

# A Systematic Literature Review of Medical Image Analysis Using Deep Learning

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**Abstract**—We review literature in top journals and conferences on the usage of deep learning for medical image analysis in modern healthcare. As a result it is shown that deep learning offers unique capabilities and breakthroughs in identifying, classifying and segmenting different kinds of medical images, especially related to cancer in the breast, lung, and brain.

**Keywords**—data-driven medical diagnosis, medical image analysis, deep learning, literature review

## I. INTRODUCTION

One of the key aspects in today's medical diagnosis process is the use of medical imaging (MI), as it is used in all fields of modern medicine [1]. Medical Image Analysis (MIA) is almost always a complex and time-consuming task which even professionals struggle with [2, 3]. Today's technological progress could make it possible to use new methods in the health domain. The use of Deep Learning (DL) could provide a wide variety of new possibilities for different medical fields, for example, it could be used to make faster and more accurate diagnoses which in turn would lead to better and more sophisticated personalized treatment [4].

The following paper will provide an overview of the current state of MIA using DL technology.

## II. RESEARCH METHODOLOGY

We conducted a systematic literature search capturing deep learning-related work in the digital health domain until 09/12/2019, including four meta-databases (IEEE, ACM, AIS Basket of 8, SpringerLink/"Image Processing" and "Data Mining and Knowledge Discovery") that met the inclusion criteria of "deep learning" AND medical AND "image analysis". Only international peer-reviewed publications were considered.

## III. RESULTS

### A. Cancer Tissue Identification

The most commonly researched field of DL in MIA is *cancer* and *tumour* detection and classification. The accurate identification and categorization of cancer structures and subtypes plays a major role in histological images. Studies showed promising results for the automatic analysis of *cancer tissue* by using DL approaches. For example, with the use of FAST [5] it is possible to make volume segmentation in CTs. The image is

passed to a patch generator and a volume renderer. This patch generator splits the volume into sub-volumes which in turn is passed onto the convolutional neural network (CNN). The segmentation network then processes the volume which is then stitched back together with the help of a patch stitcher. This processed volume is then rendered to highlight the segmented tumor [5]. In another example, CNNs were able to organize and extract the discriminative information from the data [6]. One of the most reliable cancer diagnosis methods is the histopathological examination. The main downside to it is that it is time consuming and prone to errors because it requires a detailed inspection by a pathologist. With the help of DL in MIA it is possible to make this process faster and more efficient since it wouldn't rely necessarily on human interference [7]. The use of CNNs makes it possible to segment cancerous tissue and to classify it in many cases [8].

### B. Mammographic Applications

Another popular use for DL in MIA is *breast cancer* detection. Breast cancer is one of the main causes of death in females and there is a rise in the number of diagnoses, but there are not enough specialists to take care of the patients. The use of DL applications would be a possible solution to speed up the process and therefore offer the possibility of earlier treatment [9]. In addition, Yunchao and Faqiang [10] presented an approach for predicting breast cancer with the help of DL in MIA, which is more reliable than before and thus is able to replace most of the pathologists' workload.

In the medical subdiscipline of breast cancer detection, a fusion of a CNN and a Support Vector Machine (SVM) was able to detect cancerous tissue with an accuracy of 92% [11]. With the help of the CNN it was possible to narrow down the region of interest (ROI) in digital mammograms [12].

### C. Medical Imaging of Lungs

Anthimopoulos et al. [14] described an approach to *lung* pattern classification using CNNs. They designed a network that captures low level textural features of the lung tissue. These image patches were generated through the annotations of a CT slice. In this example, the CNN processed the patches and assigned them to the proper area of unhealthy tissue. The patches have a 100% overlap with the lung, at least 80% with the ground-truth and 0% with each other [13].

In the past years, DL techniques in general became the state-of-art in the widespread field of image classification. For example, diffuse lung diseases are a challenge for medical doctors since the disease comes in many different forms. ML applications are able to help physicians with their work by drastically reducing the time needed to perform a MIA of the lung patterns [14]. Currently, CT and PET scans are used to detect lung cancer, as by combining the imagery it allows the detection of metabolically active lesions. With the use of CNNs it was possible to retrieve the information from the CTs only. This cuts the time of the analysis of the MI and thus make it more efficient while maintaining a high level of accuracy [15].

#### D. Brain Tissue Segmentation

4D CT imaging could be essential in the future for workups after a stroke [16]. One major breakthrough was made in *dementia* diagnosis. With the help of DL applications in MIA it was possible to detect early signs of dementia in testing samples [17]. These predictions of *Alzheimers disease* would help enable earlier treatment and therefore slow down the degenerative process and provide support in sustaining patients' quality of life for as long as possible [17].

#### E. Miscellaneous Applications

Another medical field in which the pattern of the images is diverse and subtle is *fetal anatomy* assessment. DL applications in MIA are able to detect the organs of the yet unborn fetus [18]. In clinical routines, it is common to use a 2D Ultrasound (US) for check-ups during pregnancy but some monitoring tasks can not be done properly because of multiple factors including, for example, the orientation of the fetus. One possible solution would be the use of a 3D US, but to fully check on the fetus, prior organ localization is needed. This information is difficult to obtain since every fetus varies in size and orientation. Raynaud et al. [18] propose the use of DL applications in the analysis of the 3D US imagery as a solution to accommodate for this problem. One of their examples is a combined spine detector. With the use of DL and morphological filter responses is it possible to combine the two outputs and obtain a more robust spine binary mask.

For *Cervical Histopathology Image Classification* (CHIC), many ML methods were developed and applied to image segmentation, feature extraction or classification tasks. So, the ML methods are constantly updated with the current state-of-art of technology in CHIC [19].

*Gastrointestinal (GI) disease* is a widely occurring illness which is examined by GI endoscopy. With the help of CNNs it is possible to detect, classify and segment GI diseases [20].

Stacked Sparse Autoencoder (SSAE) is a DL application which allows high level features to be captured through learning processing from low level features (pixel). In *nucleus and cytoplasm morphology* analysis, SSAE helps the physicians to indicate if the cells are normal or abnormal [21]. Methods for *thoracic aorta calcifications* (TAC) or *coronary artery calcification* (CAC) also rely on segmentation to separate the ROIs in the images [22].

DL approaches are also widely used in *cardiac image segmentation* were the high accuracy of those methods have

made them the state-of-the-art in this field. For example, the CNNs used by Duan et al. [23] were able to localize landmarks and segment the CMRs simultaneously. The network combined the computational advantage of a 2D network with the capability of addressing 3D spatial consistency issues without a loss of precision [23].

Another use of the automatic segmentation ability of CNNs is the field of *retinal vessel segmentation*. The retinal vessel segmentation is a fundamental part in the diagnosis of eye-related diseases. Both the thick and the thin vessels are important for symptom detection. With the help of DL models it is possible to automate this process. Usually an attempt is made to segment both vessel types simultaneously. For example, Yan et al. [24] proposed a method in which they segment thick and thin vessels separately. Zang et al. [25] proposed an accelerated matrix for the same task.

## IV. DISCUSSION

In summary we found that deep learning offers unique capabilities and breakthroughs in identifying, classifying and segmenting different kinds of medical images, especially related to cancer in the breast, lung, and brain.

## V. LIMITATIONS AND FUTURE WORK

### A. Limitations

Some research publications with interesting material is potentially excluded because the analysis was limited to peer-reviewed publications with completed research work. This excluded material could potentially lead to a publication bias.

### B. Future Work

In future work we will systematically compare ML methods in experimental laboratory settings, including the evaluation of mental concepts such as cognitive workload [26, 27], concentration [28], and personality [29, 30] in multi-agent-settings [31-34]. We will therefore triangulate objective and perceived user-oriented concepts [35-37] using physiological data (i.e., electroencephalographic data and spectra [38-39], electrocardiographic data [40], electrodermal activity [41], eye fixation [42, 43], eye pupil diameter [44-46], facial data [47]), and evaluate user acceptance [48-52], trust [53, 54] in multi-agent settings [55-62].

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## REFERENCES

- [1] Editors, "Looking Back on the Millennium in Medicine," N. Engl. J. Med., vol. 342, no. 1, pp. 42-49 2000.
- [2] W. Zhu, "Deep Learning for Automated Medical Image Analysis," Ph.D. dissertation, Univ. of California, Irvine, 2019.
- [3] E. S. Kumar and C. S. Bindu, "Medical Image Analysis Using Deep Learning: A Systematic Literature Review," in Emerging Technologies in Computer Engineering - Microservices in Big Data Analytics (ICETCE), A. K. Somani et al., vol. 985, pp. 81-97, 2019.
- [4] D. Ardila, A. P. Kiraly, S. Bharadwaj, B. Choi, J. J. Reicher, L. Peng et al., "End-to-end lung cancer screening with three-dimensional deep

- learning on low-dose chest computed tomography," *Nature Medicine*, vol. 25, pp. 954-961, 2019.
- [5] E. Smistad, A. Ostvik, and A. Pedersen, "High Performance Neural Network Inference, Streaming, and Visualization of Medical Images Using FAST," *IEEE Access*, vol. 7, pp. 136310-136321, 2019.
  - [6] N. Brancati, G. de Pietro, M. Frucci, and D. Riccio, "A Deep Learning Approach for Breast Invasive Ductal Carcinoma Detection and Lymphoma Multi-Classification in Histological Images," *IEEE Access. Special Section on Deep Learning for Comput.-Aided Med. Diagnosis*, vol. 7, pp. 44709-44720, 2019.
  - [7] C. T. Sari and C. Gunduz-Demir, "Unsupervised Feature Extraction via Deep Learning for Histopathological Classification of Colon Tissue Images," *IEEE Transactions on Med. Imaging*, vol. 38, no. 5, pp. 1139-1149, 2019.
  - [8] T. Qaiser, Y.-W. Tsang, D. Epstein, and N. Rajpoot, "Tumor Segmentation in Whole Slide Images Using Persistent Homology and Deep Convolutional Features," in *Medical Image Understanding and Analysis (MIUA)*, M. V. Hernandez and V. Gonzalez-Castro, vol. 723, pp. 320-329, 2017.
  - [9] H. Lin, H. Chen, S. Graham, Q. Dou, N. Rajpoot and P.-A. Heng, "Fast ScanNet: Fast and Dense Analysis of Multi-Gigapixel Whole-Slide Images for Cancer Metastasis Detection," *IEEE Transactions on Med. Imaging*, vol. 38, no. 8, pp. 1948-1958, 2019.
  - [10] G. Yunchao and S. Faqiang, "Data Acquisition and Processing of Breast Cancer Assisted Diagnosis Based on Ultrasound Imaging," in *Proc. ICBDE*, pp. 82-97, 2019.
  - [11] J. Vizcarra, R. Place, L. Tong, D. Gutman, and M. D. Wang, "Fusion in Breast Cancer Histology Classification," in *Proc. BCB*, pp. 485-493, 2019.
  - [12] X. Zhao, X. Wang, and H. Wang, "Classification of Benign and Malignant Breast Mass in Digital Mammograms with Convolutional Neural Networks" in *Proc. ISICDM*, pp. 47-50, 2018.
  - [13] M. Anthimopoulos, S. Christodoulidis, L. Ebner, A. Christe, and S. Mougiakakou, "Lung Pattern Classification for Interstitial Lung Diseases Using a Deep Convolutional Neural Network," *IEEE Transactions on Med. Imaging*, vol. 35, no. 5, pp. 1207-1216, 2016.
  - [14] I. Cardoso, E. Almeida, H. Allende-Cid, A. C. Frery, R. M. Rangayyan, P. M. Azevedo-Marques et al., "Evaluation of Deep Feedforward Neural Networks for Classification of Diffuse Lung Diseases," in *Pattern Recognition, Image Analysis, Computer Vision, and Applications (CIARP)*, M. Mendoza and S. Velastin, vol. 10657, pp. 152-159, 2017.
  - [15] K. Pawelczyk, M. Kawulok, J. Nalepa, M. P. Hayball, S. J. McQuaid, V. Prakash et al., "Towards Detecting High-Uptake Lesions from Lung CT Scans Using Deep Learning," in *Image Analysis and Processing (ICIAP)*, S. Battiato et al., vol. 10485, pp. 310-320, 2017.
  - [16] S. C. van de Leemput, M. Meijs, A. Patel, F. J. A. Meijer, B. van Ginneken, and R. Manniesing, "Multiclass Brain Tissue Segmentation in 4D CT Using Convolutional Neural Networks," *IEEE Access*, vol. 7, pp. 51557-51569, 2019.
  - [17] M. R. Ahmed, Y. Zhang, Z. Feng, B. Lo, O. T. Inan, and H. Liao, "Neuroimaging and Machine Learning for Dementia Diagnosis: Recent Advancements and Future Prospects," *IEEE Reviews in Biomed. Eng.*, vol. 12, pp. 19-33, 2019.
  - [18] C. Raynaud, C. Ciofalo-Veit, T. Lefèvre, R. Ardon, A. Cavallaro, I. Salim et al., "Multi-organ Detection in 3D Fetal Ultrasound with Machine Learning," in *Fetal, Infant and Ophthalmic Medical Image Analysis (OMIA, FIFI)*, M. J. Cardoso et al., vol. 10554, pp. 62-72, 2017.
  - [19] C. Li, H. Chen, L. Zhang, N. Xu, D. Xue, Z. Hu et al., "Cervical Histopathology Image Classification Using Multilayer Hidden Conditional Random Fields and Weakly Supervised Learning," *IEEE Access*, vol. 7, pp. 90378-90397, 2019.
  - [20] W. Du, Nini Rao, D. Liu, H. Jiang, C. Luo, Z. Li et al., "Review on the Applications of Deep Learning in the Analysis of Gastrointestinal Endoscopy Images," *IEEE Access*, vol. 7, pp. 142053-142069, 2019.
  - [21] R. Mufidah, I. Wasito, N. Hanifah, M. Faturrahman, and F. D. Ghaisani, "Automatic Nucleus Detection of Pap Smear Images using Stacked Sparse Autoencoder (SSAE)," *ICACS*, pp. 9-13, 2017.
  - [22] N. Lessmann, B. van Ginneken, M. Zreik, P. A. de Jong, B. D. de Vos, M. A. Viergever et al., "Automatic Calcium Scoring in Low-Dose Chest CT Using Deep Neural Networks With Dilated Convolutions," *IEEE Transactions on Med. Imaging*, vol. 37, no. 2, pp. 615-625, 2018.
  - [23] J. Duan, G. Bello, J. Schlemper, W. Bai, T. J. W. Dawes, C. Biffi, et al., "Automatic 3D Bi-Ventricular Segmentation of Cardiac Images by a Shape-Refined Multi-Task Deep Learning Approach," *IEEE Transactions on Med. Imaging*, vol. 38, No. 9, pp. 2151-2164, 2019.
  - [24] Z. Yan, X. Yang, and K.-T. Cheng, "A Three-Stage Deep Learning Model for Accurate Retinal Vessel Segmentation," *IEEE J. of Biomed. and Health Inform.*, vol. 23, no. 4, pp. 1427-1436, 2019.
  - [25] Y. Zhang, J. Lian, L. Rong, W. Jia, C. Li, and Y. Zheng, "Even faster retinal vessel segmentation via accelerated singular value decomposition," *Neural Computing and Appl.*, vol. 32, pp. 1893-1902, 2020.
  - [26] R. Buettner, "Investigation of the Relationship Between Visual Website Complexity and Users' Mental Workload: A NeuroIS Perspective", in *Information Systems and Neuro Science*, vol. 10 of LNISO, pp. 123-128, 2015.
  - [27] R. Buettner, S. Sauer, C. Maier, and A. Eckhardt, "Real-time Prediction of User Performance based on Pupillary Assessment via Eye Tracking", *AIS Trans Hum.-Comput Interact*, vol. 10, no. 1, pp. 26-56, 2018.
  - [28] R. Buettner, H. Baumgartl, and D. Sauter, "Microsaccades as a Predictor of a User's Level of Concentration," ser. LNISO, vol. 29, pp. 173-177, 2018.
  - [29] R. Buettner, "Innovative Personality-based Digital Services," in *PACIS '16 Proc.*, paper 278, 2016.
  - [30] R. Buettner, "Personality as a predictor of business social media usage: An empirical investigation of XING usage patterns," in *PACIS '16 Proc.*, paper 163, 2016.
  - [31] R. Buettner, "A Classification Structure for Automated Negotiations," in *IEEE/WIC/ACM WI-IAT 2006 Proc.*, pp. 523-530, 2006.
  - [32] R. Buettner and S. Kirn, "Bargaining Power in Electronic Negotiations: A Bilateral Negotiation Mechanism," in *EC-Web '08 Proc.*, ser. LNCS, vol. 5183, pp. 92-101, 2008.
  - [33] R. Buettner, "Cooperation in Hunting and Food-sharing: A Two-Player Bio-inspired Trust Model," in *BIONETICS '09 Proc.*, Avignon, France, December 9-11, 2009, pp. 1-10, 2009.
  - [34] J. Landes and R. Buettner, "Argumentation-Based Negotiation? Negotiation-Based Argumentation!" in *EC-Web '12 Proc.*, pp. 149-162, 2012.
  - [35] R. Buettner, "Asking both the User's Brain and its Owner using Subjective and Objective Psychophysiological NeuroIS Instruments," in *ICIS 2017 Proceedings*, 2017.
  - [36] R. Buettner, "Getting a job via career-oriented social networking markets: The weakness of too many ties," *Electronic Markets*, vol. 27, no. 4, pp. 371-385, 2017.
  - [37] R. Buettner, "Predicting user behavior in electronic markets based on personality-mining in large online social networks: A personalitybased product recommender framework," *Electronic Markets*, vol. 27, no. 3, pp. 247-265, 2017.
  - [38] D. Raab, H. Baumgartl, and R. Buettner, "Machine Learning based Diagnosis of Binge Eating Disorder using EEG recordings," in *PACIS 2020 Proc.*, 2020, in press.
  - [39] H. Baumgartl, F. Dikici, D. Sauter, and R. Buettner, "Detecting Antisocial Personality Disorder using a Novel Machine Learning Algorithm based on Electroencephalographic Data," in *PACIS 2020 Proc.*, 2020, in press.
  - [40] R. Buettner and M. Schunter, "Efficient machine learning based detection of heart disease," in *IEEE Healthcom 2019 Proc.*, pp. 1-6, 2019.
  - [41] A. Eckhardt, C. Maier, and R. Buettner, "The Influence of Pressure to Perform and Experience on Changing Perceptions and User Performance: A Multi-Method Experimental Analysis," in *ICIS '12 Proc.*, 2012.
  - [42] R. Buettner, "Cognitive Workload of Humans Using Artificial Intelligence Systems: Towards Objective Measurement Applying Eye-

- Tracking Technology,” in *KI 2013 Proc.*, ser. LNAI, vol. 8077, pp. 37-48, 2013.
- [43] A. Eckhardt, C. Maier, J. P.-A. Hsieh, T. Chuk, A. B. Chan, A. B. Hsiao, and R. Buettner, “Objective measures of IS usage behavior under conditions of experience and pressure using eye fixation data,” in *ICIS '13 Proc.*, 2013.
  - [44] R. Buettner, “Social inclusion in eParticipation and eGovernment solutions: A systematic laboratory-experimental approach using objective psychophysiological measures,” in *EGOV/ePart 2013 Proc.*, ser. LNI, vol. P-221, pp. 260-261, 2013.
  - [45] R. Buettner, B. Daxenberger, A. Eckhardt, and C. Maier, “Cognitive Workload Induced by Information Systems: Introducing an Objective Way of Measuring based on Pupillary Diameter Responses,” in *Pre-ICIS HCI/MIS 2013 Proc.*, paper 20, 2013.
  - [46] R. Buettner, S. Sauer, C. Maier, and A. Eckhardt, “Towards ex ante Prediction of User Performance: A novel NeuroIS Methodology based on Real-Time Measurement of Mental Effort,” in *HICSS-48 Proc.*, pp. 533-542, 2015.
  - [47] R. Buettner, “Robust user identification based on facial action units unaffected by users’ emotions,” in *HICSS-51 Proc.*, pp. 265-273, 2018.
  - [48] R. Buettner, B. Daxenberger, and C. Woesle, “User acceptance in different electronic negotiation systems - a comparative approach,” in *ICEBE '13 Proc.*, pp. 1-8, 2013.
  - [49] R. Buettner, “Towards a New Personal Information Technology Acceptance Model: Conceptualization and Empirical Evidence from a Bring Your Own Device Dataset,” in *AMCIS '15 Proc.*, 2015.
  - [50] R. Buettner, “Analyzing the Problem of Employee Internal Social Network Site Avoidance: Are Users Resistant due to their Privacy Concerns?” in *HICSS-48 Proc.*, pp. 1819-1828, 2015.
  - [51] R. Buettner, “Getting a Job via Career-oriented Social Networking Sites: The Weakness of Ties,” in *HICSS-49 Proc.*, pp. 2156-2165, 2016.
  - [52] R. Buettner, “Online user behavior and digital footprints,” Trier, Germany, 2019.
  - [53] F. Meixner and R. Buettner, “Trust as an Integral Part for Success of Cloud Computing,” in *ICIW 2012 Proc.*, pp. 207-214, 2012.
  - [54] R. Buettner, R., “The impact of trust in consumer protection on internet shopping behavior: An empirical study using a large official dataset from the European Union,” in *BDS-2020 Proc.*, 2020, in press.
  - [55] R. Buettner, “The State of the Art in Automated Negotiation Models of the Behavior and Information Perspective,” *ITSSA*, vol. 1, no. 4, pp. 351-356, 2006.
  - [56] R. Buettner, “Electronic Negotiations of the Transactional Costs Perspective,” in *IADIS'07 Proc.*, vol. 2, pp. 99-105, 2007.
  - [57] R. Buettner, “Imperfect Information in Electronic Negotiations: An Empirical Study,” in *IADIS'07 Proc.*, vol. 2, pp. 116-121, 2007.
  - [58] J. Landes and R. Buettner, “Job Allocation in a Temporary Employment Agency via Multi-dimensional Price VCG Auctions Using a Multi-agent System,” in *MICAI '11 Proc.*, pp. 182-187, 2011.
  - [59] R. Buettner and J. Landes, “Web Service-based Applications for Electronic Labor Markets: A Multi-dimensional Price VCG Auction with Individual Utilities,” in *ICIW 2012 Proc.*, pp. 168-177, 2012.
  - [60] R. Buettner, “A Systematic Literature Review of Crowdsourcing Research from a Human Resource Management Perspective,” in *HICSS-48 Proc.*, pp. 4609-4618, 2015.
  - [61] R. Buettner, “Automatisierte Verhandlungen in Multi-Agenten-Systemen: Entwurf eines argumentationsbasierten Mechanismus für nur imperfekt beschreibbare Verhandlungsgegenstände,” Springer: Berlin, 2010.
  - [62] S. Rodermund, R. Buettner, and I. J. Timm, “Towards Simulation-based Preplanning for Experimental Analysis of Nudging,” in *WI-2020 Proc.*, 2020.