

Detection of Alzheimer's Disease and Dementia States Based on Deep Learning from MRI Images: A Comprehensive Review

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Abstract

Many studies have been conducted to examine abnormal conditions in brain structures and to detect Alzheimer's and Dementia states using features derived from medical images. From these data, it is very important to detect the diagnosis of Alzheimer's and Dementia disease early and to provide appropriate treatment to the patients. Quality magnetic resonance (MR) images are requested to make this diagnosis. But while producing a quality image, it also brings less spatial coverage and longer scanning and identification time. In this context, biomedical image processing has undergone a serious expansion and has become an interdisciplinary research field that includes many fields. Computer Aided systems have become an important part in the diagnosis process. With the development of computer aided systems, producing quality information for the diagnosis of disease in image processing applications has caused various problems. Such difficulties are tried to be overcome with artificial intelligence technology and super-resolution (SR), which has gained great importance in image processing lately. Using the super resolution methodology, a high resolution image is obtained from the low resolution image. Thus, the image processing timing is shortened and an image with desired features can be obtained. This shortens the irritating and long-lasting MR imaging process. In addition, it provides convenience for the diagnosis of the disease with the improvements it provides on MR images. Recovering the image is an important step in this process. The quality of the reconstructed image depends on the restoration methods. The functionality of artificial intelligence technology in image processing and biomedical fields is increasing day by day. The deep learning method is preferred in techniques aimed at obtaining a reconstructed quality image. At the same time, various artificial intelligence methods are widely used for classifying and detecting the data obtained. One of the most

common of these is neural network (NN) methods.. Deep learning, a special method of neural networks, is widely used in classification methods due to its superior structural properties. When studies are examined, it is seen that DL methods are widely used. The success of the proposed methods is increasing day by day.

Keywords

Alzheimer's disease; Dementia disease; super-resolution; Deep learning

1. Introduction

Computer-aided detection systems (CADs) are tools for detecting abnormal conditions in medical imaging processes and increasing diagnostic accuracy using advanced image processing and pattern recognition techniques.

Due to the prevalence of Alzheimer's and dementia disease, many studies have been carried out on the automatic detection of these diseases with computer-aided systems in recent years. Biomedical imaging has been developed for the purpose of patient care as well as for the study of biological structure and function, and to address some disorders. In this context, it can help diagnose Alzheimer's and Dementia diseases in the early stages, which can lead to more effective treatments. Advances in magnetic resonance imaging (MRI), a technique of digital radiography, serve this purpose.

1.1. Magnetic Resonance Imaging (MRI)

Magnetic resonance imaging (MRI) is a medical imaging technique that uses synchronously with computer-assisted systems to produce images of various regions of the body. It is often used for disease detection and diagnosis. It is based on a technology that takes advantage of the movement change in the axis of the protons in the water that make up living tissues.

An MRI scanner contains two powerful circular magnets. The human body is largely made up of water molecules consisting of hydrogen and oxygen atoms. At the center of each atom are protons that act as a magnet and are sensitive to the magnetic field. These protons are stimulated by the application of radiofrequency current on the patient. This application spins the various protons of the body. When the radiofrequency field is closed, the signal detected by the receiver part of the MRI scanner is exposed. Signals are processed using biomedical imaging methods and an MR image is created.

The patient should remain still for a long time during the imaging process, so as not to blur the MR image. This brings some problems with it. In order to obtain the image quickly, the intravenous method is used on the patient before MRI or

during the procedure. This process improves image quality and acquisition speed. However, recently, various biomedical imaging methods have been used to increase the speed and quality of image acquisition. The scheme of the operation of the computer-aided system is shown in figure 1.

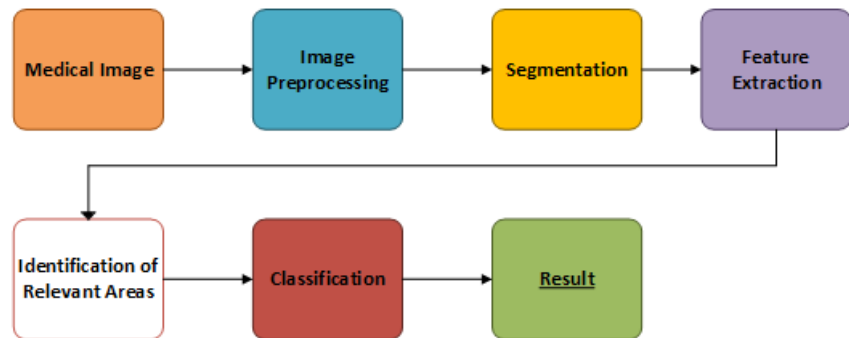


Figure 1. Functioning scheme of computer aided system

1.2. MRI Applications

MRI scan helps to provide a precise diagnosis in many cases and positively affects treatment. In addition, MRI is frequently used in the follow-up of healed diseases. The main usage areas are listed below.

- **Cardiovascular:** Magnetic resonance imaging (MRI) is preferred in cardiovascular imaging due to its high efficiency for blood circulation and cardiac motion analysis [10].
- **Musculoskeletal:** MRI has experienced great advances in image contrast. This feature is used for disease detection and diagnosis. It provides evaluation of tissue structures and bone marrow deformations in bones. It provides visualization of structures around the bone marrow, such as adipose tissue. Also provides visualization of muscle diseases [11].
- **Liver and gastrointestinal:** Liver and gastrointestinal tumors are independent tumors that spread in the region where they are located. Cancer cells can spread out of the tissue in which they are located, or directly to other areas by blood-lymph vessels. With MRI, it is possible to detect these cancerous areas in the gastrointestinal region and liver [12].
- **Angiography:** Magnetic resonance angiography (MRA) is used effectively in the diagnosis of vascular-related diseases in the brain. The vascular network is desired to be examined in three dimensions in the brain. In this case, due to the excess data in the three-dimensional MR image, the scanning time will be longer [13].
- **Neuroimaging:** MRI is preferred for the diagnosis of many diseases as it

provides visualization of brain anatomy containing brainstem and cerebellum. It allows diagnosis of many neurological diseases, especially Alzheimer's disease and dementia. Therefore, MRI is frequently preferred for Neuroimaging.

1.3. Recent Studies

Alzheimer's disease is a neurological disorder. As time progresses, these disorders increase their effects and cause a loss of memory and thinking skills. As these disorders progress, the patient's cognitive loss increases and leads to dementia. [14]. In research and results, progressive brain disorder is the most common cause of dementia in older people [20]. AD destroys nerve cells and causes tissue loss in the relevant region. Thus, it significantly reduces the volume of the disease-related region in the brain over time [9]. This decrease in volume disrupts the functions that will take place in that region or completely eliminates its effect. Besides Alzheimer's psychological activities, it also significantly affects people's daily routine. It is estimated that the number of people affected by the disease will double in the coming years, and one in 85 people is expected to have AD [15]. Morbidity represents the frequency of a disease or the amount of the disease, if any, and the symptoms caused by the disease [20]. Figure 2 also shows the morbidity rate of the most common neuro-psychiatric and neurological diseases.

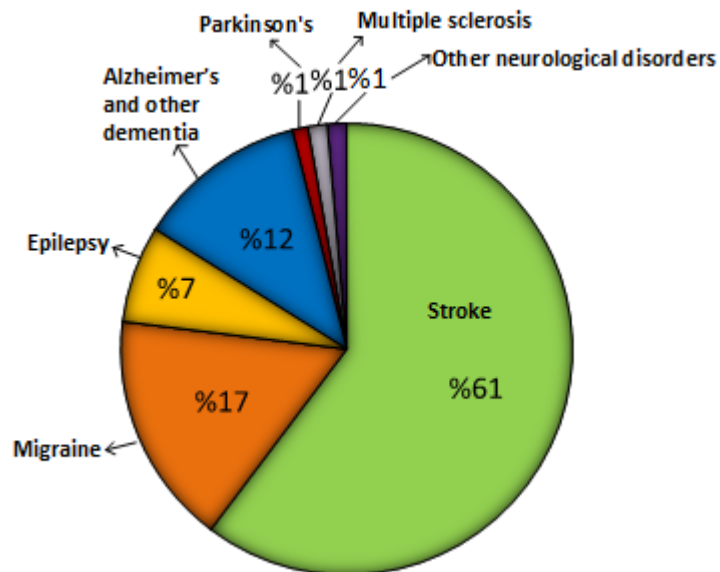


Figure 2. Morbidity rate of Neuro-psychiatric and Neurological Diseases [64]

In many studies in the literature, imaging biomarkers of Alzheimer's and dementia diseases have been identified using the magnetic resonance imaging (MRI) method.

In current studies, computer-aided detection systems continue to be developed for Alzheimer's and dementia disease detection using brain MR images.

2. Models

In recent years, Deep Learning (DL) techniques based on Artificial Neural Network (ANN) have been successfully applied for diagnosis and detection of diseases. The basic operation in the networks established in these techniques is designed by inspiring the work of the human brain. Nonlinear operations are applied using many signals entering the system and output signals are created. These techniques reflect representative learning. Because these models do not use an explicit method when extracting features from the data. The feature extraction process is performed by using hidden data layers from the data [20]. For the feature extraction process in deep learning models, it is not necessary to extract features separately as in the machine learning technique [20]. In previous studies, techniques such as support vector machine, k-tools and neural network were used in the diagnosis of neurological diseases [16]. In addition, the techniques used in the diagnosis of some disorders that cause disease in humans have been inspired by nature and have provided important achievements [17, 18]. Early diagnosis in Alzheimer's disease (AD) is very important in preventing the progression of the disease and ensuring its treatment. For AD detection, features extracted from brain MR images are classified [19]. Features should accurately represent AD-related regions of structures such as ventricular size, hippocampus shape, cortical thickness, and brain volume [19]. The MRI neuroimaging tool has been explored to develop such a system [21, 22]. Features derived from existing MRI images are divided into three groups: Alzheimer's disease, mild cognitive impairment (MCI), normal control (NC) group and classification is established [23, 24, 25, 26]. This article examines the deep learning methods used to diagnose Alzheimer's and dementia based on MRI scans. Deep Neural Network (DNN) [28, 34-42], Deep Belief Networks (DBNs) [20, 43, 44], Restricted Boltzmann Machines (RBMs) [20], Deep Auto-encoder (DA) [32, 45-48], Recurrent Neural Network (RNN) [49], Deep Boltzmann Machine (DBM) [50], and Convolutional neural networks (CNN) [33, 51-61] are important and widely preferred deep learning models. As a result of the research, it has been seen that the methods are also applied in different diseases.

In this section, 4 different models emerging within the scope of deep learning are examined. Deep Neural Network (DNN), Convolutional Neural Networks (CNN), Deep Automatic Encoder (DA), Deep Boltzmann Machine (DBM) models are described below.

2.1. Model-1: Deep Neural Networks (DNN)

Deep Neural Network (DNN) has a structure consisting of input layer, output layer and hidden layers [27]. Deep learning allows to learn the representations of the data obtained thanks to the numerous calculation layers in its structure [28]. DNN uses the feed forward feature of the neural network for data detection [20]. In the method, feature extraction and clustering are carried out with an unsupervised approach. In the field of biomedical imaging, DNN is mostly used for classification and reducing the symptoms of the related disease. Figure 3 shows the architecture of the DNN learning process.

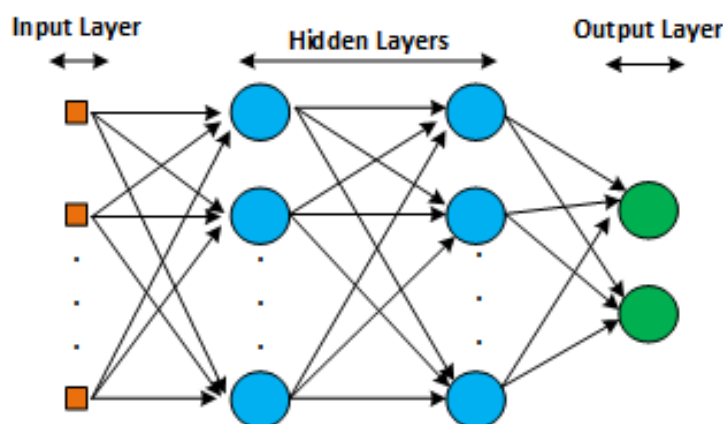


Figure 3. Architecture of DNN

2.2. Model-2: Convolutional Neural Networks (CNN)

CNN is one of the most widely used deep learning techniques. CNN is a model in which the image information is taken from the input in the method, transferred to the output in one direction and obtained the output according to the desired classification. CNN consists of input, output and multiple hidden layers [20]. CNN topology uses relationships between data to reduce the number of parameters required in education. At the same time, thanks to these relationships, forward feeding improves [14]. An exemplary model is shown in Figure 4. This proposed model is called 3D adaptable CNN. While the lower layers of a 3D-CNN developed for a specific purpose separate and detect the general properties, the upper layers make it easier to classify the problem by using these properties [29].

2.3. Model-3: Deep Automatic Encoder (DA)

Features extracted using data-based learning in artificial intelligence architecture can often be more accurate. However, Deep Auto Encoder (DA) has the same number of input and output nodes and is trained to rebuild the input vector instead

of assigning a class tag [30]. For this reason, Deep Auto Encoder (DA) is a type of neural network performed unsupervised. The Deep Autoencoder model consists of three layers: input (encoder), output (decoder) with the same number of nodes as input, and hidden (code). The number of hidden layer nodes is less than the same number of input and output node layers. Encoder layers encode the input data (MRI). With this coding process, the size of the input is reduced. In the next process, this data is presented to the decoder by the hidden layer. In the last step, the decoder layers return the input data to their original form [20]. The architecture of DA is shown in Figure 5 [31]. The feature extraction method in the DA model is not linear [32]. Therefore, it does not need to classify the data used in education.

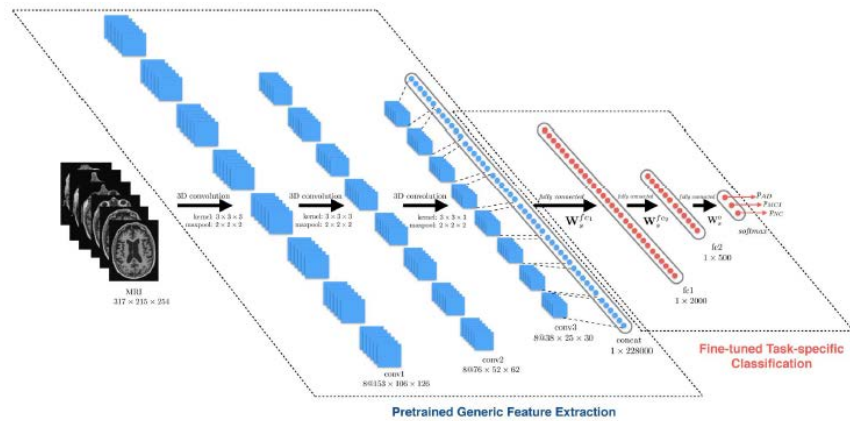


Figure 4. Proposed CNN structure [19]

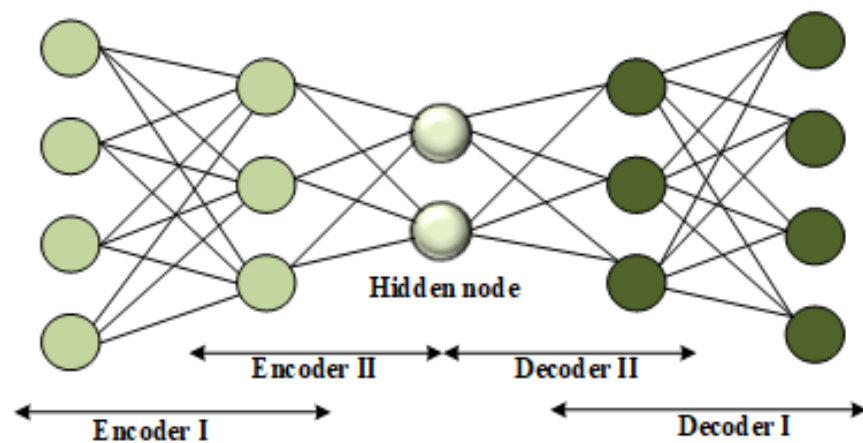


Figure 5. Architecture of DA

2.4. Model-4: Deep Boltzmann Machine (DBM)

The Deep Boltzmann Machine (DBM) is an unsupervised model, such as DNN, that does not have a complete relationship between its layers. This model contains a large number of hidden unit nodes. The Deep Boltzmann machine (DBM) is formed by the combination of the Restricted Boltzmann Machines (RBMs). Multiple RBM structures are combined into consecutive structures to establish DBM structures [20]. There is no interlayer link in RBMs. Existing connections are only available between adjacent layer units. The same applies to DBM as its structure has RBM structure properties. The DBM method can be routed for complex and ambiguous inputs and provides robust output data. The structure of DBM is shown in Figure 6.

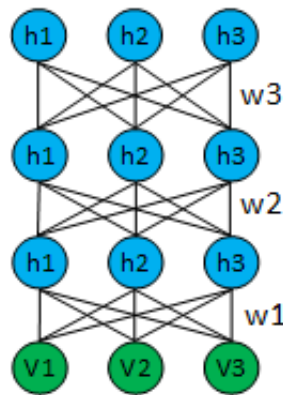


Figure 6. Architecture of DBM

3. Results and Comparisons

In all models examined, the data are from Alzheimer's Disease Neuroimaging Initiative (ADNI) [33]. Results and comparisons were made in this dataset. In the table, the accuracy rate of the disease diagnosis, instances and modality are given for the four models examined. Accuracy values are represented in percentage (%).

Table 1. Accuracy Diagnostic Studies of Models Examined on AD.

	Model			
	DNN[35]	CNN[52]	DA[45]	DBM[50]
Accuracy	99.2	99.9	91.95	95.35
Instances	1409	302	338	
Modality	MRI	MRI and Functional MRI	MRI	PET and MRI

When the accuracy rates in the table are examined, different effects of deep learning

methods are observed. The imaging technique used and the number of samples used by the models may be different. The differences between the sample numbers affect the accuracy rates.

In this study examines four deep learning models for Alzheimer's and Dementia disease. In the literature review, it is observed that these four models are widely used for the diagnosis of other neurological disorders. The ratio of the studies carried out using these four DL techniques between 2010-2019 in Google Scholar is shown in Figure 7.

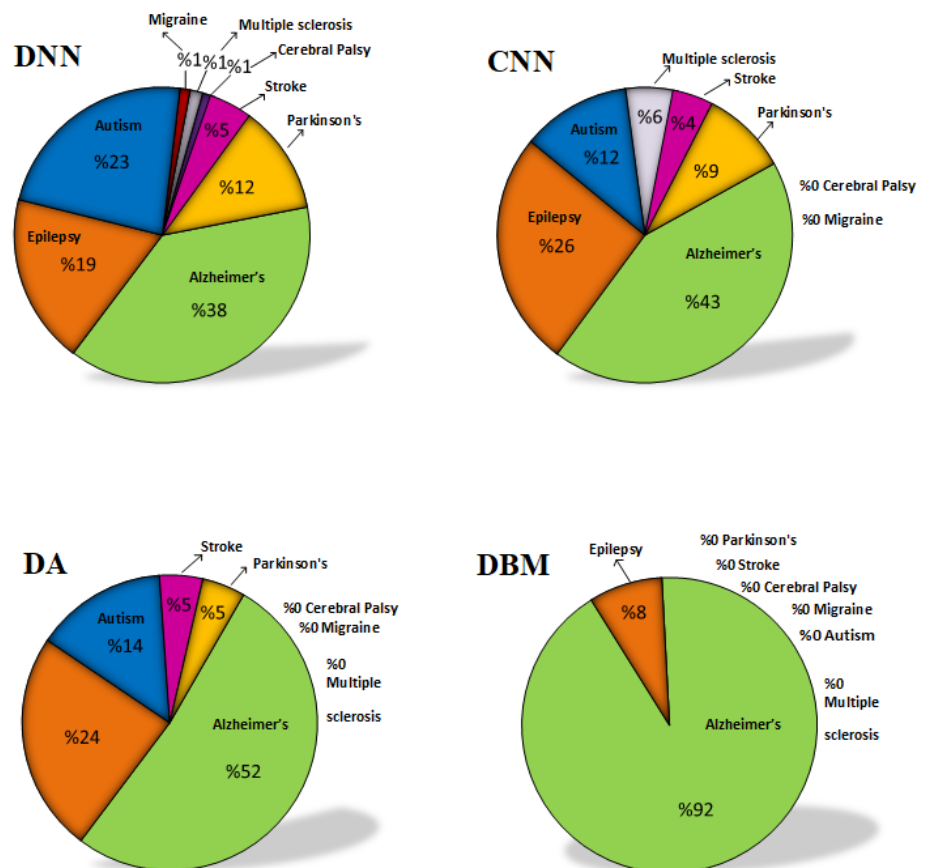


Figure 7. The ratio of the articles on other diseases in the Google Scholar database (2010-2019) of the four Deep Learning techniques examined in our study [20].

4. Conclusion

Neurological disorders negatively affect human life and even threaten human life in the future. In recent years, with the development of computer-aided systems, DL techniques have also been improved and have been widely preferred in establishing neurological diagnoses. The data used in the diagnosis of the disease are the MR

image data that are consumed in large quantities in storage environments that are significantly higher. The processing and training of this large volume of data is one of the challenges that can be encountered in techniques.

In this study, a compilation of DL methods used for the diagnosis of Alzheimer's and Dementia diseases commonly seen from Neurological and Neuropsychiatric Disorders is presented. It has been observed that MRI image data are used effectively in the diagnosis of Alzheimer's and Dementia diseases. Four deep learning techniques commonly used for the detection of these diseases are briefly described. The high accuracy rates of these DL methods applied to MRI images in disease diagnosis are remarkable. In addition, the usage rates of these four techniques for other major Neurological and Neuropsychiatric Disorders have been found. This study did not aim to compare methods with each other. Undoubtedly, biomedical imaging and deep learning techniques are improving day by day and new methods will continue to be discovered.

Conflicts of Interest

There is no conflict of interest.

References

- [1] Kennedy, D. P., and Adolphs, R. (2012) The social brain in psychiatric and neurological disorders. *Trends Cogn Sci* 16(11):559–572.
- [2] Orru, G., Pettersson-Yeo, W., Marquand, A. F., Sartori, G., and Mechelli, A., (2012) Using SVM to identify imaging biomarkers of neurological and psychiatric disease: A critical review. *Neurosci Biobehav Rev* 36(4):1140–1152.
- [3] Spasov, S., Passamonti, L., Duggento, A., Lio, P., and Toschi, N. (2019) Alzheimer's Disease Neuroimaging Initiative. A parameter-efficient deep learning approach to predict conversion from mild cognitive impairment to Alzheimer's disease. *NeuroImage* 189: 276–287.
- [4] Cheng, D., and Liu, M. (2017) CNNs based multi-modality classification for AD diagnosis. In 2017 10th international conference on image and signal processing, BioMedical Engineering and Informatics, IEEE, 1-5.
- [5] Farooq, A., Anwar, S., Awais, M., and Rehman, S. (2017) A deep CNN based multi-class classification of Alzheimer's disease using MRI. In 2017 IEEE international conference on imaging systems and techniques, IEEE, 1-6.
- [6] Gunawardena, K. A. N. P., Rajapakse, R. N., and Kodikara, N. D. (2017) Applying convolutional neural networks for pre-detection of Alzheimer's disease from structural MRI data. In 2017 24th international conference on mechatronics and machine vision in practice, IEEE, 1-7.
- [7] Billones, C. D., Demetria, O. J. L. D., Hostallero, D. E. D., and Naval, P. C. (2016) DemNet: A Convolutional Neural Network for the Detection of Alzheimer's Disease and Mild Cognitive Impairment. In IEEE Region 10

- Conference, 3724–3727.
- [8] Payan, A., and Montana, G. (2015) Predicting Alzheimer's disease: a neuroimaging study with 3D convolutional neural networks. arXiv preprint arXiv:1502.02506:1–9.
 - [9] McKhann, G. et al., (1984) Clinical diagnosis of Alzheimer's disease: Report of the NINCDSADRDA Work Group* under the auspices of Department of Health and Human Services Task Force on Alzheimer's Disease. *Neurology*, vol. 34, no. 7, pp. 939.
 - [10] Christodoulou, AG., Zhang, H., Zhao, B., Hitchens, TK., Ho, C. and Liang, ZP. (2013) High-resolution cardiovascular MRI by integrating parallel imaging with low-rank and sparse modeling. *IEEE Trans Biomed Eng.* 2013 Nov;60(11):3083-92. doi: 10.1109/TBME.2013.2266096.
 - [11] Helms, C., (2008). *Musculoskeletal MRI*. Saunders. ISBN 978-1-4160-5534-1.
 - [12] Baboi, LM. et al., (2007) Characterization of neuro-endocrine tumors in an athymic nude mouse model using dedicated synchronization strategies for T2-weighted MR imaging at 7T. *Conf Proc IEEE Eng Med Biol Soc.* 2007;2007:2879-82.
 - [13] Zhaoyang, J., Yiping, P.D. (2012) Application of partial-echo compressed sensing in MR angiography. 5th International Conference on BioMedical Engineering and Informatics. doi: 10.1109/BMEI.2012.6513105.
 - [14] Sarraf, S. and Tofighi, G. (2016) Classification of Alzheimer's Disease using fMRI Data and Deep Learning Convolutional Neural Networks. *Computer Vision and Pattern Recognition.* arXiv:1603.08631 [cs.CV]
 - [15] Alzheimer's Association et al., (2014) Alzheimer's disease facts and figures, *Alzheimers Dement*, vol. 10, no. 2, pp. e47–e92, 2014.
 - [16] Sharma, M., Singh, G., and Singh, R., (2017) Stark assessment of lifestyle based human disorders using data mining based learning techniques. *IRBM.* 38:305–324.
 - [17] Gautam, R., Kaur, P., and Sharma, M., (2019) A comprehensive review on nature-inspired computing algorithms for the diagnosis of chronic disorders in human beings. *Progress in Artificial Intelligence.* 8(4):1–24.
 - [18] Kaur, P., and Sharma, M., (2019) Diagnosis of human psychological disorders using supervised-learning and nature-inspired computing techniques: A meta-analysis. *J Med Syst* 43(7):204.
 - [19] Hosseini-Asl, E., Keynton, R., El-Baz, A., (2016) Alzheimer's disease diagnostics by adaptation of 3D convolutional network. *IEEE International Conference on Image Processing (ICIP).*
 - [20] Gautam, R. and Sharma, M. (2020) Prevalence and Diagnosis of Neurological Disorders Using Different Deep Learning Techniques: A Meta-Analysis. *Journal of Medical Systems* volume 44, Article number: 49 (2020). <https://doi.org/10.1007/s10916-019-1519-7>.
 - [21] Jack, C.R. et al., (2011) Introduction to the recommendations from the National Institute on Aging-Alzheimer's Association workgroups on diagnostic guidelines for Alzheimer's disease. *Alzheimers Dement*, vol. 7, no. 3, pp. 257–262.

- [22] McKhann, G.M. et al., (2011) The diagnosis of dementia due to Alzheimers disease: Recommendations from the National Institute on Aging-Alzheimers Association workgroups on diagnostic guidelines for Alzheimer's disease. *Alzheimers Dement*, vol. 7, no. 3, pp. 263–269.
- [23] Bron, E.E. et al., (2015) Standardized evaluation of algorithms for computer-aided diagnosis of dementia based on structural MRI: The CADDementia challenge. *NeuroImage*.
- [24] Cuingnet, R. et al., (2011) Automatic classification of patients with Alzheimer's disease from structural MRI: a comparison of ten methods using the ADNI database. *NeuroImage*, vol. 56, no. 2, pp. 766–781.
- [25] Falahati, F. et al., (2014) Multivariate Data Analysis and Machine Learning in Alzheimer's Disease with a Focus on Structural Magnetic Resonance Imaging. *J. of Alzheimer Dis*, vol. 41, no. 3, pp. 685–708.
- [26] Sabuncu, M.R. et al., (2015) Clinical Prediction from Structural Brain MRI Scans: A Large-Scale Empirical Study. *Neuroinformatics*, vol. 13, no. 1, pp. 31–46.
- [27] Deng, L., (2014) A tutorial survey of architectures, algorithms, and applications for deep learning. *APSIPA Transactions on Signal and Information Processing* 3:1–29.
- [28] Gulhare, K. K., (2017) Shukla, S. P., and Sharma, L. K., Deep Neural Network Classification method to Alzheimer's Disease Detection. *International Journals of Advanced Research in Computer Science and Software Engineering* 7(6):1–4
- [29] Long, M. and Wang, J. (2015) Learning transferable features with deep adaptation networks. *arXiv:1502.02791 [cs.LG]*.
- [30] Ravi, D., Wong, C., Deligianni, F., Berthelot, M., Andreu-Perez, J., Lo, B., and Yang, G., (2017) Deep learning for health informatics. *IEEE Journal Of Biomedical and Health Informatics* 21(1):4–21.
- [31] Pouyanfar, S., Sadiq, S., Yan, Y., Tian, H., Tao, Y., Reyes, M. P., and Iyengar, S. S., (2018) A survey on deep learning: Algorithms, techniques, and applications. *ACM Computing Surveys (CSUR)* 51(5): 92.
- [32] Dolph, C. V., Alam, M., Shboul, Z., Samad, M. D., and Iftekharuddin, K. M., (2017) Deep learning of texture and structural features for multiclass Alzheimer's disease classification. In *2017 International Joint Conference on Neural Networks*, 2259–2266.
- [33] Rieke, J., Eitel, F., Weygandt, M., Haynes, J. D., and Ritter, K., (2018) Visualizing convolutional networks for MRI-based diagnosis of Alzheimer's disease. In *Understanding and Interpreting Machine Learning in Medical Image Computing Applications*, Springer, Cham, 24-31.
- [34] Benyoussef, E. M., Elbyed, A., and El Hadiri, H., (2018) 3D MRI classification using KNN and deep neural network for Alzheimer's disease diagnosis. In *International Conf. on Advanced Intelligent Sys. Sustainable Development*, Springer, Cham, 154–158.
- [35] Basaia, S., Agosta, F., Wagner, L., Canu, E., Magnani, G., and Santangelo, R., (2019) Alzheimer's Disease Neuroimaging Initiative: Automated classification of Alzheimer's disease and mild cognitive impairment using a single MRI and

- deep neural networks. *NeuroImage: Clinical* 21(1-8):101645.
- [36] Kim, D., and Kim, K. (2018) Detection of Early Stage Alzheimer's Disease using EEG Relative Power with Deep Neural Network. In 40th Annual International Conference of the IEEE Engg. in Medicine and Bio. Society, 352–355.
- [37] Cireşan, D. C., Giusti, A., Gambardella, L. M., and Schmidhuber. (2013) Mitosis detection in breast cancer histology images with deep neural networks. In *International Conf. on Med. Image-Computing and Computer-assisted Intervention*, Springer, Heidelberg, 411–418.
- [38] Ramesh, S., Caytiles, R. D., and Iyengar, N. C. S. (2017) A deep-learning approach to identify diabetes. *Advanced Science and Tech Letter* 145:44–49.
- [39] Ma, X., Yang, H., Chen, Q., Huang, D., and Wang, Y. (2016) Depaudionet: An efficient deep model for audio-based depression classification. In *Proceedings of the 6th International Workshop on Audio/Visual Emotion Challenge*, ACM, 35–42.
- [40] Heinsfeld, A. S., Franco, A. R., Craddock, R. C., Buchweitz, A., and Meneguzzi, F. (2018) Identification of autism spectrum disorder using deep learning and the ABIDE dataset. *NeuroImage: Clinical* 17:16–23.
- [41] Daoud, H. G., Abdelhameed, A.M., and Bayoumi, M. (2018) Automatic epileptic seizure detection based on empirical mode decomposition and deep neural network. In *2018 IEEE 14th International Colloquium on Signal Processing & Its Applications (CSPA) IEEE*, 182–186.
- [42] Kadam, V. J., and Jadhav, S. M. (2019) Feature Ensemble Learning Based on Sparse Auto-encoders for Diagnosis of Parkinson's Disease. In *Computing, Communication and Signal Processing*, Springer, Singapore. 810:567–581.
- [43] Kim, J., Kang, U., and Lee, Y. (2017) Statistics and deep belief network based cardiovascular risk prediction. *Healthcare Informatics Research* 23(3):169–175.
- [44] Cai, H., Sha, X., Han, X., Wei, S., and Hu, B. (2016) Pervasive EEG diagnosis of depression using Deep Belief Network with three electrodes EEG collector. In *2016 IEEE International Conference on Bioinformatics and Biomedicine*, 1239–1246.
- [45] Shi, B., Chen, Y., Zhang, P., Smith, C. D., and Liu, J. (2017) Alzheimer's Disease Neuroimaging Initiative: Nonlinear feature transformation and deep fusion for Alzheimer's Disease staging analysis. *Pattern Recogn* 63:487–498.
- [46] Bhatkoti, P., and Paul, M. (2016) Early diagnosis of Alzheimer's disease: A multi-class deep learning framework with modified k-sparse autoencoder classification. In *International Conference on Image and Vision Computing New Zealand*, IEEE, 1-5.
- [47] Xu, J., Xiang, L., Liu, Q., Gilmore, H., Wu, J., Tang, J., and Madabhushi, A. (2016) Stacked sparse autoencoder for nuclei detection on breast cancer-histopathology images. *IEEE Trans Med Imaging* 35(1):119–130.
- [48] Chen, L., Zhou, M., Su, W., Wu, M., She, J., and Hirota, K. (2018) Softmax regression-based deep sparse autoencoder network for facial emotion recognition in human-robot interaction. *Inf Sci* 428:49–61.
- [49] Cui, R., Liu, M. (2019) Alzheimer's disease neuroimaging initiative: RNN-based longitudinal analysis for diagnosis of Alzheimer-disease.

- Computerized Med. Imaging and Graphics, 1-25.
- [50] Suk, H. I., Lee, S. W., and Shen, D. (2014) Alzheimer's disease neuroimaging Initiative Hierarchical feature representation and multimodal fusion with DL for AD/MCI diagnosis. *Neuroimage*. 101: 569–582.
- [51] Awate, G., Bangare, S., Pradeepini, G., and Patil, S. (2018) Detection of Alzheimers Disease from MRI using Convolutional Neural Network with Tensorflow. arXiv preprint arXiv:1806.10170.
- [52] Sarraf, S., DeSouza, D. D., Anderson, J., and Tofighi, G. (2017) DeepAD: Alzheimer's disease classification via deep convolutional neural networks using MRI and fMRI. *BioRxiv*, 070441:1 - 15.
- [53] Kruthika, K. R., and Maheshappa, H. D. (2019) Alzheimer's disease neuroimaging initiative: CBIR system using capsule networks and 3D CNN for Alzheimer's disease diagnosis. *Informatics in Medicine Unlocked* 14:59–68.
- [54] Aderghal, K., Khvostikov, A., Krylov, A., Benois-Pineau, J., Afdel, K., and Catheline, G., (2018) Classification of Alzheimer Disease on Imaging Modalities with Deep CNNs Using Cross-Modal Transfer Learning. In 2018 IEEE 31st international symposium on computer-based medical systems, IEEE, 345-350.
- [55] Tang, H., Yao, E., Tan, G., and Guo, X. (2018) A Fast and Accurate 3D Fine-Tuning Convolutional Neural Network for Alzheimer's Disease Diagnosis. In *International CCF Conference on Artificial Intelligence*, Springer, Singapore, 115–126.
- [56] Cui, R., and Liu, M. (2018) Hippocampus Analysis by Combination of 3D DenseNet and Shapes for Alzheimer's Disease Diagnosis. *IEEE J Biomed Health Inform* 23(5):1–8.
- [57] Wang, H., Shen, Y., Wang, S., Xiao, T., Deng, L., Wang, X., and Zhao, X. (2019) Ensemble of 3D densely connected convolutional network for diagnosis of mild cognitive impairment and Alzheimer's disease. *Neurocomputing* 333:145–156.
- [58] Kumar, S., Negi, A., and Singh, J. N. (2019) Semantic Segmentation Using Deep Learning for Brain Tumor MRI via Fully Convolution Neural Networks. In *Information and Communication Technology for Intelligent Systems*, Springer, Singapore, 11–19.
- [59] Ren, X., Xiang, L., Nie, D., Shao, Y., Zhang, H., Shen, D., and Wang, Q. (2018) Interleaved 3D-CNN s for joint segmentation of small volume structures in head and neck CT images. *Med Phys* 45(5): 2063–2075.
- [60] Yang, H., Zhang, J., Liu, Q., and Wang, Y. (2018) Multimodal MR based classification of migraine: using deep learning CNN. *Biomedical engineering online* 17 (1): 138.
- [61] Li, G., Liu, M., Sun, Q., Shen, D., and Wang, L., (2018) Early diagnosis of autism disease by multi-channel CNNs. In *International Workshop on Machine Learning in Medical Imaging*, Springer, Cham, 303–309.
- [62] Wang, S.H., Phillips, P., Sui, Y., Liu, B., Yang, M., and Cheng, H. (2018) Classification of Alzheimer's disease based on eight-layer convolutional neural network with leaky rectified linear unit and max pooling. *J Med Syst*

42(5):85.

- [63] Islam, J., and Zhang, Y. (2017) A novel deep learning based multi-class classification method for Alzheimer's disease detection using brain MRI data. In International Conference on Brain Informatics, Springer, Cham, 213–222.
- [64] Feigin, V. L., Abajobir, A. A., Abate, K. H., Abd-Allah, F., Abdulle, A. M., Abera, S. F., and Aichour, M. T. E., Global, regional, and national burden of neurological disorders during 1990–2015: A systematic analysis for the global burden of disease study 2015. *The Lancet Neurology* 16(11):877–897, 2017.