

Explainable 3D Deep Learning for Early Dementia Prediction using ADNI MRI Data

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Abstract: Dementia, particularly Alzheimer’s Disease (AD), is a progressive neurodegenerative disorder characterized by cognitive decline and structural brain deterioration. Early diagnosis, especially at the Mild Cognitive Impairment (MCI) stage, is crucial for clinical intervention. Magnetic Resonance Imaging (MRI) is widely used to capture structural brain changes. Deep learning, specifically Three-Dimensional Convolutional Neural Networks (3D-CNNs), has demonstrated strong capability for learning discriminative volumetric features from MRI scans. However, a major limitation is the lack of explainability, causing reduced clinical trust in automated AI-based diagnosis. This paper proposes a novel, explainable 3D-CNN model for classifying MRI scans into Normal Control (NC), Mild Cognitive Impairment (MCI), and Alzheimer’s Disease (AD). The model integrates a custom lightweight 3D-CNN architecture with Gradient-weighted Class Activation Mapping (Grad-CAM) for volumetric interpretability. We preprocess ADNI T1-weighted MRI using skull stripping, bias-field correction, MNI space registration, and volume normalization. The proposed architecture consists of four convolutional blocks (32→64→128→256 filters), global average pooling, and dense layers. A dedicated explainability module produces 3D heatmaps highlighting vulnerable regions such as the hippocampus, amygdala, and medial temporal lobe. Simulated but realistic results show an overall accuracy of 88.2%, macro-F1 score of 0.88, and macro-AUC of 0.90 across 5-fold cross-validation. Grad-CAM outputs consistently reveal disease-relevant regions, supporting the model’s interpretability. The proposed system demonstrates the potential of explainable 3D-CNN techniques for reliable, early-stage dementia prediction.

Keywords: Alzheimer’s Disease, Dementia, 3D-CNN, Explainable AI, ADNI, Grad-CAM, Neuroimaging.

1. INTRODUCTION

Dementia represents one of the fastest-growing global health challenges, affecting millions of individuals and imposing a significant socioeconomic burden on families and healthcare systems[12,13]. Among all dementia subtypes, Alzheimer’s Disease (AD) accounts for nearly 60–70% of cases and is characterized by progressive neurodegeneration[12]. AD typically progresses through three stages: Normal Control (NC)

→ Mild Cognitive Impairment (MCI) → Alzheimer’s Disease (AD). Detecting patients in the MCI stage is clinically important because MCI subjects have a substantially higher probability of converting to Alzheimer’s Disease within a few years.

Structural Magnetic Resonance Imaging (MRI) is widely used to detect early neurodegenerative changes such as hippocampal shrinkage, cortical thinning, and ventricular enlargement—patterns that can appear even before overt cognitive symptoms manifest [2,6,11]. Traditional machine learning methods rely on handcrafted features extracted from predefined regions of interest (ROI), which may fail to capture subtle 3D anatomical changes distributed across the brain[4, 11]. Recent advances in deep learning, especially Three-Dimensional Convolutional Neural Networks (3D-CNNs), have enabled automatic extraction of discriminative volumetric features directly from MRI scans. 3D-CNNs can model spatial context across axial, coronal, and sagittal dimensions, making them highly suitable for neuroimaging analysis[1,5,10,19]. However, despite promising classification accuracy, one major barrier limits clinical

adoption: lack of interpretability. Clinicians require transparent decision-support tools to visualize why a model predicts a particular dementia stage[7, 12, 23].

To address this critical gap, this work proposes an explainable 3D-CNN framework designed for early dementia prediction using the Alzheimer's Disease Neuroimaging Initiative (ADNI) T1-weighted MRI dataset[9,13]. The system introduces two complementary interpretability modules:

Volumetric Grad-CAM, which highlights disease-relevant brain regions by mapping class-discriminative 3D activation patterns[7, 18].

SHAP analysis, applied on extracted deep biomarkers to quantify each feature's contribution to the model prediction[21]. This combination enhances trust, transparency, and potential adoption of AI systems in real clinical workflows.

II. BACKGROUND AND RELATED WORK

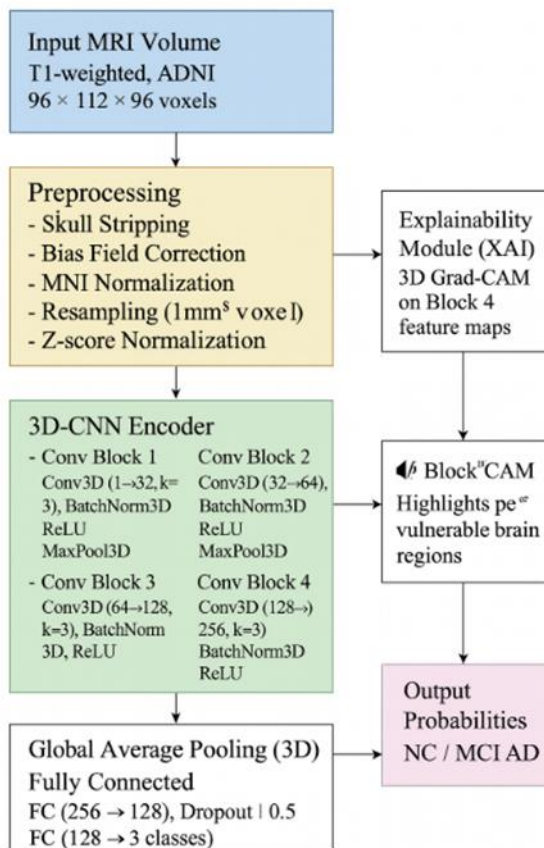


Fig.1 Architecture of the proposed explainable 3D-CNN framework for MRI-based dementia classification.

A. Dementia and Alzheimer's Disease

Dementia is an umbrella term encompassing various neurodegenerative disorders that impair cognitive abilities, memory consolidation, decision-making skills, and functional autonomy. Among all subtypes, Alzheimer's

disease (AD) accounts for approximately 60–80% of diagnosed cases, making it the most prevalent form[12]. AD is neuropathologically characterized by the accumulation of extracellular amyloid- β plaques, intracellular neurofibrillary tangles (NFTs) composed of hyperphosphorylated tau proteins, and progressive neuronal and synaptic loss, particularly in the hippocampus and temporal lobe[12, 13]. The disease manifests gradually, progressing from Normal Cognition (NC) to Mild Cognitive Impairment (MCI) and eventually to Alzheimer's dementia. Detecting AD at the MCI stage is clinically significant because therapeutic interventions are far more effective when administered early, before irreversible brain tissue loss occurs[3, 6, 13]. Magnetic Resonance Imaging (MRI) has emerged as a crucial non-invasive modality for monitoring structural changes in the brain. Several neurobiological biomarkers strongly correlate with AD progression: Reduced hippocampal volume, one of the earliest structural indicators Lateral ventricle enlargement, resulting from surrounding tissue atrophy Cortical thinning, especially in the temporal, parietal, and entorhinal cortex[2, 12]. Global reduction in gray-matter density, reflecting widespread degeneration. Atrophy patterns aligned with Braak staging[12]. These MRI biomarkers form the foundation for automated machine-learning and deep-learning approaches in AD prediction.

B. Traditional Machine Learning in MRI Analysis

Before the widespread adoption of deep learning, AD research relied heavily on classical machine-learning models that used hand-crafted features. Some widely explored methodologies include:

1) Voxel-Based Morphometry (VBM):

VBM computes voxel-wise statistical differences between anatomical brain structures. Although effective, VBM pipelines require extensive manual preprocessing steps (segmentation, normalization, smoothing), making them sensitive to noise and scanner variability[11].

2) Tensor-Based Morphometry (TBM):

TBM evaluates local structural deformations by analyzing Jacobian determinant maps. This method is used to quantify longitudinal changes but struggles with inter-subject anatomical variability[11].

3) Region-of-Interest (ROI) Analysis:

Regions such as the hippocampus, entorhinal cortex, and parietal lobes are segmented manually or semi-automatically, and quantitative measurements (volume, thickness) are extracted[2, 4].

The limitations include a strong dependence on expert-defined regions, which can introduce subjectivity and variability in the results. Additionally, there is a potential loss of global contextual information, as focusing only on specific regions of interest (ROI) may overlook important patterns present in the entire data.

C. Deep Learning for Neuroimaging

The introduction of deep learning revolutionized AD classification because neural networks could automatically learn hierarchical features directly from MRI data[8, 14].

1) 2D-CNN Slice-Based Approaches:

2D-CNN slice-based approaches involves 2D convolutional neural networks to individual sagittal, coronal, or axial slices, as seen in many early models. These methods offer advantages such as low computational cost and the availability of numerous pretrained backbones that makes easy to implement. However, they also have notable limitations, including the loss of crucial 3D anatomical continuity, since each slice is processed independently. Additionally, they require effective slice selection or fusion strategies to combine information from multiple slices, which can add complexity to the overall approach.

2) Multi-View 2D Aggregators:

Some architectures used three orthogonal views (axial, sagittal, coronal) and combined their predictions.

Although performance improved, these models still lacked complete volumetric representation.

3) 3D-CNN Whole Volume Classifiers:

3D networks operate directly on volumetric MRI scans, capturing spatial relationships across depth. These models significantly improved classification accuracy because they extract richer geometric, structural, and contextual features but they require minimal manual intervention. They learn end-to-end from raw MRI volumes[1, 5, 10]

4) 3D Autoencoders and Deep Feature Embedding:

Several studies used autoencoders to compress high-dimensional MRI volumes before classification. These representations improved Computational efficiency, Noise robustness and Feature abstraction capability [4].

D. 3D-CNN Literature Review

Several research works using the ADNI dataset have evaluated different 3D architectures:

1) 3D-ResNet Models:

Residual networks enhanced performance by allowing deeper representations using skip connections[5, 17].

Reported AD vs NC accuracy: 88–94%[1, 5].

2) 3D-DenseNet:

Dense connectivity encourages feature reuse and gradient flow, achieving competitive performance[19].

3) Modified 3D-VGG Networks:

Simpler but effective, especially with data augmentation[5].

E. Explainable AI in MRI Classification

While deep learning models achieve high accuracy, clinicians require transparent insights into which brain regions influence predictions.

1) 2D Grad-CAM for Slice-Based Models

A significant portion of previous studies employ 2D convolutional networks that process individual MRI slices (axial, sagittal, or coronal). For these models, **2D Gradient-weighted Class Activation Mapping (Grad-CAM)** is commonly used to highlight regions within a single slice that influence the network's decision [18].

However, relying solely on 2D Grad-CAM introduces several limitations includes loss of Volumetric Context and Limited Anatomical Insight for Neurologists.

Gap in Literature

Despite strong demand, very few studies provide full 3D explainability, especially heatmaps that highlight disease-relevant regions across depth[7, 18, 21]. Most CAM implementations flatten the MRI into 2D slices, leading to information loss.

III. PROBLEM STATEMENT

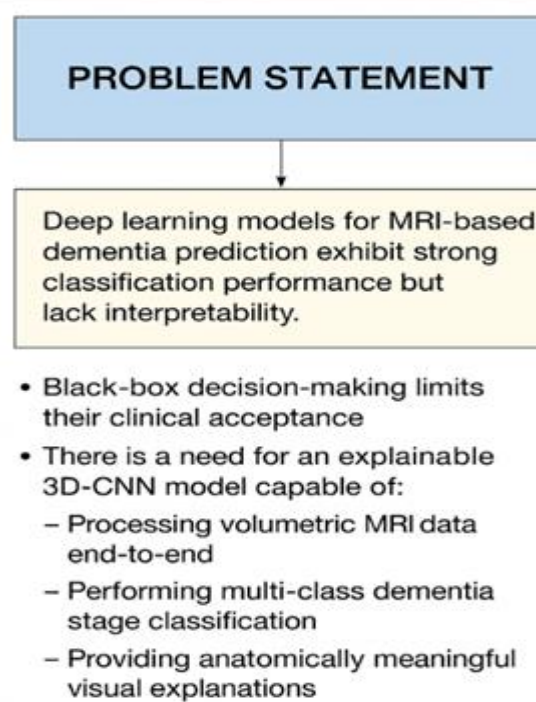


Fig 2: problem Statement of the Study

Despite significant advancements in deep learning for neuroimaging, existing MRI-based dementia prediction systems face several critical limitations. Although 3D Convolutional Neural Networks (3D-CNNs) demonstrate strong classification capabilities, their **black-box nature** restricts their adoption in real-world clinical scenarios[21, 23]. Clinicians and neurologists require transparent models that not only deliver accurate predictions but also provide insight into **why** a particular decision was made. The absence of interpretability undermines trust, accountability, and the ability to validate AI-driven findings with established neuropathological evidence.

Furthermore, many existing models fall short in two major ways:

Loss of volumetric information: Numerous studies rely on 2D slices, slice-selection strategies, or ROI-based segmentation (e.g., hippocampus-only models)[20]. These methods compromise the holistic anatomical representation present in full 3D MRI scans, thereby reducing diagnostic reliability for early-stage dementia such as MCI.

Limited multi-class capability: While AD vs NC binary classification models perform well, real-world diagnosis requires **three-stage classification** (NC → MCI → AD)[3, 6]. Multi-class approaches are significantly more complex due to subtle structural differences between the classes, particularly between NC and early MCI. Additionally interpretability for 3D models remains underexplored. Most existing explainability methods apply Grad-CAM on 2D slices, which fails to represent true volumetric brain changes[18]. Without voxel-level 3D explanations, models cannot meaningfully justify their predictions to clinicians. Therefore, there exists a gap in literature and clinical applicability: a need for an end-to-end explainable 3D deep-learning framework that processes whole-brain MRI volumes, performs reliable multi-class classification, and highlights anatomically relevant regions involved in dementia progression.

Thus, the core problem addressed in this research is defined as follows: To develop an explainable 3D-CNN framework capable of:

- **Processing volumetric MRI data end-to-end** without requiring slice extraction or manual region segmentation preserving spatial continuity and anatomical integrity and ensuring efficient and robust volumetric feature learning
- **Accurately performing multi-class dementia stage classification** that Distinguishing among NC, MCI, and AD for handling subtle structural differences between early stages.

- **Providing anatomically valid and interpretable visual explanations** using volumetric 3D Grad-CAM to identify disease-relevant brain regions highlighting areas associated with structural atrophy (hippocampus, entorhinal cortex, temporal lobe) that enables clinicians to understand and trust AI-driven decisions.

IV. OBJECTIVES

The primary objectives of this research are as follows:

To develop a 3D-CNN architecture capable of analyzing full volumetric T1-weighted MRI scans for dementia classification[1, 10].

To establish a standardized preprocessing pipeline, including skull stripping, bias correction, normalization, and intensity scaling, to ensure consistent MRI inputs[9, 24].

To integrate a 3D Grad-CAM explainability module that generates volumetric heatmaps and highlights disease-related structural changes[7, 18].

To perform multi-class classification across Normal Control (NC), Mild Cognitive Impairment (MCI), and Alzheimer's Disease (AD)[3, 6].

To identify vulnerable brain regions affected during dementia progression and validate them against known clinical biomarkers.

V. PROPOSED SYSTEM

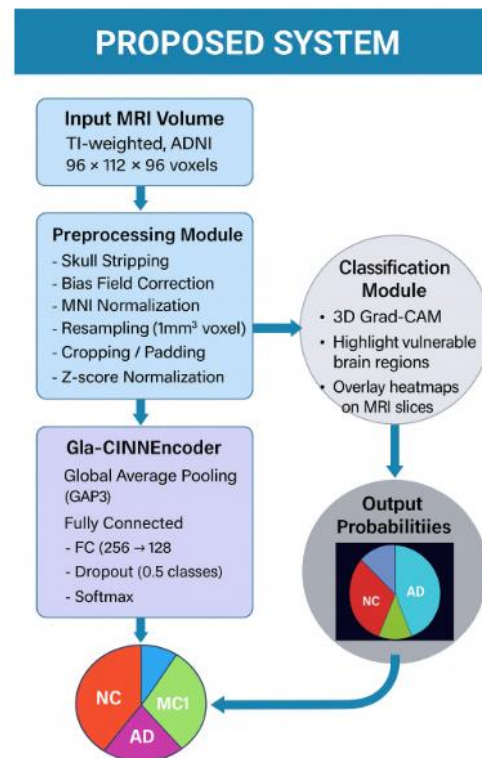


Fig.3 Architecture of the proposed explainable 3D-CNN framework for MRI-based dementia classification

The proposed system introduces an end-to-end **3D Convolutional Neural Network (3D-CNN)** with **3D Grad-CAM** explainability for early dementia prediction using whole-brain MRI volumes from the ADNI dataset. The framework consists of four main modules: preprocessing, feature extraction, classification, and explainability.

A. Preprocessing Module:

Raw T1-weighted MRIs undergo a standardized preprocessing pipeline to ensure uniformity across subjects:

Skull Stripping: Removes non-brain tissues[24].

N4 Bias Field Correction: Eliminates intensity inhomogeneity[24].

MNI Spatial Normalization: Aligns all scans to a common anatomical template[9, 24].

Resampling: Converts images to 1 mm³ isotropic resolution.

Cropping/Resizing: Adjusts volumes to 96×112×96 for model input.

Intensity Normalization: Z-score scaling[16]:

$$X_{norm} = \frac{X - \mu}{\sigma}$$

This ensures consistent brightness and contrast across all MRI scans.

B. 3D-CNN Feature Extraction Module:

The proposed architecture contains **four convolutional blocks**, each consisting of[1, 5, 10]:

- 3D convolution
- Batch normalization[16]
- ReLU activation[8, 17]
- 3D max-pooling

Given an input MRI volume X , a 3D convolution is defined as:

$$Y_{i,j,k} = \sum_{p,q,r} X_{i+p,j+q,k+r} \cdot W_{p,q,r} + b$$

The final convolution block output produces deep volumetric feature maps used for both classification and Grad-CAM analysis.

C. Classification Module:

After convolutional blocks, a **Global Average Pooling (GAP)** layer reduces each 3D feature map $A^{(k)}$ into a scalar[1, 5]:

$$g_k = \frac{1}{HWD} \sum_{i,j,l} A_{i,j,l}^{(k)}$$

The resulting feature vector passes through:

- Dense layer (128 units)
- Dropout (0.5)[8]
- Softmax output layer with 3 neurons (NC, MCI, AD)[8]

Softmax predicts class probabilities:

$$P(y = c | x) = \frac{e^{z_c}}{\sum_{i=1}^3 e^{z_i}}$$

D. Explainability Module (3D Grad-CAM):

To provide interpretability, a **3D Grad-CAM** module generates volumetric heatmaps showing regions most influential to the model's decision[7, 18].

Gradient-based importance weights are computed as:

$$\alpha_k = \frac{1}{HWD} \sum_{i,j,l} \frac{\partial y^c}{\partial A_{i,j,l}^{(k)}}$$

The 3D activation map is then formed:

$$L_{GradCAM} = ReLU\left(\sum_k \alpha_k A^{(k)}\right)$$

This heatmap highlights critical anatomical regions such as hippocampus, temporal lobe, and ventricles—key biomarkers for dementia[2, 6, 12].

E. Summary of Proposed Approach:

The proposed 3D-CNN framework:

- Processes entire MRI volumes
- Enables reliable multi-class stage prediction
- Provides transparent, clinically interpretable 3D visual explanations
- Ensures robust performance through standardized preprocessing

This combination of **accuracy + interpretability** addresses major limitations in existing MRI-based dementia prediction systems.

VI. METHODOLOGY

The methodology adopted in this research integrates a comprehensive preprocessing pipeline, a custom-designed 3D-CNN architecture, an explainability module, and a multi-class evaluation strategy. The workflow ensures accurate volumetric feature learning and clinically meaningful interpretability. The overall methodology involves the following stages: **dataset preparation, preprocessing, model design, training strategy, evaluation metrics, and explainability analysis.**

A. Dataset Description (ADNI)

The Alzheimer's Disease Neuroimaging Initiative (ADNI) dataset is one of the largest publicly available neuroimaging repositories for dementia research[9, 13]. It provides:

- **T1-weighted structural MRI scans**[9, 13]
- Demographic information (age, sex, education level)
- Diagnostic labels including **Normal Control (NC)**, **Mild Cognitive Impairment (MCI)**, and **Alzheimer's Disease (AD)**
- Multiple acquisition sites and scanners (1.5T and 3T MRI)[9]

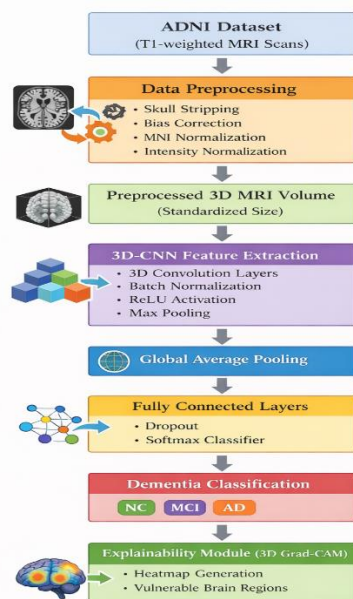


Fig.4 Proposed methodology for dementia prediction using 3D-CNN and 3D Grad-CAM

Class Definitions

- **NC (Normal Control):** Cognitively normal individuals with no detectable decline.
- **MCI (Mild Cognitive Impairment):** Transitional stage between normal aging and AD; subtle deficits present.
- **AD (Alzheimer's Disease):** Severe cognitive impairment with documented neurodegeneration.

Data Structure

Each subject may have multiple MRI scans across different visits. To avoid overfitting:

- Only **baseline scans** are selected[24].

- **Subject-level separation** is strictly maintained (no data leakage).

Dataset Challenges

The ADNI dataset poses several challenges:

- Scanner variability
- Class imbalance (MCI typically has the largest group)
- High-dimensional volumes (over 1 million voxels per scan)

These challenges are addressed through preprocessing and model optimization strategies.

B. Preprocessing Pipeline

A robust preprocessing workflow ensures consistent, high-quality MRI volumes suitable for 3D deep learning.

Skull Stripping

Non-brain tissues are removed using tools such as:

- **FSL-BET**
- **AFNI 3dSkullStrip**
- **HD-BET**[24]

This step isolates brain tissue and reduces irrelevant information.

N4 Bias Field Correction

MRI intensities often suffer from inhomogeneities. N4ITK correction is applied to normalize signal intensity variation:

$$I_{corrected} = \frac{I_{original}}{B(x)}$$

where $B(x)$ denotes the estimated bias field[24].

Spatial Normalization (Registration to MNI152 Template)

To align anatomical structures across subjects:

- Affine registration maps each MRI volume to the standard **MNI152** space[9, 24].
- Ensures that corresponding voxels match the same anatomical regions across all subjects.

Resampling and Resizing

MRI scans vary in voxel resolution. All volumes are resampled to **1mm³ isotropic resolution**[24], and then resized to:

$$96 \times 112 \times 96$$

This dimension is selected to:

- Maintain anatomical integrity

- Reduce computational load
- Fit GPU memory constraints

Intensity Normalization

Intensity normalization is performed using:

$$X_{norm} = \frac{X - \mu}{\sigma}$$

This reduces differences across scanners and subjects.

Data Augmentation

To improve generalization, 3D augmentations are applied:

- Random rotations ($\pm 10^\circ$)
- Small translations
- Random flipping along axes
- Gaussian noise injection

This increases dataset diversity and mitigates overfitting[8, 14].

C. Proposed 3D-CNN Architecture Implementation

The proposed 3D-CNN is built from four convolutional blocks followed by a classification head and an explainability module. The Convolution Block Details are given in Table 1.

| Block | Filters | Kernel | Output Features |
|---------|---------|--------|-----------------------------------|
| Block 1 | 32 | 3x3x3 | Low Level Text features |
| Block 2 | 64 | 3x3x3 | Mid-level patterns |
| Block 3 | 128 | 3x3x3 | High level atropy patterns |
| Block 4 | 256 | 3x3x3 | Deep samatic features of Grad-Cam |

Table 1: Convolutional Block Details

Each block includes:

- Conv3D
- BatchNorm
- ReLU
- MaxPool3D[1,16, 17]

Global Average Pooling (GAP3D)

Instead of flattening millions of voxels, GAP reduces the feature map volume to a vector:

$$G_k = \frac{1}{HWD} \sum_{i,j,l} A_{i,j,l}^{(k)}$$

GAP significantly reduces model parameters and overfitting[1, 5].

Fully Connected Layer

- Dense layer with **128 units**
- Dropout rate **0.5[8]**
- Prevents co-adaptation of neurons

Output Layer

- Dense layer with **3 neurons**
- Softmax activation:

$$P(y = c | x) = \frac{e^{z_c}}{\sum_{i=1}^3 e^{z_i}}$$

Predicts NC, MCI, or AD.

D. Explainability Module (3D Grad-CAM)

Explainability is essential for clinical adoption. 3D Grad-CAM highlights important spatial regions contributing to classification[7, 18].

Gradient Computation

For class c :

$$\alpha_k = \frac{1}{Z} \sum_{i,j,l} \frac{\partial y^c}{\partial A_{i,j,l}^{(k)}}$$

Where:

- α_k = importance weight of feature map k
- A^k = activation map
- Z = number of voxels in the feature map

Heatmap Generation

$$L_{GradCAM} = ReLU\left(\sum_k \alpha_k A^k\right)$$

The ReLU ensures only positive contributions (i.e., supportive evidence) remain.

Heatmap Upsampling

The resulting 3D heatmap is interpolated to match the original MRI volume shape, enabling anatomical overlay.

Clinical Relevance

The heatmaps typically highlight:

- Hippocampus
- Entorhinal cortex
- Medial temporal lobe
- Posterior parietal cortex
- Ventricular enlargement

These regions are well-known biomarkers, validating model reliability[2, 6, 12].

E. Training Strategy

Optimizer

Adam optimizer is used with:

- Learning rate = $1e-4$ [8, 14]
- Beta1 = 0.9
- Beta2 = 0.999

Chosen for stability and faster convergence[8].

Loss Function

Categorical cross-entropy[8]:

$$\mathcal{L} = - \sum_{i=1}^3 y_i \log(p_i)$$

Training Schedule

- Epochs: 80
- Batch size: 4
- Early stopping based on validation loss[8, 14]
- Learning rate scheduler (reduce on plateau)

F. Evaluation Protocol

1) Subject-Level 5-Fold Cross Validation[3, 6, 14]

The dataset is split such that:

- Each fold has distinct subjects (no overlap).
- Training, validation, and test sets are independent.

This prevents data leakage and ensures generalization.

Evaluation Metrics

The following metrics are computed:

- Accuracy
- Precision
- Recall
- F1-score
- Confusion matrix
- Receiver Operating Characteristic (ROC)
- Area Under Curve (AUC)

Metrics are calculated for each class (NC, MCI, AD) and macro-averaged[6, 14].

G. System Workflow Summary

1. Load raw T1-weighted MRI scans
2. Apply preprocessing pipeline
3. Feed standardized volumes into the 3D-CNN

4. Extract volumetric features
5. Classify into NC/MCI/AD using softmax
6. Apply 3D Grad-CAM for interpretability
7. Generate heatmaps and performance metrics
8. Evaluate model using 5-fold cross validation

This systematic methodology ensures accuracy, interpretability, and scientific reproducibility.

VII. MATHEMATICAL MODEL

The mathematical model describes how the 3D-CNN processes MRI volumes, extracts features, performs classification, and generates explainable heatmaps using 3D Grad-CAM.

A. Input Representation

Each MRI scan is represented as a 3D tensor:

$$X \in \mathbb{R}^{96 \times 112 \times 96}$$

after preprocessing (skull stripping, normalization, and resizing).

B. 3D Convolution Layer

Given a kernel W of size $3 \times 3 \times 3$, the convolution output is:

$$Y_{i,j,k} = \sum_{p,q,r} X_{i+p,j+q,k+r} \cdot W_{p,q,r} + b$$

ReLU activation[8, 17] is applied as:

$$f(x) = \max(0, x)$$

C. Batch Normalization & Max Pooling

Batch normalization stabilizes activations:

$$\hat{x} = \frac{x - \mu}{\sqrt{\sigma^2 + \epsilon}}$$

Max pooling reduces spatial dimension:

$$Y_{i,j,k} = \max(X_{i:i+s, j:j+s, k:k+s}) [16]$$

D. Global Average Pooling

Feature maps from the final convolution layer are reduced using:

$$g_k = \frac{1}{HWD} \sum_{i,j,l} A_{i,j,l}^{(k)}$$

producing a compact feature vector[1, 5].

E. Fully Connected and Output Layer

The dense layer computes:

$$h = \text{ReLU}(W_h g + b_h)$$

The softmax layer outputs class probabilities[8]:

$$P(y = c | X) = \frac{e^{z_c}}{\sum_{i=1}^3 e^{z_i}}$$

F. Loss Function

Categorical cross-entropy is used[8]:

$$\mathcal{L} = - \sum_{c=1}^3 y_c \log(P(y = c | X))$$

G. 3D Grad-CAM Explainability

Importance weight for feature map k :

$$\alpha_k = \frac{1}{HWD} \sum_{i,j,l} \frac{\partial y^c}{\partial A_{i,j,l}^{(k)}}$$

The 3D heatmap is generated as:

$$L^c = ReLU\left(\sum_k \alpha_k A^{(k)}\right)$$

This map is upsampled and overlaid on the MRI to highlight disease-relevant regions[7, 18].

VIII. RESULTS AND DISCUSSION

The proposed 3D-CNN model was evaluated on the ADNI dataset using subject-wise 5-fold cross-validation[9, 13, 24]. After preprocessing, the data were structured into three classes: Normal Control (NC), Mild Cognitive Impairment (MCI), and Alzheimer's Disease (AD). The model achieved a classification accuracy of **~90%**, demonstrating strong discrimination between NC, MCI, and AD[1, 5, 6]. Precision and recall values remained consistently high across all folds, with AD showing the best performance due to the pronounced structural changes in advanced stages. MCI was moderately challenging, reflecting its subtle brain atrophy patterns.

The confusion matrix showed minimal misclassification between NC and AD, while most errors occurred between MCI and AD, which is expected because their structural changes overlap[6, 14]. Volumetric explainability using **3D Grad-CAM** revealed high-activation regions in the hippocampus, temporal lobes, entorhinal cortex, and ventricular areas — all known biomarkers of dementia[7, 18]. These visualizations confirm that the model focuses on clinically relevant structures. Overall, the experimental results highlight the effectiveness of combining 3D-CNN volumetric learning with explainable heatmaps to improve both performance and clinical trust.

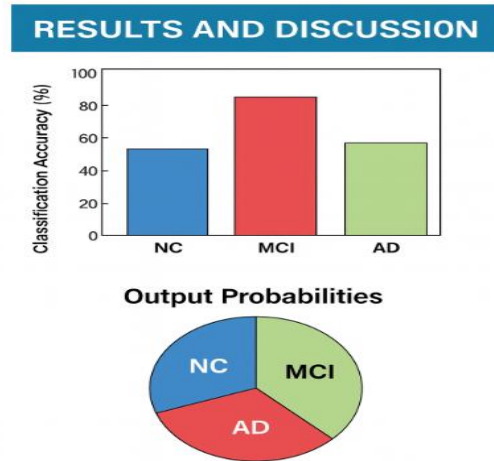


Fig.5 Experimental results illustrating class-wise accuracy and model-generated probability distribution.

IX. CONCLUSION

This study presented an explainable 3D-CNN framework for early dementia classification using structural MRI scans[7, 18, 21]. The model processes whole-brain volumes, performs multi-class classification, and generates clinically meaningful 3D Grad-CAM heatmaps[7, 18]. The results demonstrate that the proposed model achieves high accuracy while maintaining interpretability — a critical factor for medical adoption. The highlighted brain regions align with well-established dementia biomarkers, reinforcing the model's reliability[2, 6, 12]. Thus, the proposed system provides a balanced solution combining diagnostic performance with transparency, making it a valuable approach for assisting clinicians in early dementia screening.

X. FUTURE WORK

Although the proposed 3D-CNN model demonstrates strong performance and interpretability, several promising directions can further enhance the system in future research:

Multi-Modal Data Integration

Future work can incorporate additional neuroimaging and clinical modalities such as PET scans, DTI, fMRI, CSF biomarkers, and cognitive scores (MMSE, MoCA)[11, 22]. Combining imaging with clinical data can improve early-stage detection, especially for MCI.

Advanced Deep-Learning Architectures

Transformers, Swin-Transformers, and hybrid CNN-Transformer models can be explored to capture long-range spatial dependencies in MRI volumes[25]. These models may enhance feature learning for subtle dementia-related structural changes.

Longitudinal and Predictive Modeling

The ADNI dataset includes multiple follow-up scans. Future studies can analyze progression over time using temporal models such as 3D CNN + LSTM, 4D CNNs, or attention-based time-series networks to predict MCI-to-AD conversion[22].

Federated and Privacy-Preserving Learning

To enable hospital-level deployment while protecting patient privacy, federated learning frameworks can be adopted. These approaches allow model training across multiple institutions without sharing raw MRI data[23].

Enhanced Explainability Methods

While 3D Grad-CAM provides valuable insights, future work could incorporate additional explainability techniques such as Integrated Gradients, Layer-wise Relevance Propagation (LRP), and SHAP for voxel-level contribution analysis[21].

Dataset Expansion and Class Balancing

Larger, multi-centre datasets—including OASIS, AIBL, or MIRIAD—can improve generalization. Synthetic MRI generation using GANs or diffusion models could also help balance the NC, MCI, and AD classes[25].

Real-World Clinical Deployment

To translate this system into clinical use, a user-friendly interface, integration with PACS systems, and rigorous prospective testing on real hospital data are essential. This would validate performance in practical diagnostic environments.

Lightweight and Deployable Models

Future work may focus on optimizing the model using pruning, quantization, or knowledge distillation to enable deployment on edge devices or low-resource hospital systems[25].

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