# Deep Convolutional Second Generation Curvelet Transform-based MR Image for Early Detection of Alzheimer's Disease

Takrouni Wiem<sup>1</sup> and Douik Ali<sup>2</sup>

<sup>1</sup>University of Sousse, ISITCom, 4011, Sousse, Tunisia, Networked Objects Control and Communication Systems Laboratory (NOCCS-ENISO), 4054, Sousse, Tunisia <sup>2</sup>University of Sousse, ENISO, Networked Objects Control and Communication Systems Laboratory (NOCCS),

Keywords: Second-Generation Curvelet (SGC), Mild Cognitive Impairment (MCI), Multiclass Classification.

Merging neuroimaging data with machine learning has an important potential for the early diagnosis of Abstract: Alzheimer's Disease (AD) and Mild Cognitive Impairment (MCI). The applicability of multiclass classification and the prediction to define the progress of different stages of the disease have been relatively understudied. This paper presents a short review of the deep learning history and introduces a new solution for delineating changes in each stage of AD. Our Deep Convolutional Second-Generation Curvelet Transform Network (SGCTN) is divided into both levels: The feature learning level is the first task that can combine a Second-Generation Curvelet (SGC) with autoencoder trained features. Then, for each hidden layer, a pooling is used to obtain our convolutional neural network. This network is used to learn predictive information for binary and multiclass classification. Our experiments test uses a different number of Cognitively Normal (CN), AD, early EMCI, and Later LMCI subjects from the AD Neuroimaging Initiative (ADNI). Magnetic Resonance Imaging (MRI) information modalities are considered as input. The proposed DSGCCN achieves 98.1% accuracy for delineating the early MCI from CN. Furthermore, for detecting the distinctive level of AD, a multiclass classification test realizes the global accuracy of , and it more particularly differentiates MCI and AD groups from the CN group with 96% accuracy. Compared to the state-of-the-art deep approach, our results indicate that our architecture can achieve better performance for the same databases. Model analysis based (SGC) can improve the classification performance via comparison experiments.

# **1 INTRODUCTION**

Recent research by Alzheimer's Statistics reports that for Alzheimer's Disease (AD) in the world, almost 50 million people have Alzheimer's or related dementia with only one in four people with AD have been diagnosed. AD and other dementia are the top reason for disabilities in later life. Seventy-two percent of the projected rise in the global burden of dementia and pervasiveness by 2050 will take place in low and middle-income countries (Ryu et al., 2017). AD is neuropathologically identified by grievous cell loss and cortical atrophy along with an elevated dementia index as calculated by numbers of neuritic plaques and neurofibrillary tangles in the hippocampus and the neocortex. This disease gets worse with time and later declines cognitive functions and behavioral impairments that touch memory, language and thought, including forgetfulness. Cure strategies for AD are concentrating on preventing the AD evolution or speeding up the clearance of these aggregates. AD, composed of different neurodegenerative levels, which represent the mutation from one stage to another and identifies each one by the specificity of the biomarker. It is indicated that 15-25% of people aged 60 or older have a prodromal stage of AD that can be related to Mild Cognitive Impairment (MCI), which is a transitional stage between dementia and the normal cognitive function. A patient diagnosed with MCI can either have later MCI (LMCI) or Early MCI (EMCI) due to age-linked memory degradation, hence the accent on the importance of early diagnosis of the disease. Thus, accurate and early diagnosis of MCI can help path disease progression, supply better treatment paradigms for patients, and decrease medical costs. Nevertheless, neuroimaging and clinical studies have exposed differences between MCI and Cognitively Normal (CN) (Scheltens and Korf, 2000), (Silverman, 2009). To identify pathological biomarkers to under-

285

Wiem, T. and Ali, D.

DOI: 10.5220/0010228902850292

Copyright © 2021 by SCITEPRESS - Science and Technology Publications, Lda. All rights reserved

Deep Convolutional Second Generation Curvelet Transform-based MR Image for Early Detection of Alzheimer's Disease.

In Proceedings of the 16th International Joint Conference on Computer Vision, Imaging and Computer Graphics Theory and Applications (VISIGRAPP 2021) - Volume 4: VISAPP, pages 285-292 ISBN: 978-989-758-488-6

stand the mechanism causes and monitor early brain changes for each stage of the disease, the quantitative prognosis of AD/MCI by analyzing different types of neuroimaging modalities is necessary for the early classification of AD. Magnetic Resonance Imaging (MRI) allows measuring spatial patterns of atrophy and their growth with disease evolution. It also supplies visual information concerning the macroscopic tissue atrophy, which results from the cellular changes and detects neuronal injury and degeneration underlying AD/MCI (Davatzikos et al., 2011). Positron Emission Tomography (PET) (Nozadi et al., 2018) can be used for the examination of the cerebral glucose metabolism which indicates the functional brain activity. Thereby, a trustworthy diagnosis from brain modalities is necessary, and a sturdy Computer-Aided Diagnosis (CAD) by the data anatomization of neuroimaging will enable for a more reliable and informative approach and can increase potentially diagnostic accuracy. A classical interpretation process for exploring biomarkers for the neuropsychiatric analysis disorders has been founded on the univariate mass statistics within the assumption that various regions of the brain act independently. Nevertheless, this assumption (Fox et al., 2005) is not suitably specified for our present comprehensive brain functioning.

#### 2 RELATED WORK

Machine Learning (ML) methods, which can lay hold of the correlation between regions into account, have become the basic integration and attraction of CAD techniques (Davatzikos et al., 2008), (Suk et al., 2017) and have been broadly used for the automated diagnosis and interpretation of brain disorders. Furthermore, various ML classification models have been used to develop automated neurological disorder prediction. Both major research orientations include Support Vector Machine (SVM) (Pinaya et al., 2016) and Deep Learning (DL) based diagnostic models (Greenspan et al., 2016), (Litjens et al., 2017). SVMbased models of brain disorders have been criticized for their poor performance on raw data and for requiring the expert use of design techniques to extract informative handcrafted features. In contrast, DL models enable a system to use raw data as input, thus allowing them to automatically find highly discriminating features in the training dataset (Shen et al., 2017). As a recent successful category of unsupervised learning models, autoencoders, such as convolutional (Masci et al., 2011), variational (Kingma and Welling, 2013), k-sparse (Makhzani and Frey, 2013), contractive (Rifai et al., 2011) and denois-

ing ones (Vincent et al., 2008), perform an important role in feature extraction, dimension reduction and generative tasks. Higher data dimensionality is an endemic characteristic of medical data, and learning efficient coding for image classification is the goal of these autoencoders. It is crucial to note that these approaches distort spatial locality (neighbor relations) in brain-imaging data (Suk et al., 2017), (Liu et al., 2019), (Payan and Montana, 2015) over the feature extraction level. Many automated systems have been developed in the last years for binary classification (normal or abnormal) of brain MRI, which have made remarkable progress. However, multiclass classification into a specific grade of brain diseases is comparatively more challenging and has great clinical significance. Numerous methods have been utilized to analyze using the wavelet or its variants to extract features for the task of binary and multiclass brain MRI, despite their defeat to capture directional features at numerous levels of resolution (Gudigar et al., 2019), (Jia et al., 2019), (Nayak et al., 2017). In the last mentioned references, the employed classifiers, like the SVM, the fuzzy neural network, and Least-squares SVM endure critical issues such as high computational complexity, poor scalability and slow learning speed. In (Zhang and Suganthan, 2016), the random vector functional link network was a classifier that afforded great generalization performance at the speed learning property. The hybrid approach (Gao et al., 2018) combined the MRI texture features of the contourlet-based hippocampal, the regional CMgl measurement based on fluorine-18 fluorodeoxyglucose-positron emission tomography, medical history, the morphometric volume, the neuropsychological tests of symptoms with the multivariant models to enhance the AD classification and the prediction of MCI conversion, and to appraise whether the partial least squares and the Gaussian process were realizable in developing multivariate models in such a situation. Hence, this situation had various limitations, only hippocampal MRI texture features were explored, the size was approximately modest, and the power statistical might be restricted. Thus, this model was insensitive to high-dimensional data, and dimensionality reduction might upgrade the predictive achievement of this model. However, very few studies have been announced up to now concerning multiclass AD classification, predicting MCI conversion and precision detection. Even more, it remains unknown which method is more appropriate for processing high-dimensional data in this context. To address the above problems, in the present study, we propose a new combination of a Second Generation Curvelet Transform Network (SGCTN) and Deep

Convolutional Autoencoder (DCA) for early detection of AD and prediction of MCI conversion. A DCA consists of operating the classification of a particular class against all the other classes of the dataset by the reconstruction of Deep Convolutional (SGCTN). This reconstruction is achieved using a sequence of a stacked autoencoder and a linear classifier. The remaining part of this paper is composed of four sections. The Deep Convolutional (SGCTN) is proposed in section III. An overview of the dataset utilized in the classification test is given in section IV. Section V includes the experiments and the results of the test. Finally, Section VI summarizes and concludes this paper.

### 3 PROPOSED DEEP CONVOLUTIONAL (SGCTN)

To improve the rate of convergence and classification performance, a new approach called Deep Convolutional (SGCTN) is constructed. In this section, we describe the theoretical background steps that lead to our approach.



Figure 1: The SGCTN for one element of class.

• Step 1: We construct the SGCTN for each object of a class using the best algorithm (Dubois et al., 2015), (Ma and Plonka, 2010). Figure 1 is constituted for three layers. The SGCTN is defined by a much simpler and more natural indexing structure with three parameters: scale, orientation (angle) and location, so curved singularities can be well approximated with very few coefficients and in a non-adaptive manner. We define an SGCTN by pondering a series of second generation curvelets (SGC) interpreted and widened from one mother SGC atom with weight values to approximate a determined signal f:

$$\hat{f} = \sum_{i=1}^{n} (a_i \varphi_i) \tag{1}$$

• Step 2: Every SGCTN is shown in the form of a table (Figure 2). The neuron number that contains a SGC can vary for each SGCTN created in step 1.



Figure 2: Tables of each SGCTN for a class.

• Step 3: In this stage, we select the best SGC (chosen in step 2) to produce an SGCTN for a class. Thereby, we account that our brain dataset is composed of two classes:



Figure 3: Tables of the number of appearance of each SGC for a class.

Class 1 which we will represent in a SGCTN.
Class 2 which includes all the other classes of the dataset.

We build tables for all the SGC in the dataset of SGC for the two classes (Class 1 and Class 2). The tables hold the number of emergences of each SGC in each position in all the SGCTN utilizing the tables in step 2 (Figure 3). Subsequently, we calculate a coefficient for every SGC in that manner: for  $\varphi_i$  in class 1, the coefficient is determined by the sum of all the values of  $\varphi_i$  that are multiplied by (1+ (n - i)) in each position (i) and the identical operation is used for  $\varphi_i$  in class 2. This is indicated as follows

$$Class1_{coef_{\varphi_i}} = Class2_{coef_{\varphi_i}} = \sum_{i=1}^{n} (V_i * (1 + (n-i)))$$
(2)

Whence, the global coefficient for  $\varphi_i$  is defined as follows

$$Glob_{coef_{\varphi_i}} = Class_{1_{coef_{\varphi_i}}} - Class_{2_{coef_{\varphi_i}}}$$
(3)

Next, we calculate all global coefficients of all SGC and arrange them on a table ordered from the biggest to the smallest (Figure 4). The method of coefficient calculation for all SGC enables penalizing the SGC frequently utilized in other classes (in our instance, the classes are combined in one class: class 2) and the preferred SGC used in the



Figure 4: Tables of the global coefficient for each SGC.

working class (Class1). For example, if  $\varphi_i$  is the ideal SGC utilized in class 1 in the first position with 30 emergences and the ideal SGC in class 2 in the second position with 200 emergences, we calculate the  $\varphi_i$  coefficient to punish it and avert it in the first position.



Figure 5: SGCTN for a class.

- **Step 4:** We construct a novel SGCTN for a class (Figure 5) utilizing the ideal SGC from the table in step 3. The total of the SGC utilized for the overall SGCTN is the average of the SGC numbers used in step 2.
- Step 5: In this stage, we construct a network (with the use of the SGC in our SGCTN that produces a class in step 4) using an autoencoder algorithm. It is important to note that an encoder level generates a feature vector (hidden layer) from the input vector with a primal SGC and a decoder level that reconstructs the input vector from the vector of a feature with a dual SGC. In this phase, the neurons of the hidden layer incorporate a linear function. We take into consideration that:

$$f = x_1, x_2, x_3, \dots, x_n$$

$$\hat{f} = \hat{x}_1, \hat{x}_2, \hat{x}_3, \dots, \hat{x}_n$$

$$\phi_i = w_{1i}, w_{2i}, w_{3i}, \dots, w_{ni}$$

$$\hat{\phi}_i = \hat{w}_{1i}, \hat{w}_{2i}, \hat{w}_{3i}, \dots, \hat{w}_{ni}$$
(4)

Then,

$$a_i = \prec f, \varphi_i \succ \Longrightarrow a_i = \sum_{j=1}^n (w_{ji} x_j)$$
 (5)

$$\hat{x}_i = \sum_{j=1}^n \hat{w}_{ji} a_j \tag{6}$$

• Step 6: The hidden layer is used to construct our SGCTN and it is considered for the second training of the input layer. The hidden layer is used to construct our deep convolutional SGCTN and it is considered for the second training of the input layer. The construction of an SGCTN with both hidden layers and a linear classifier is displayed in Figure 6. This process is enforced for all ele-



Figure 6: A deep SGCTN with both hidden layers.

ments of a class until the creation of our deep convolutional SGCTN, which figures the entire class. After that, we replace the linear function by the sigmoid function in the hidden layers to enforce fine-tuning. The choice of using the sigmoid function is to promote the significant features and to derivate an activation function in the backpropagation step. A few new components are shown to be very effective when connected to a convolutional neural network:

\*Local Contrast Normalization (LCN): This component is an efficient technique that makes a deep architecture more sturdy to illumination changes that have not been seen under training. It adapts local subtractive and divisive normalizations, which impose a kind of local competition between features at the identical spatial position in various feature maps and adjacent features in a feature map. It is determined by this function:

$$X_{i+1,x,y} = \frac{X_{i,x,y} - m_{i,N(x,y)}}{\sigma_{i,N(x,y)}}$$
(7)

\*Geometric Lp-norm Pooling (GLP): Pooling is the reducing step of spatial resolution, which aggregates local features over the region of interest into a statistic through a certain spatial pooling operation. The GLP method can preserve the specific-class spatial/geometric information on the pooled features and appreciably boosts the



Figure 7: Geometric Lp-norm pooling of adjacent SGC.

discriminating ability of the resultant features for image classification. It is determined by this function:

$$Out = \left(\sum \sum I(i,j)^{GLP} * G_k(i,j)\right)^{\frac{1}{GLP}}$$
(8)

where I is the input feature map, Gk is a Gaussian kernel, and Out is the output feature map. In our contribution, we achieve intelligence pooling. We operate it if and only if the SGC that are discovered in the neurons are adjacent, whether of an identical type or an identical scale (Figure 7). For that reason, for each hidden layer, GLP and LCN are employed of our SGCTN and will be considered as one step (Figure 8). In addition, our deep convolutional SGCTN will be created.



Figure 8: Typical architecture of Deep convolutional SGCTN.

• **Step 7:** For the last stage, a linear weak classifier is applied, which is defined to be a classifier that is only slightly correlated with true classification. It plainly classifies data with a unique threshold on a certain data dimension.

### **4 DATASET OVERVIEW**

In this work, we use the ADNI http://adni.loni. usc.edu/ dataset including various phases (ADNI-1,

ADNI-2, and ADNI-GO) which is preprocessed by Freesurfer (v5.3). The ADNI has gathered 1167 scans of adults aged between 55 and 90, composed of cognitively normal older persons, individuals with early AD, and individuals with early or late MCI. The demographic details of subjects are provided in Table I. In this paper, we analyze the performance of our architecture on both T1 and T2 weighted MRI images collected from the same set of subjects and evaluate different parameters under both binary and multi class classification. To train our data, parallel processing is needed, so we use open source package python 3.0 to perform the training and validation of the classifier (GPU: 1xTesla K80, having 2496 CUDNN cores, compute 3.7, 24GB(23.439GB Usable) GDDR5 VRAM). We use Keras library over Tensorflow modules to design our proposed architecture.

#### 5 EXPERIMENTS AND RESULT

We compare the performance of our model while training and testing with brain MRI together with (Original Features Curvelet Network (OFCN), SGCTN), and segmented hippocampal regions.

#### 5.1 Devising Training and Test Set

We augment the MRI data to resist the model from geometry changes and noise. We use 5482 MRI slides in our experiment. We split the data into 85:15 training and testing sets based on subjects to defeat the intra relation amid the split data to carry out the classification without biasing.

# 5.2 Classification Results and Discussion

The focus of this work is established on demonstrating how the proposed approach can resolve the most discriminating elements related to the progression of MCI and ameliorate binary and multiclass classification performance. After the selection of features, we train the initial SGCTN to learn the fundamental vector of features. Those features are connected to the other input of the autoencoder to learn secondary features in order to discriminate the various levels of the disease. Our network includes seven hidden layers and a softmax layer for output. We test the classification performance in both cases, which is OFCN and SGCTN. In this latter, we choose five scales and eighteen orientations. The results provide the confusion matrix to perform the quality of our architecture,

Categories	Age	MMSE	CDR	CDR GS
AD (284 sc)	$75.85 \pm 7.94$	$23.4{\pm}2.1$	4.5 to 9	$0.7 \pm 0.3$
LMCI (169 sc)	$72.99 \pm 7.67$	$27.1 \pm 1.9$	2.5 to 4.5	$0.5 \pm 0.2$
EMCI (328 sc)	$72.62 \pm 7.33$	$28.4{\pm}1.5$	0.5 to 2.5	$0.5 {\pm} 0.0$
Contols (371 sc)	$75.68 \pm 8.01$	29.1±0.9	0 to 0.5	$0.0{\pm}0.0$

Table 1: Demoghraphic details about the participant study.

which contains actual and predicted class information. The following five metrics are considered:

$$Accuracy(ACC) = \frac{TP + TN}{TP + TN + FP + FN}$$
(9)

$$Sensitivity(SEN) = \frac{TP}{TP + FN}$$
(10)

$$Specificity(SPEC) = \frac{TN}{TN + FP}$$
(11)

$$PositivePredictiveValue(PPV) = \frac{TP}{TP + FP} \quad (12)$$

Table II proves the results of our network, witch allows distinguishing between CN/AD, CN/EMCI and EMCI/LMCI. The results are more accurate with 99.1%, 98.1% and 93.3 % in the tasks of CN vs AD, CN vs EMCI and EMCI vs LMCI, respectively for the deep convolutional SGCTN using whole image compared with the OFCN in the same tasks with hippocampal patch are 92.4% 90.8 % and 89.7% is more accurate than the OFCN using whole image. On the



Figure 9: ROC curves to multiclass classification.

other hand, the OFCN model achieves good performance in both tasks. This can be explained by the autoencoder selecting only each class and separating the approximation to the components of the other class. The selection based on the principal contribution of each best SGC to the construction of the SGCTN for each component of a class. We note that the SGC stays unchanged and that the weight changes from one slide to another for both classes. Then, based on a SGC, which, in fact includes an extension of the isotropic multiresolution analysis concept to include anisotropic scaling and angular dependence (directionality) while preserving rotational invariance. The SGC is also faster and less redundant compared to its original features curvelet (OFC), It does not exhibit blocking artefacts due to special partitioning.

We use the learning curve of the multiclass classification using the proposed deep convolutional SGCTN (Figure 9). The classifier can more accurately differentiate CN, AD and LMCI from an EMCI class, so the overall classification performance is ameliorated significantly. It can be noted that the learning curve related to the test is closer to the learning curve of validation. As there are three classes instead of both classes, the model can learn more generic discriminative elements through all three classes. Then, the confusion matrix evaluates the performance with each line corresponding to a true class. The components of the diagonal of the confusion matrix depict the point number for the predicted label which is equal to the true label, whereas the components of the off-diagonal are those that are misclassified by the classifier (Figure 10). Compared with the deep learn-

. . . . . .

	Confusion matrix on SGCIN features							
CN	0.97	0.02	0.00	0.01	-1.0 -0.8			
EMCI	0.02	0.95	0.03	0.00	-0.6			
LMCI	0.00	0.04	0.96	0.00	-0.4			
AD	0.00	0.01	0.05	0.94	-0.2			
	CN	EMCI Predic	LMCI ted label	AD				

-----

Figure 10: AD vs. LMCI vs. EMCI vs. CN classification confusion matrix.

ing classification model and geometric transformbased feature extraction pattern analysis based on ADNI data (Fang et al., 2020), (Wee et al., 2019), (Ramzan et al., 2020), it should be noted that in Table III our proposed architecture still achieves better performance in the terms of classification accuracy for the two tasks (EMCI vs. LMCI and, CN vs. EMCI). This can be considered for the use of autoencoder based SGC, which gives better feature extraction than other method based on a deep model with

Task	Features	OFCN			SGCTN				
		ACC	SEN	SPEC	PPV	ACC	SEN	SPEC	PPV
CN/AD	hippocampal	92.4 %	93.4%	90.2%	90.4%	97.3%	98.3%	96.1%	96.5%
	whole slide	91.2%	91.9%	89.9%	90.2%	99.1%	99.8%	98.2%	98.6%
CN/EMCI	hippocampal	90.8%	91.3%	89.3%	89.7%	96.9%	97.2%	95.2%	95.6%
	whole slide	90.6%	92.1 %	88.8%	88.9%	98.1%	98.8%	97.2%	97.6%
LMCI/EMCI	hippocampal	89.7%	90.1%	87.9%	88.0%	91.9%	93.2%	90.8%	90.9%
	whole slide	86.8%	88.1%	85.8%	85.9%	93.3%	94.5%	92.7%	92.9%

Table 2: Binary classification performance comparison of OFCN features and SGCTN features.

	CN vs. AD	CN vs. EMCI	EMCI vs. LMCI
(Fang et al., 2020)	-	79.25%	83.33%
(Wee et al., 2019)	81.0%	-	-
(Ramzan et al., 2020)	-	96.85%	88.6%
Proposed	99.1%	98.1%	93.3%

or without a transform based on multiresolution analysis(Gao et al., 2018), (Hofer et al., 2020), (Swain et al., 2020).

# 6 CONCLUSIONS

In this study, a novel deep convolutional SGCTN for brain disease image classification, which combines the flexibility of SGC with autoencoder technique to extract and reduce these features. By this method, a series of trained autoencoders are accomplished by a linear classifier and are stacked to build a deep convolutional neural network. The obtained classification CN vs. EMCI results indicate that our architecture can perform well the delineation of the fluffiest changes associated with the EMCI group. After the reconstruction of deep convolutional SGCTN layers, high accuracy of 98.2% is obtained, which display the potential of the proposed approach for clinical diagnosis of the early level of AD. Also, the deep convolutional SGCTN achieves high accuracy in the task of EMCI vs. LMCI of 93.3 %, as well as an AUC score of 96.1 % CN vs. EMCI and EMCI vs. LMCI classification results are considered as the best classification performance obtained so far. A future work, we aim to focus on considering the advantage of the proposed approach to build a computer-aided diagnosis system that can help in the EMCI delineating group in the process of multiclass classification, which can be helpful in the planning of early therapy and ameliorative interventions.

#### REFERENCES

Davatzikos, C., Bhatt, P., Shaw, L. M., Batmanghelich, K. N., and Trojanowski, J. Q. (2011). Prediction of mci to ad conversion, via mri, csf biomarkers, and pattern classification. *Neurobiology of aging*, 32(12):2322–e19.

- Davatzikos, C., Fan, Y., Wu, X., Shen, D., and Resnick, S. M. (2008). Detection of prodromal alzheimer's disease via pattern classification of magnetic resonance imaging. *Neurobiology of aging*, 29(4):514–523.
- Dubois, S., Péteri, R., and Ménard, M. (2015). Characterization and recognition of dynamic textures based on the 2d+ t curvelet transform. *Signal, Image and Video Processing*, 9(4):819–830.
- Fang, C., Li, C., Forouzannezhad, P., Cabrerizo, M., Curiel, R. E., Loewenstein, D., Duara, R., Adjouadi, M., Initiative, A. D. N., et al. (2020). Gaussian discriminative component analysis for early detection of alzheimer's disease: A supervised dimensionality reduction algorithm. *Journal of Neuroscience Methods*, 344:108856.
- Fox, M. D., Snyder, A. Z., Vincent, J. L., Corbetta, M., Van Essen, D. C., and Raichle, M. E. (2005). The human brain is intrinsically organized into dynamic, anticorrelated functional networks. *Proceedings of the National Academy of Sciences*, 102(27):9673–9678.
- Gao, N., Tao, L.-X., Huang, J., Zhang, F., Li, X., O'Sullivan, F., Chen, S.-P., Tian, S.-J., Mahara, G., Luo, Y.-X., et al. (2018). Contourlet-based hippocampal magnetic resonance imaging texture features for multivariant classification and prediction of alzheimer's disease. *Metabolic Brain Disease*, 33(6):1899–1909.
- Greenspan, H., Van Ginneken, B., and Summers, R. M. (2016). Guest editorial deep learning in medical imaging: Overview and future promise of an exciting new technique. *IEEE Transactions on Medical Imaging*, 35(5):1153–1159.
- Gudigar, A., Raghavendra, U., San, T. R., Ciaccio, E. J., and Acharya, U. R. (2019). Application of multiresolution analysis for automated detection of brain abnormality using mr images: A comparative study. *Future Generation Computer Systems*, 90:359–367.
- Hofer, C., Kwitt, R., Höller, Y., Trinka, E., and Uhl, A. (2020). An empirical assessment of appearance descriptors applied to mri for automated diagnosis of

tle and mci. Computers in Biology and Medicine, 117:103592.

- Jia, W., Muhammad, K., Wang, S.-H., and Zhang, Y.-D. (2019). Five-category classification of pathological brain images based on deep stacked sparse autoencoder. *Multimedia Tools and Applications*, 78(4):4045–4064.
- Kingma, D. P. and Welling, M. (2013). Auto-encoding variational bayes. arXiv preprint arXiv:1312.6114.
- Litjens, G., Kooi, T., Bejnordi, B. E., Setio, A. A. A., Ciompi, F., Ghafoorian, M., Van Der Laak, J. A., Van Ginneken, B., and Sánchez, C. I. (2017). A survey on deep learning in medical image analysis. *Medical image analysis*, 42:60–88.
- Liu, M., Zhang, J., Lian, C., and Shen, D. (2019). Weakly supervised deep learning for brain disease prognosis using mri and incomplete clinical scores. *IEEE Transactions on Cybernetics*.
- Ma, J. and Plonka, G. (2010). A review of curvelets and recent applications. *IEEE Signal Processing Magazine*, 27(2):118–133.
- Makhzani, A. and Frey, B. (2013). K-sparse autoencoders. arXiv preprint arXiv:1312.5663.
- Masci, J., Meier, U., Cireşan, D., and Schmidhuber, J. (2011). Stacked convolutional auto-encoders for hierarchical feature extraction. In *International conference on artificial neural networks*, pages 52–59. Springer.
- Nayak, D. R., Dash, R., Majhi, B., and Prasad, V. (2017). Automated pathological brain detection system: A fast discrete curvelet transform and probabilistic neural network based approach. *Expert Systems with Applications*, 88:152–164.
- Nozadi, S. H., Kadoury, S., Initiative, A. D. N., et al. (2018). Classification of alzheimer's and mci patients from semantically parcelled pet images: a comparison between av45 and fdg-pet. *International journal of biomedical imaging*, 2018.
- Payan, A. and Montana, G. (2015). Predicting alzheimer's disease: a neuroimaging study with 3d convolutional neural networks. arXiv preprint arXiv:1502.02506.
- Pinaya, W. H., Gadelha, A., Doyle, O. M., Noto, C., Zugman, A., Cordeiro, Q., Jackowski, A. P., Bressan, R. A., and Sato, J. R. (2016). Using deep belief network modelling to characterize differences in brain morphometry in schizophrenia. *Scientific reports*, 6:38897.
- Ramzan, F., Khan, M. U. G., Rehmat, A., Iqbal, S., Saba, T., Rehman, A., and Mehmood, Z. (2020). A deep learning approach for automated diagnosis and multiclass classification of alzheimer's disease stages using resting-state fmri and residual neural networks. *Journal of Medical Systems*, 44(2):37.
- Rifai, S., Mesnil, G., Vincent, P., Muller, X., Bengio, Y., Dauphin, Y., and Glorot, X. (2011). Higher order contractive auto-encoder. In *Joint European conference on machine learning and knowledge discovery in databases*, pages 645–660. Springer.
- Ryu, C., Jang, D. C., Jung, D., Kim, Y. G., Shim, H. G., Ryu, H.-H., Lee, Y.-S., Linden, D. J., Worley, P. F.,

and Kim, S. J. (2017). Stim1 regulates somatic ca2+ signals and intrinsic firing properties of cerebellar purkinje neurons. *Journal of Neuroscience*, 37(37):8876–8894.

- Scheltens, P. and Korf, E. S. (2000). Contribution of neuroimaging in the diagnosis of alzheimer's disease and other dementias. *Current opinion in neurology*, 13(4):391–396.
- Shen, D., Wu, G., and Suk, H.-I. (2017). Deep learning in medical image analysis. *Annual review of biomedical* engineering, 19:221–248.
- Silverman, D. (2009). *PET in the Evaluation of Alzheimer's Disease and Related Disorders*. Springer.
- Suk, H.-I., Lee, S.-W., Shen, D., Initiative, A. D. N., et al. (2017). Deep ensemble learning of sparse regression models for brain disease diagnosis. *Medical image analysis*, 37:101–113.
- Swain, B. K., Sahani, M., and Sharma, R. (2020). Automatic recognition of the early stage of alzheimer's disease based on discrete wavelet transform and reduced deep convolutional neural network. In *Innovation in Electrical Power Engineering, Communication, and Computing Technology*, pages 531–542. Springer.
- Vincent, P., Larochelle, H., Bengio, Y., and Manzagol, P.-A. (2008). Extracting and composing robust features with denoising autoencoders. In *Proceedings of the* 25th international conference on Machine learning, pages 1096–1103.
- Wee, C.-Y., Liu, C., Lee, A., Poh, J. S., Ji, H., Qiu, A., Initiative, A. D. N., et al. (2019). Cortical graph neural network for ad and mci diagnosis and transfer learning across populations. *NeuroImage: Clinical*, 23:101929.
- Zhang, L. and Suganthan, P. N. (2016). A comprehensive evaluation of random vector functional link networks. *Information sciences*, 367:1094–1105.