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Deep learning-based classification of dementia using image representation of subcortical signals



Shivani Ranjan¹⁺, Ayush Tripathi¹⁺, Harshal Shende¹, Robin Badal³, Amit Kumar², Pramod Yadav³, Deepak Joshi² and Lalan Kumar^{1,4*}

Abstract

Background Dementia is a neurological syndrome marked by cognitive decline. Alzheimer's disease (AD) and frontotemporal dementia (FTD) are the common forms of dementia, each with distinct progression patterns. Early and accurate diagnosis of dementia cases (AD and FTD) is crucial for effective medical care, as both conditions have similar early-symptoms. EEG, a non-invasive tool for recording brain activity, has shown potential in distinguishing AD from FTD and mild cognitive impairment (MCI).

Methods This study aims to develop a deep learning-based classification system for dementia by analyzing EEG derived scout time-series signals from deep brain regions, specifically the hippocampus, amygdala, and thalamus. Scout time series extracted via the standardized low-resolution brain electromagnetic tomography (sLORETA) technique are utilized. The time series is converted to image representations using continuous wavelet transform (CWT) and fed as input to deep learning models. Two high-density EEG datasets are utilized to validate the efficacy of the proposed method: the online BrainLat dataset (128 channels, comprising 16 AD, 13 FTD, and 19 healthy controls (HC)) and the in-house IITD-AIIA dataset (64 channels, including subjects with 10 AD, 9 MCl, and 8 HC). Different classification strategies and classifier combinations have been utilized for the accurate mapping of classes in both data sets.

Results The best results were achieved using a product of probabilities from classifiers for left and right subcortical regions in conjunction with the DenseNet model architecture. It yield accuracies of 94.17% and 77.72% on the Brain-Lat and IITD-AIIA datasets, respectively.

Conclusions The results highlight that the image representation-based deep learning approach has the potential to differentiate various stages of dementia. It pave the way for more accurate and early diagnosis, which is crucial for the effective treatment and management of debilitating conditions.

Keywords Dementia, Continuous wavelet transform, Deep learning, Frontotemporal dementia, Alzheimer's disease, Mild cognitive impairment

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Introduction

Background & related work

Dementia represents a neurological syndrome impairing cognitive functioning, behaviour, and daily activities [1]. It leads to nerve cell degeneration and disrupted brain communication [2]. The number of people with dementia is expected to double worldwide by 2050 [3], with Alzheimer's disease (AD) being the most prevalent form, significantly contributing to this increase. Despite progress in diagnosing and managing AD, no definitive cure for AD exists. Thus, early or timely detection is a global research priority [4].

Mild cognitive impairment (MCI) is an intermediate stage between healthy ageing and dementia, with a 3-15% annual conversion rate to AD compared to 1-2%in the general population [5]. Frontotemporal dementia (FTD), the second most common form, is characterized by changes in language, behaviour, executive function, and motor symptoms [6]. AD and FTD present similar early symptoms, often leading to misdiagnosis and complicating treatment due to their distinct progression patterns and causes [7].

Diagnostic methods face challenges due to a lack of optimal behavioural tests and the high cost of cerebrospinal fluid (CSF) and blood marker tests [8]. Screening tools such as the Clinical Dementia Rating (CDR) [9], Mini-Mental State Exam (MMSE) [10], Montreal Cognitive Assessment (MoCA) [11], and Addenbrooke's Cognitive Examination III (ACE-III) [12] are useful but have limitations. These limitations include time-consuming administration, reliance on subjective judgments, influence by education level and premorbid intelligence, and less sensitivity at early stages [12]. There is a growing focus on identifying non-invasive brain markers to detect disease pathology before behavioural symptoms appear [13].

Mainstream early diagnosis relies on pathological biomarkers like β -Amyloid and tau Positron Emission Tomography (PET) neuroimaging [14]. AD stages are primarily associated with β -amyloid plaques and tau tangles [15], while FTD involves tau or TDP-43 protein abnormalities [16]. Imaging methods like Computed Tomography (CT), PET [14], and functional Magnetic Resonance Imaging (fMRI) have been used in literature, with fMRI showing higher sensitivity in some cases [17]. Machine learning and MRI-based differentiation [18–21] offer high accuracy in distinguishing these conditions [22]. However, the practical utility of these neuroimaging methods is restricted by high infrastructure costs, less favourability in terms of patient tolerance, and braincomputer interface applications.

Electroencephalogram (EEG) has gained significant attention as a noninvasive tool to analyze brain activity [23, 24]. It has proven reliable in distinguishing dementia

patients from controls [25, 26]. The suitability of EEG for repeated studies and patient monitoring makes it useful in early diagnosis and continuous tracking of AD. EEG detects changes in frequency bands, each corresponding to different functional brain alterations. These include δ (0.5–4 Hz) for slow activity, θ (4–8 Hz) for sleep-wake transitions, α (8–12 Hz) for resting states, β (12–30 Hz) for attention, and γ (above 30 Hz) for complex cognitive processes [27, 28]. This capability aids in defining the neurophysiological profile of AD stages and differentiating it from FTD [29, 30].

However, artifacts from physiological and external sources can obscure or distort crucial frequency bands of EEG signals. This affects neuronal information clarity and integrity. Advancements in signal processing and the use of Machine Learning tools have improved the ability of EEG to differentiate AD from other conditions [26, 31]. These improvements enhance classification accuracy and artifact removal. These tools may also aid in automation and the discovery of new neurophysiological markers [32].

Previous research on differentiating AD from FTD [22, 29-31, 33] and AD from MCI [26, 27, 35-37] has primarily utilized EEG features such as subbands power, Global Field Power (GFP), spectral ratios, and connectivity features, as detailed in Table 1. Despite these insights, diagnosing dementia remains challenging due to the extensive signal analysis required. Effective diagnosis requires a combination of complex features, including time-domain, frequency-domain, and connectivity metrics. As may be noted from Table 1, the current studies have primarily targeted sensor information or variations in cortical regions. However, deep brain regions (subcortical regions), especially the hippocampus, are crucial for accurate AD and FTD classification due to their early involvement in disease progression [38, 39]. AD often begins with neurodegeneration in subcortical areas like the hippocampus before affecting the cerebral cortex [40]. This early involvement makes deep brain regions essential for early diagnosis and precise differentiation between AD and FTD. Detecting changes in these regions can significantly enhance classification accuracy and provide earlier diagnostic insights [39, 41]. Additionally, it has been established that subcortical signals can be detected using surface EEG [42]. Motivated by these, the current study focuses on utilizing time-series signals from deep brain regions, specifically the hippocampus, amygdala, and thalamus for Dementia classification.

Objectives and contributions

In this work, an image representation-based framework has been presented for the classification [43] of three stages of dementia on two different high-density

Approach	Extracted features	Cases	Domain	Reference	Classification problem	Accuracy(%)
sLORETA and ROC	GFP	19 FTD, 19 AD, 21 HC	Source (Cortex)	Nishida et al. [33]	FTD/HC	85.80
					AD/HC	92.8
					FTD/AD	89.8
Decision Trees, Ran- dom Forests	mean, variance, IQR, frequency	19 FTD, 16 AD, 19 HC	Sensor (19)	Miltiadous et al. [29]	AD/HC	86.3
					FTD/HC	78.5
DICEnet	band power, coher- ence	23 FTD, 36 AD, 29 HC	Sensor (19)	Miltiadous et al. [31]	AD/HC	83.3
					FTD/HC	74.9
Gaussian Naïve Bayes	phase lock value, con- nectivity	23 FTD, 36 AD, 29 HC	Sensor (19)	Si et al. [34]	FTD/HC	81.1
					AD/HC	86.2
					FTD/AD	84.7
SVM	graph	23 FTD, 36 AD, 29 HC	Sensor (19)	Rostamikia et al. [30]	FTD/AD	87.8
					AD+FTD/HC	93.5
eloreta	connectivity	75 MCI, 75 HC	Source (Cortex)	Babiloni et al. [35]	MCI/HC	71.0
Logistic Regression	spectral ratios, con- nectivity	64 MCI, 60AD, 65 HC	Sensor (32)	Farina et al. [36]	AD/HC	97.0
					MCI/HC	77.0
					AD/MCI	87.0
Logistic Regression	coherence, spectral power	53 MCI, 26 AD, 55 HC	Sensor (20)	Meghdadi et al. [37]	AD/HC	85.0
					MCI/HC	60.0
Extreme Learning Machine	FuzzyEn, PLV	28 MCI, 21 HC	Sensor (16)	Su et al. [27]	MCI/HC	89.8
CEEDNet	EEG signals, age	417 MCl, 230 AD, 459 HC	Sensor(19)	Kim et al. [26]	Dementia/MCI/AD	76.7

Table 1 A brief description of neurodegenerative disorders classification using EEG

EEG datasets. In the online dataset, the 3-class classification task involves HC and subjects with FTD and AD dementia. The framework has additionally, been validated for in-house collected EEG dataset comprising of subjects with MCI, AD, and HC. The pipeline starts with the extraction of the scout time series corresponding to the left and right regions of the thalamus, hippocampus, and amygdala, using the sLORETA technique. Subsequently, the time series epochs are converted to image representation using a continuous wavelet transform. By utilizing the multi-resolution CWT-based image representation, the time-frequency maps of signals corresponding to the three categories are efficiently learned. In order to learn this mapping, the images corresponding to the left and right regions are fed to standard deep learning model architectures from the computer vision domain such as Xception [44], ResNet [45], InceptionResNet [46], MobileNet [47], NasNetMobile [48], EfficientNet [49], and DenseNet [50]. For classifying the corresponding images, different classification strategies have been adopted. First, the models from the left and right regions are used in isolation for prediction. Subsequently, the sum and product of the posterior probabilities are utilized for the classification task. Additionally, two fusion techniques, namely, early fusion and tensor fusion networks, have also been explored for the purpose of dementia classification on both datasets.

The remainder of the paper is organized as follows: "Materials and methods" section provides the description of the datasets utilized in this study ("Dataset description" section), the scout time series extraction process ("Scout time series extraction" section), preparation of image data ("Image data preparation" section), and the adopted classification strategy ("Classification strategy" section). Experimental details and results are presented in "Experiments and results" section, and "Conclusions" section concludes the paper.

Materials and methods

In this section, the datasets utilized in the study, EEG preprocessing steps, scout time series extraction, image data preparation and classification strategy have been elaborated. A block diagram representing the complete pipeline is presented in Fig. 1.

Dataset description

BrainLat dataset

The dataset utilized in this study is the section of publicly available, preprocessed EEG recordings released by the Latin American Brain Health Institute (Brain-Lat) [51]. The selected subset comprises five-minute EEG recordings from the Latin American population. More specifically, resting-state, eyes-closed recordings from 48 subjects (AD = 16; FTD = 13; HC = 19) were used for the experiments. The recordings were obtained using a 128-channel Biosemi Active II system with pintype active, sintered Ag-AgCl electrodes referenced to contralateral linked mastoids. External electrodes were also placed periocularly to capture blinks and eye movements. Analog filters with a frequency cutoff of 0.03-100 Hz were used to reduce noise. The EEG was monitored online to detect drowsiness, muscle activity, and sweat artefacts.

The recorded data was processed offline using EEGLab [52]. The first step involved in processing steps was average referencing of the EEG data. Subsequently, a bandpass filter was applied between 0.5 and 40 Hz using a zero-phase shift Butterworth filter of order 8. The data was then downsampled from 2048 to 512 Hz.

Independent Component Analysis (ICA) was employed to detect artefacts induced by blinking and eye movements. The components identified as artefacts were then removed to obtain clean EEG data. Malfunctioning channels were identified using a semiautomatic detection method and replaced using weighted spherical interpolation. Finally, the processed EEG signals were stored for subsequent scout time series extraction.

IITD-AIIA dataset

The second dataset utilized in this study consists of inhouse collected resting-state, eyes-closed EEG data recorded from 27 (AD = 10; MCI = 9; HC = 8) righthanded participants aged 60-80 years. The data collection protocol was approved by the Institute Ethics Committee, All India Institute of Ayurveda, New Delhi. EEG data was recorded using a 64-channel Ag/AgCl active electrode EEG setup (actiCHamp, Brain Products GmbH, Germany) with Fz as the reference electrode. The signals were recorded at a sampling rate of 1000 Hz, and the 10:10 EEG electrode placement system was adopted. A conductive EEG gel was applied under each electrode to maintain resistance below 10 k Ω , ensuring a high signal-to-noise ratio. No internal filters were used during the recording. The diagnosis of the AD, MCI, and HC groups were based on the criteria of the MMSE [10] and MoCA screening tools [11]. Only participants whose

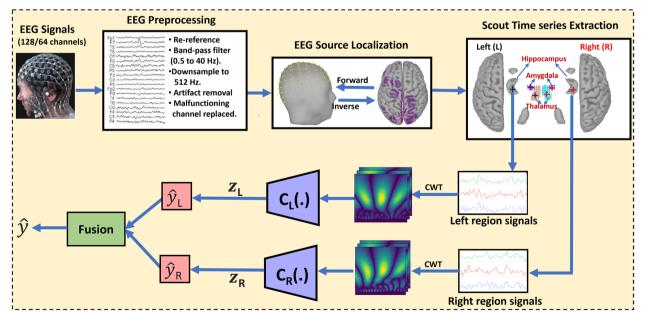


Fig. 1 Block diagram depicting the proposed method. The processed EEG signals are utilized to extract scout time series from the hippocampus, amygdala, and thalamus using sLORETA. The signals are segmented and divided into left and right regions. Subsequently, the CWT-based images are fed to separate classifiers for images corresponding to left and right regions. z_L and z_R represent the latent representation of the classifiers, while \hat{y}_L and \hat{y}_R denote classifier predictions. The latent embeddings are fused using Early and Tensor Fusion, while the individual classifier outputs are fused using probability sum and product

category was consistent across both scales (MMSE: AD < 18, MCI 18–25, HC > 25; MoCA: AD < 21, MCI 21–26, HC > 26) were included in the analysis. HC group participants reported no history of neurological or psychiatric disorders. All participants provided informed consent prior to the study.

Preprocessing and analysis of EEG data were conducted using EEGLAB [52] and MATLAB 2022b. Five-minute segments of continuous EEG data were bandpass filtered between 0.5 Hz and 40 Hz using a fourth-order Butterworth filter to remove irrelevant noise and enhance the signal of interest. ICA was implemented within EEGLAB to visually identify and remove components associated with blinks, eye movements, and muscle artefacts. The preprocessed data were re-referenced to an average reference. Finally, the EEG data for each subject were downsampled to 512 Hz for further scout time series extraction. For participants whose EEG recordings were shorter than five minutes, the maximum available duration was utilized during preprocessing.

Scout time series extraction

EEG source localization aims to identify the primary brain current sources that generate the measured scalp potentials [53]. This process involves solving both the forward and inverse problems. The disparity between the number of EEG channels (128 or 64) and the number of current dipoles to be estimated (approximately 30,020) renders the source localization problem severely underdetermined. Nevertheless, in literature, it has been reported that source localization can be reasonably accurate with 64/128 channels [54].

Forward problem

The forward problem defines the relationship between cortical currents and scalp potentials through a lead field matrix [53]. This matrix models the propagation of currents through head tissues using Neumann and Dirichlet boundary conditions [55]. This relationship can be mathematically expressed as:

$$V = A\tilde{S} + Z$$

where *V* represents the scalp potentials, *A* is the lead field matrix, \tilde{S} denotes the cortical source currents, and *Z* is the sensor noise matrix.

The head model was computed using the Brainstorm toolbox, which involved a mixed model of cortical and deep structures [56, 57]. This model included 30,020 vertices, combining 15,002 from the default cortex and 15,018 from the aseg atlas. The ICBM152 MRI template and the aseg atlas were used to compute the head model for both cortical and subcortical structures, employing OpenMEEG with default conductivity parameters. To

focus on specific regions, an aseg subatlas was created, including the hippocampus (surface scout), thalamus, and amygdala (volume scouts).

Inverse problem

The inverse problem estimates cortical source currents *S* using the lead field matrix *A*. The standard low-resolution electrical tomography (sLORETA) method was employed for this purpose. sLORETA assumes spatial smoothness and coherence among adjacent brain regions [58].

The source currents are estimated by solving the following optimization problem:

$$\min_{S} F = ||V - AS||_2 + \lambda ||S||$$

The solution is given by:

$$\tilde{S} = A^T \left[A A^T + \lambda H \right]^+ V = A_{sLORETA} V$$

where *H* is the average reference operator, and $A_{sLORETA}$ is the inverse kernel relating the recorded scalp potentials *V* to the cortical and subcortical source current estimate \tilde{S} . A sample plot of the grand average brain activation for AD, FTD, and HC cases from the BrainLat dataset is depicted in Fig. 2.

The brain was parcellated into left and right regions for the hippocampus, amygdala, and thalamus using the created aseg subatlas. The constrained current signals were then computed for these regions. For each of the six regions (hippocampus, amygdala, and thalamus, bilaterally) in AD, FTD, and HC cases from the BrainLat Dataset and AD, MCI and HC from the IITD-AIIA Dataset, the sources current signal belonging to a particular region are averaged out to obtain a 6-dimensional time series matrix denoted by \hat{S} . This averaged time series matrix was subsequently used for image data preparation. The EEG source localization plots for BrainLat Dataset that depict the activation difference in the brain regions specifically hippocampus, thalamus, and amygdala for AD, FTD, and HC cases is illustrated in Fig. 2.

Image data preparation

Signals corresponding to the left and right regions of the thalamus, hippocampus and amygdala were extracted using the aforementioned scout time series extraction procedure. The signals are divided into 0.25 seconds epochs which correspond to 128 samples. This corresponds to a 128×6 dimensional matrix \hat{S} , where 6 is the number of signals (corresponding to the left and right thalamus, hippocampus and amygdala). The individual time series is then converted into corresponding image representation by using the Continuous Wavelet Transform (CWT). The underlying principle behind CWT is

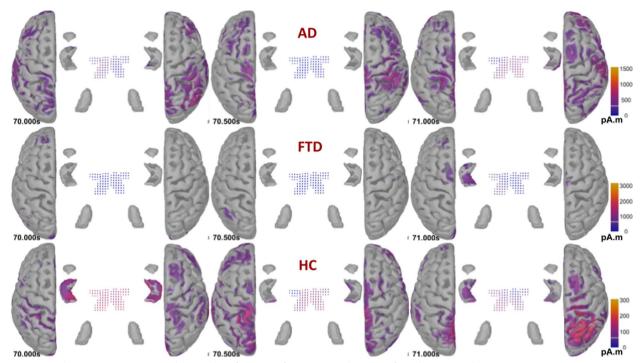


Fig. 2 Grand Average EEG Source Localization plots (front view) for AD, FTD, and HC cases from the BrainLat dataset at timestamps 70s, 70.5s, and 71s. These plots are generated using the Brainstorm Toolbox. The activation maps were set to 20% amplitude, with the amplitude threshold parameter set to "Maximum: Global" for each case

to provide a multi-resolution representation of the time series by varying translation and scale parameters of a mother wavelet [59]. The basis functions for CWT are obtained by scaling and shifting the mother wavelet and can be mathematically expressed as:

$$\Psi_{\sigma,\tau}(t) = \frac{1}{\sqrt{\sigma}} \Psi\left(\frac{t-\tau}{\sigma}\right) \tag{1}$$

Here, the translation is governed by parameter τ which shifts the mother wavelet in time while σ is the scale factor. Normalization by $\frac{1}{\sqrt{\sigma}}$ is done to ensure that the basis function always has unit energy. Once the basis function is defined, the CWT is computed using inner product of the signal with the basis function at different translations and scaling values. For a signal *s*(*t*), this is represented mathematically as,

$$W_{y}^{\Psi}[\sigma,\tau] = s(t) \cdot \Psi_{\sigma,\tau}(t) = \frac{1}{\sqrt{\sigma}} \int_{-\infty}^{\infty} s(t) \Psi^{*}\left(\frac{t-\tau}{\sigma}\right) dt$$
(2)

Finally, wavelet coefficients are obtained by taking all shifts and scales of the Morlet mother wavelet. The aforementioned procedure results in a $128 \times 128 \times 6$ dimensional representation \mathcal{I} of the input data X. This is subdivided into two parts: \mathcal{I}_L and \mathcal{I}_R corresponding to left and right regions, respectively, which are stored subsequently. It is to be noted that each \mathcal{I}_L and \mathcal{I}_R have a dimension of $128 \times 128 \times 3$.

Classification strategy

In order to classify the images into different classes of neurodegenerative disorders (AD, FTD, HC in the BrainLat dataset and AD, MCI, HC in the IITD-AIIA dataset), several different approaches are adopted. In the first approach, two different classifier networks $C_L(.)$ and $C_R(.)$, are individually trained to map \mathcal{I}_L and \mathcal{I}_R to a 3 dimensional vector representing the probabilistic distributions $P(\hat{y}_L = c_i | \mathcal{I}_{\mathcal{L}}; \Theta_L)$ and $P(\hat{y}_R = c_i | \mathcal{I}_{\mathcal{R}}; \Theta_R)$. Here, \hat{y}_L and \hat{y}_R is the predicted class from the left and right classifiers, and $c_i = \{AD, FTD/MCI, HC\}$ is the set of classes. Θ_L and Θ_R are the set of parameters for the classifiers corresponding to the left and right sets of images. Furthermore, a combination of posterior probabilities obtained from individual classifiers is also utilized for the classification task. The sum and product of class probabilities from individual classifiers are computed, and the input X is assigned to class \hat{y}_{sum} or \hat{y}_{mul} based on a maximum of the computed probabilities. Mathematically, the probabilities are computed as $P(\hat{y}_{sum} = c_i | \mathcal{I}_L, \mathcal{I}_R) = P(\hat{y}_L = c_i | \mathcal{I}_L; \Theta_L) + P(\hat{y}_R = c_i | \mathcal{I}_R; \Theta_R)$ and

 $P(\hat{y}_{mul} = c_i | \mathcal{I}_L, \mathcal{I}_R) = P(\hat{y}_L = c_i | \mathcal{I}_L; \Theta_L) * P(\hat{y}_R = c_i | \mathcal{I}_R; \Theta_R)$ for the sum and product cases respectively.

Additionally, two other approaches for using the latent representations of the two classifiers are utilized for the classification task. The latent outputs of the classifiers are fused by using two different strategies: Early Fusion and Tensor Fusion Network [60] to obtain predictions denoted by \hat{y}_{ef} and \hat{y}_{tfn} respectively. The two approaches have been pictorially depicted in Fig. 3. It is to be noted that both early fusion and tensor fusion networks are trained in an end-to-end manner. Several different standard architectures and their variants were utilized for the classifier block:

- Xception [44]: Utilizes depthwise separable convolutions, which improve computational efficiency without sacrificing accuracy.
- ResNet [45]: Introduces residual connections to combat the vanishing gradient problem, enabling deeper networks to be trained effectively.
- InceptionResNet [46]: Combines the strength of Inception modules for multi-scale feature extraction with residual connections.
- MobileNet [47]: Focuses on lightweight architecture through depthwise separable convolutions, suitable for resource-constrained environments.
- NASNetMobile [48]: An architecture discovered through Neural Architecture Search, designed for efficient mobile deployment.
- EfficientNet [49]: Scales network depth, width, and resolution systematically for optimal performance.
- DenseNet [50]: Employs dense connectivity, where each layer is connected to every other layer, promoting feature reuse and improving gradient flow.

For all the classifiers, weights are initialized from the pre-trained models on the ImageNet task. Subsequently, Adam optimizer is used to minimize the cross-entropy loss to learn the final model parameters.

Experiments and results

Experimental details

As elaborated in "Materials and methods" section, the scout time series corresponding to the left and right thalamus, hippocampus and amygdala are segmented to form epochs of 0.25 seconds. For the BrainLat dataset, this segmentation process yields a total of 15408, 12048, and 21648 epochs corresponding to AD, FTD and HC, respectively. Similarly, for the IITD-AIIA dataset, a total of 11088 AD, 10800 MCI and 9600 HC epochs are obtained. For both datasets, from the set of epochs, 80% are randomly selected for training the deep learning models, while evaluation is done on the remaining 20% epochs. Since there is a significant class imbalance in both datasets, different class weights are assigned to samples belonging to different classes based on samples in the majority class to the number of samples in a particular class. This leads to class weights of 1.405, 1.797, and 1 for AD, FTD and HC classes, respectively in the BrainLat dataset. Class weights of 1, 1.027, and 1.155 are assigned to AD, MCI and HC samples of the IITD-AIIA dataset. The model parameters are learnt by using the Adam optimizer while minimizing cross-entropy loss. Furthermore, at the end of each training step, the validation accuracy is monitored on a set of randomly selected 20% samples from the training set. The training process is stopped if the validation accuracy does not improve over a set of 20 continuous training steps.

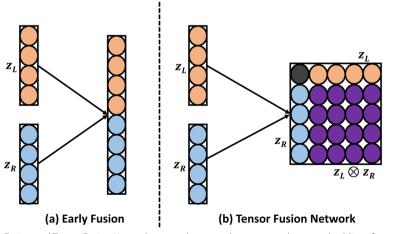


Fig. 3 Depiction of the Early Fusion and Tensor Fusion Network approaches. z_L and z_R represent latent embeddings from the left and right classifiers, respectively

Results and discussion

The performance of the different model architectures and different approaches utilized for the task of dementia classification are presented in Tables 2 and 3 for the BrainLat and IITD-AIIA data, respectively. The results of the individual classifiers (for left and right regions), fusion using sum and product of posterior probabilities and the latent embedding fusion approaches (using early and tensor fusion) are presented. Among the different approaches, using the product of posterior probabilities consistently yields the best classification accuracy for most of the model architectures (12 out of 14 for both datasets). The DenseNet201 emerges as the bestperforming model architecture, yielding accuracies of 94.17% and 77.72% in conjunction with the product of the probabilities approach on the two datasets. The superior performance of DenseNet201 can be attributed to several key architectural features. The dense connectivity of DenseNet201, which connects each layer to every other layer in a feed-forward fashion, ensures that features learned in earlier layers are reused throughout the network. This improves gradient flow and mitigates the vanishing gradient problem, enabling the model to learn more diverse and complex features critical for identifying subtle patterns in EEG-derived images. Despite its depth, DenseNet201 is parameter-efficient, with fewer parameters than models like ResNet, reducing the risk of overfitting, particularly with relatively small datasets. Additionally, its ability to learn hierarchical features across scales-from low-level patterns to high-level abstractions-proves essential in distinguishing neurodegenerative conditions by capturing subtle differences in the activation patterns of subcortical regions.

It may be observed that the classification accuracy on the IITD-AIIA dataset is low compared to the BrainLat dataset. This may be attributed to two main factors. First, the number of samples used for training the model is considerably lower in the case of the IITD-AIIA dataset. In order to learn the complex dynamics from image data, a large number of samples is required, which impacts the overall efficacy of the model. Second, the subcortical source localization in this dataset is done based on a lower number of EEG sensors (64 sensors). A lower number of EEG sensors leads to less accurate localization [54] of the subcortical sources, and hence, the consequent image representations lead to a comparatively lower accuracy score. Nevertheless, an accuracy of 77.72% can be considered a reasonably good performance of the model architecture.

The classification accuracies achieved using IITD-AIIA and BrainLat datasets emphasize the critical role of subcortical regions in the progression and classification of neurodegenerative disorders. Subcortical structures such as the hippocampus, amygdala, and thalamus play a pivotal role in maintaining cognitive functions, including memory, executive functioning, attention, and emotional regulation. Neurodegenerative conditions like AD [19], FTD [21], and MCI [20] are associated with neuronal loss and structural degeneration in these regions, leading to disrupted cognitive functions [1] and altered activation patterns. The pronounced sensitivity of the model to subcortical dynamics, particularly in the right hemisphere, aligns with prior findings of lateralized degeneration. This may be attributed to the fact that the right hippocampus often exhibits greater atrophy in FTD (e.g., 21% vs. 16% tissue loss in FTD) compared to the left hemisphere [38, 41]. This asymmetry in degeneration disrupts neural networks and produces distinct patterns of activation that are effectively leveraged by the model for classification. These findings are consistent with existing literature, which highlights the vulnerability of the right hemisphere in FTD and AD neurodegenerative diseases [19, 20]. The ability of DenseNet201 to leverage subcortical dynamics, combined with its hierarchical feature-learning capabilities, contributes to the accurate classification of neurodegenerative disorders. This integration underscores the

Model	Left	Right	Early Fusion	Tensor Fusion	Sum of Prob	Product of Prob
Xception	89.19	90.80	91.52	90.69	90.97	90.98
ResNet101	89.53	90.16	89.10	86.68	91.68	91.68
ResNet152	88.15	89.62	90.94	90.56	89.91	90.08
InceptionResNetV2	87.82	91.95	90.62	89.29	92.55	92.55
MobileNet	89.10	91.10	91.02	90.89	92.22	92.26
MobileNetV2	89.31	91.78	92.52	90.02	92.91	92.90
NASNetMobile	87.60	90.27	90.37	90.65	90.73	90.72
EfficientNetB2	88.73	91.57	89.97	92.29	92.51	92.55
EfficientNetB3	89.61	90.36	91.33	90.42	92.26	92.33
EfficientNetB4	89.46	91.55	92.15	91.12	92.39	92.39
EfficientNetB5	89.27	90.16	91.44	91.25	91.54	91.54
DenseNet121	90.36	91.25	92.31	90.98	92.68	92.69
DenseNet169	90.70	91.62	92.63	92.50	93.53	93.52
DenseNet201	91.04	92.85	92.32	91.88	94.16	94.17

Table 2 Recognition accuracies using different classifiers utilizing different classification strategies for the BrainLat dataset. The results for the best classification strategy for each model are depicted in blue, and the best result overall is in bold

significance of subcortical regions and the importance of advanced architectures in dementia classification.

In Fig. 4, the confusion matrices for the three class classification tasks on both datasets are presented by using the best-performing model. It may be noted that for the BrainLat dataset, the model is particularly adept at recognizing the Healthy cases of the three classes. The majority of confusion in model predictions comes from the classification of FTD and AD classes. For the IITD-AIIA dataset, the classification accuracies of the individual classes are almost similar to each other. In order to better understand the classification heuristics, a scatter plot obtained by applying t-SNE [61] for dimensionality reduction on the latent embedding vectors is presented in Fig. 5 for both datasets. From the scatter plot, it may be observed that for the BrainLat dataset, the clusters corresponding to AD and FTD (depicted by A and F, respectively) have a significant overlap between them. This is particularly different from the Healthy cases (depicted by H) for which the cluster is significantly different from the other two classes. Subsequently, for the IITD-AIIA dataset, there is a significant overlap between all three clusters (AD, MCI and HC). Therefore, the misclassification trend observed in the confusion matrix is supported by the

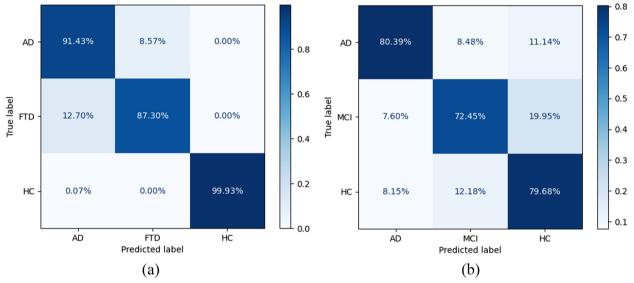


Fig. 4 Confusion matrix using a combination of DenseNet201 and \hat{y}_{mul} for a BrainLat dataset and b IITD-AllA dataset

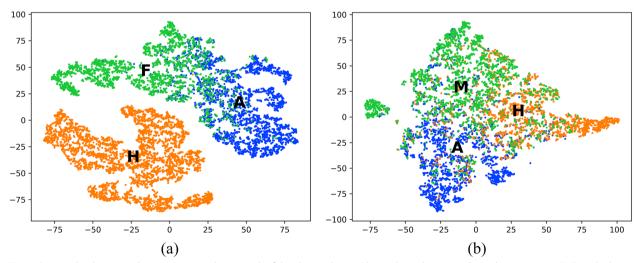


Fig. 5 Scatter plot depicting clusters corresponding to each of the classes obtained by applying dimensionality reduction using t-SNE on the latent embedding vector for a BrainLat dataset and b IITD-AIIA dataset

clusters corresponding to the three classes as presented in Fig. 5. In Fig. 6, the Receiver Operator Characteristics (ROC) and Precision-Recall Curves are depicted for the BrainLat and IITD-AIIA datasets, respectively. The average curves, along with class-wise curves, are depicted in the figures. Average area under ROC values of 0.99 and 0.92 are obtained for the two datasets. Additionally, the average precision value for the two datasets is 0.99 and 0.86. The observations from the curves complement the confusion matrices and the conclusions drawn from the scatter plots.

To evaluate the effectiveness of the continuous wavelet transform (CWT) compared to the short-time Fourier transform (STFT), we performed a classification analysis using the DenseNet201 architecture. For the Product of Probabilities fusion approach, CWT achieved an accuracy of 94.17%, significantly outperforming STFT, which achieved 91.71%. This may be attributed to the dynamic resolution of CWT which enables it to provide a more comprehensive analysis of transient and localized spectral variations [59], that are critical for distinguishing neurodegenerative conditions. Unlike STFT, which relies on a fixed time-frequency resolution determined by the window size, CWT adapts its resolution dynamically [62]. This adaptability allows CWT to effectively represent high-frequency, short-duration components and low-frequency, long-duration components within the same framework. These capabilities make CWT particularly suitable for capturing the nuanced and complex patterns in subcortical EEG data, where transient dynamics are pivotal in classification tasks.

Conclusions

In this work, a dementia classification framework using time-series signals from deep brain regions, specifically the hippocampus, amygdala, and thalamus, is presented. EEG source localization using sLORETA was leveraged to transform the average scout time series signals into image representations using CWT. The images were fed to standard model architectures from the image domain to learn the complex attributes present in the data for reliable dementia classification. The efficacy of the proposed framework was validated on two high-density EEG datasets. An online BrainLat dataset that includes subjects with AD, FTD, and HC, and an in-house collected IITD-AIIA dataset that comprises of subjects with MCI, AD, and HC, were used for the experiments. Various deep learning models, including Xception,

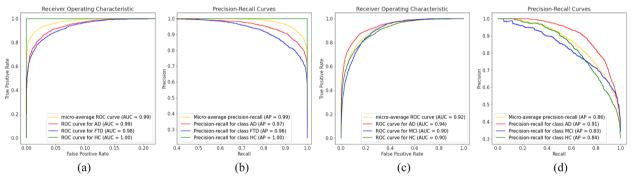


Fig. 6 a, c Multi-class Receiver Operator Characteristic, and b, d Multi-class Precision-Recall Curves for the combination of DenseNet201 and \hat{y}_{mul} on the BrainLat and IITD-AIIA datasets, respectively

Model	Left	Right	Early Fusion	Tensor Fusion	Sum of Prob	Product of Prob
Xception	69.29	69.02	73.71	70.66	76.06	76.61
ResNet101	68.50	67.39	72.02	70.64	73.50	73.85
ResNet152	66.74	66.35	71.36	62.81	73.15	73.52
InceptionResNetV2	67.91	70.74	75.85	73.88	76.31	76.66
MobileNet	67.21	68.72	73.47	68.42	74.44	74.79
MobileNetV2	66.20	66.48	71.71	67.15	72.40	73.20
NASNetMobile	60.57	57.84	70.02	62.49	64.49	65.15
EfficientNetB2	65.88	64.75	74.44	68.24	71.98	72.67
EfficientNetB3	67.10	66.50	71.82	69.83	72.01	72.15
EfficientNetB4	67.37	67.89	71.37	70.09	73.02	73.53
EfficientNetB5	67.99	67.45	72.18	68.23	73.34	73.99
DenseNet121	69.23	69.55	74.90	72.07	75.10	75.52
DenseNet169	70.74	70.45	74.75	70.67	77.17	77.80
DenseNet201	70.55	71.12	75.36	69.20	77.36	77.72

Table 3 Recognition accuracies using different classifiers utilizing different classification strategies for the IITD-AIIA dataset. The results for the best classification strategy for each model are depicted in blue, and the best result overall is in bold

ResNet, InceptionResNet, MobileNet, NasNetMobile, EfficientNet, and DenseNet, were used for classifying the images into one of the three categories for both the datasets. Different classification strategies, including isolated predictions from the left and right brain regions, sum and product of posterior probabilities, early fusion, and tensor fusion networks, were explored to yield optimum classification performance. The experimental results demonstrate that the proposed method achieves high classification accuracy, with the best performance observed using the combination of DenseNet201 and the product of posterior probabilities. The proposed method achieves a classification accuracy of 94.17% on the BrainLat dataset and 77.72% on the IITD-AIIA dataset, highlighting its effectiveness in leveraging deep brain regions for accurate differentiation between AD, FTD, and MCI. By focusing on subcortical regions, which are among the earliest sites of neurodegeneration, our approach facilitates early diagnosis, critical for timely treatment and management. This study also establishes a promising baseline for future research in dementia classification using image representations of subcortical EEG signals. However, the method is limited by the sample size of the datasets and the reliance on high-density EEG recordings, which may not be readily available in all clinical settings. Moreover, the datasets used are geographically limited, which could affect the generalizability of the results to other populations. Future work will focus on increasing the sample size, exploring more sophisticated feature extraction methods, and employing advanced deep learning architectures to enhance system performance further. We believe this methodology paves the way for more accurate and early diagnosis of neurodegenerative conditions, but additional validation across diverse populations is necessary to confirm its broader applicability.

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Authors' contributions

Conceptualization by S.R., A.T., P.Y., and L.K.; Methodology, Software, Formal analysis, Writing – original draft by A.T., and S.R.; Data Curation by S.R., and H.S.; Data acquisition by S.R., H.S., R.B., and A.K.; Clinical diagnosis by R.B. and P.Y.; Writing – review & editing and Supervision by L.K., P.K. and D.J. All authors critically reviewed the manuscript.

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Data availability

The datasets utilized or analyzed in the present study will be accessible upon reasonable request from the corresponding author.

Declarations

Ethics approval and consent to participate

The study was approved by the Institutional Ethics Committee of All India Institute of Ayurveda, New Delhi, India (Ref No. IEC-331/27.06.2023/Rp(E)-12/2023, dated: 17/08/2023). Informed consent was obtained from all participants before the trial commenced. The ethical clearance complies with the Helsinki Declaration.

Consent for publication

Not applicable.

Competing interests

The authors declare no competing interests.

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