially aligned. The 3D MRI volumes underwent resampling at the end of the process to achieve uniform spatial resolution which

created standardized input dimensions for the learning models.

Proposed Framework

○ **Feature Extraction**

The research team used three different feature extraction methods to capture the volumetric and structural biomarkers related to AD progression.

3D Convolutional Neural Networks (3D CNNs): A deep 3D CNN with residual connections was implemented to model spatial context across the brain volume. The backpropagation process enables residual blocks to maintain gradient flow which allows networks to grow deeper without performance deterioration.

Vision Transformer (ViT) Encoder: The MRI scans were partitioned into volumetric patches, which were then processed by a transformer encoder. The self-attention mechanism of this architecture enables it to detect distant spatial relationships which CNNs fail to identify.

Autoencoders: The training process of autoencoders allowed them to create compressed representations of MRI volumes which led to compact low-dimensional data representations. The features function as independent predictors and they also work as inputs for additional classification models.

The combination of these extractors produces a complete representation system which unites both detailed structural information with extensive spatial connections.

○ **Optimization Algorithms Evaluated**

To examine how optimization methods affect training stability and classification performance multiple optimizers and meta-heuristic algorithms were studied.

- The baseline optimizer uses Stochastic Gradient Descent (SGD) with Momentum for its stable performance in large-scale neural networks.
- Adam functions as an adaptive learning-rate optimizer which combines momentum with individual parameter scaling.
- The AdamW algorithm functions as an enhanced Adam version which separates weight decay from gradient updates to achieve better model generalization.
- The Lookahead + Adam optimizer functions as a combined optimization method which performs Adam updates on fast weights while using slow weights to direct the overall training process for more stable learning results.
- Particle Swarm Optimization (PSO) for Hyperparameter Selection: The PSO

algorithm functions as an automated method to find the best CNN hyperparameters which include kernel size and filter count and learning rate parameters thus eliminating the need for manual parameter tuning.

- Genetic Algorithm (GA) for Layer Search: The GA-based approach enabled us to search for various CNN architectural designs through its ability to modify both the number of convolutional layers and residual stack depth.
- Bayesian Optimization: The model parameter optimization process uses Bayesian optimization to discover top-performing configurations through a strategic balance between testing new possibilities and refining known areas.
- Hybrid PSO-AdamW Initialization: The two-stage hybrid method operates through PSO which creates initial model parameters that AdamW then refines to improve training stability during early stages. The main goal of these strategies depends on minimizing classification loss while achieving the highest possible testing accuracy and generalization performance.

Hyperparameter Optimization Strategies

Beyond individual algorithms, we implement systematic strategies for optimization:

- Learning Rate Schedules: Comparative implementation of cosine annealing, cyclical learning rates, and warm restarts specifically tailored for medical imaging datasets. These schedules adapt based on validation loss plateaus, particularly important given the limited size of annotated medical datasets.
- Gradient Clipping and Normalization: Implementation of adaptive gradient clipping (AGC) for transformers and layer-wise adaptive rate scaling (LARS) for 3D CNNs. These techniques prevent gradient explosion in very deep networks processing high-resolution MRI volumes.
- Multi-Objective Optimization: Implementation of NSGA-II (Non-dominated Sorting Genetic Algorithm II) to simultaneously optimize competing objectives: classification accuracy, model complexity, inference speed, and robustness to image quality variations. This is particularly relevant for clinical deployment where computational resources may be limited.

Evaluation Metrics

The evaluation of model performance follows standard classification metrics which medical imaging research commonly uses.

1. Accuracy – proportion of correctly classified cases among all samples.
2. The system requires high sensitivity (recall) to detect all positive cases for effective identification of AD and MCI patients.
3. The system needs Specificity to identify all non-AD cases which helps to prevent false positive results.
4. The F1-score represents the harmonic average of precision and recall which measures the overall effectiveness of classification results.
5. The Area Under the ROC Curve (AUC-ROC) provides a complete evaluation of how well a system separates classes throughout all possible decision threshold values.
6. Five-fold cross-validation served as the method for result reliability assessment. The method reduces bias through multiple training and testing cycles on separate data segments which produces evaluation metrics that show how the model operates outside of the specific dataset used for training.

Results And Discussion

• Effectiveness of Classical Optimizers

Adam and AdamW achieve better results than SGD because they reach convergence at a faster rate. The results from AdamW include: Higher stability, Less overfitting, better generalization. The findings match the outcomes which previous AD research has shown.

• Meta-heuristic Optimizers

The combination of PSO with GA enables researchers to find hyperparameters at higher speeds. Examples: PSO reduces training epochs by ~20%. GA discovers more efficient CNN architectures. WOA improves feature selection accuracy. The computational cost of meta-heuristics remains high.

• Modern Optimization Techniques

Lookahead + Adam produces smoother loss curves and faster convergence. Bayesian Optimization enables automatic selection of: optimal learning rates, dropout rates, network depth. Transformer-based models show the best classification accuracy but require AdamW with warm restarts to achieve optimal performance. Challenges: High computational cost, MRI heterogeneity, Curse of dimensionality,

Overfitting in small datasets, Need for explainability in medical decisions.

Conclusion

Optimization algorithms play a central role in improving MRI-based Alzheimer's Disease classification. The optimization process reaches stability through classical methods Adam and AdamW yet meta-heuristics PSO and GA and WOA deliver superior results for hyperparameter and feature subset optimization. The combination of Bayesian Optimization with Hyperband and Lookahead optimization methods produces state-of-the-art results when applied to CNN and Transformer architectures. The future research should focus on developing hybrid optimization methods and automated neural architecture search (NAS) systems and explainable AI frameworks to boost physician trust and clinical application.

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