scientific reports

Check for updates

Lesion detection in women breast's OPEN dynamic contrast‑enhanced magnetic resonance imaging using deep learning

Sudarshan Saikia1 , Tapas Si2 , Darpan Deb3 , Kangkana Bora4 , Saurav Mallik5 , Ujjwal Maulik6 & Zhongming Zhao7*

Breast cancer is one of the most common cancers in women and the second foremost cause of cancer death in women after lung cancer. Recent technological advances in breast cancer treatment ofer hope to millions of women in the world. Segmentation of the breast's Dynamic Contrast-Enhanced Magnetic Resonance Imaging (DCE-MRI) is one of the necessary tasks in the diagnosis and detection of breast cancer. Currently, a popular deep learning model, U-Net is extensively used in biomedical image segmentation. This article aims to advance the state of the art and conduct a more in-depth analysis with a focus on the use of various U-Net models in lesion detection in women's breast DCE-MRI. In this article, we perform an empirical study of the efectiveness and efciency of U-Net and its derived deep learning models including ResUNet, Dense UNet, DUNet, Attention U-Net, UNet++, MultiResUNet, RAUNet, Inception U-Net and U-Net GAN for lesion detection in breast DCE-MRI. All the models are applied to the benchmarked 100 Sagittal T2-Weighted fat-suppressed DCE-MRI slices of 20 patients and their performance is compared. Also, a comparative study has been conducted with V-Net, W-Net, and DeepLabV3+. Non-parametric statistical test Wilcoxon Signed Rank Test is used to analyze the signifcance of the quantitative results. Furthermore, Multi-Criteria Decision Analysis (MCDA) is used to evaluate overall performance focused on accuracy, precision, sensitivity, F1-score, specifcity, Geometric-Mean, DSC, and false-positive rate. The RAUNet segmentation model achieved a high accuracy of 99.76%, sensitivity of 85.04%, precision of 90.21%, and Dice Similarity Coefcient (DSC) of 85.04% whereas ResNet achieved 99.62% accuracy, 62.26% sensitivity, 99.56% precision, and 72.86% DSC. ResUNet is found to be the most efective model based on MCDA. On the other hand, U-Net GAN takes the least computational time to perform the segmentation task. Both quantitative and qualitative results demonstrate that the ResNet model performs better than other models in segmenting the images and lesion detection, though computational time in achieving the objectives varies.

With the increase in the number of breast cancer cases, the feld of medical science heavily depends upon its early detection for efective diagnosis and treatment in its early stages, but such a task has proven to be challenging. In the year 2020, over 2.3 million women were diagnosed with breast tumors and approximately 685,000 did not survive worldwide, as reported by the World Cancer Research Fund International and the World Health Organization. Breast cancer is the most frequent tumour found in women making it the most likely cancerous disease found in women. Breast cancer has slowly proven to be one out of the four most common cancers found in women. Globally, the cases have gone up by 20% and death rates have seen an increase of 14% since 2008¹.

¹Information Technology Department, Oil India Limited, Duliajan, Assam 786602, India. ²AI Innovation Lab, Department of Computer Science & Engineering, University of Engineering & Management, Jaipur, GURUKUL, Jaipur, Rajasthan 303807, India. ³Department of Computer Application, Christ University, Bengaluru 560029, India. ⁴ Department of Computer Science and Information Technology, Cotton University, Guwahati, Assam 781001, India. ⁵Department of Environmental Health, Harvard T. H. Chan School of Public Health, Boston, MA 02115, USA. ⁶Department of Computer Science and Engineering, Jadavpur University, Kolkata, India. ⁷Center for Precision Health, School of Biomedical Informatics, The University of Texas Health Science Center at Houston, Houston, TX 77030, USA. [⊠]email: zhongming.zhao@uth.tmc.edu

Further study in this feld of discipline can potentially make huge breakthroughs and may take the diagnosis and early detection process to a whole new dimension.

With the rapid advances in bio-technologies and medical technologies, the main progress has been made in cancer detection, diagnosis, and treatment. The best way of diagnosing breast cancer is through medical imaging tests. Medical imaging techniques used for diagnosis include ultrasound², digital mammography³, magnetic resonance imaging⁴, microscopic slices⁵, and infrared thermograms^{[6](#page-17-5)}. As a method of assisting radiologists and doctors in recognizing issues, these modalities produce images that have decreased mortality rates by 30–70%⁷. Information technology is needed to speed up and increase the accuracy of diagnosis as well as provide a second opinion to the expert because picture interpretation is operator-dependent and requires ability^{[8](#page-17-7)}. Physicians can utilize computer-aided diagnosis (CAD) systems, which use computerized characteristics extraction and classifcation algorithms, to quickly identify anomalies.

It required a lot of efort to develop CAD systems based on developments in digital artifcial intelligence, image processing, and pattern recognition. According to predictions, CAD systems will increase diagnostic rates, decrease operator dependence, and reduce the cost of medical auxiliary modalities⁹⁻¹¹. Since false positive rates can result in inefficient therapy as well as psychological, physical, and monetary costs, it may help to reduce them. Additionally, it can stop false negative results that might result in skipped treatments or remissions. Detection sensitivity without CAD is reportedly around 80%, and sensitivity with it is reportedly over 90%¹². The results showed that mammography combined with CAD provided 100% sensitivity for malignancies presenting as microcalcifcations and 86% sensitivity for other mammographic cancer presentations. Consequently, CAD has become the most productive field of study in medical imaging for enhancing diagnosis accuracy^{13,14}.

The foundational effect can be witnessed in the determination of breast cancer by examining breast Magnetic Resonance Imaging (MRI) data. It has an edge over other imaging diagnosis techniques due to its high responsiveness and no harmful emission⁴. MRI is subjected to imaging irregularities in the form of motion artifacts, noise, etc., which obstructs the segmentation process and ROI detection results are less accurate. In order to get accurate outputs, image registration positively impacts by providing non-linear ordered images and a crucial step in Dynamic Contrast-Enhanced MRI (DCE-MRI) based breast tumor diagnosis¹⁵. Apart from classical image segmentation algorithms, a number of machine learning (ML) techniques have contributed to automating the segmentation process with minimum human input, some of which are mentioned in the Related Work section and are helpful for the determination of cancer lesions. The development of the Computer-Aided Diagnosis (CAD) system which is rooted in modern digital visualization, pattern identifcation, and artifcial intelligence, has helped to reduce flawed output, which results in pointless reception and wrong therapy. The most commonly used machine learning techniques are: Support Vector Machine (SVM) and Artifcial Neural Network (ANN[\)16](#page-17-14). Additional eforts have been made in building hybrid systems in order to obtain better sensitivity and accuracy. These include the frequently used CAD systems and some open-source applications that are included in a framework where the diferent models interact with each other for the automatic determination of cancerous lesions.

Deep neural networks are now trendy machine learning designs around numerous disciplines and are extensively stationed in research and related enterprises. In the medical visualization feld, deep learning is mostly applied in the form of convolutional neural networks (CNNs) and its use in clinical practice has been rapidly growing over the years in order to refne the methodology of cancer identifcation. Deep learning has a huge list of applications in medical science. Their successful application to various datasets not only improves accuracy and precision in the study results but also signifcantly reduces the intervention of humans in a subjective manner. Radiotherapy, PET-MRI attenuation correction, and image registration are some of the deep learning applications in the medical imaging discipline. In the case of MRIs, deep learning is usually centered on fractionalization and categorization of rejuvenated intensity images¹⁷

U-shaped Deep CNN model, namely U-Net¹⁸ is commonly implemented in biomedical image segmentation. One main advantage of U-Net is that it can be trained with a few images. Piantadosi et al.^{[19](#page-17-17)} proposed a Three Time Points U-Net (3TP U-Net) for lesion segmentation in breast DCE-MRI and this proposal was evaluated using exclusive metadata consisting of coronal T1-weighted FLASH 3D breast DCE-MRI data. Afer reviewing rigorous existing literature, we found that there is no deep investigation of diferent U-Net models in breast lesion detection using DCE-MRI. This research intends to advance the state of the art and conduct deeper analysis regarding the application of U-Net models in lesion detection in women's breast DCE-MRI. For this, U-Net models are selected: UNet, Dense UNet Attention UNet, UNet++, MultiResUNet, RAUNet, Inception U-Net, U-Net GAN. The models are evaluated on 100 slices of 20 women's breast 2D Sagittal T2-Weighted fat-suppressed DCE-MRI. For comparative study purposes, V-Net, W-Net, and DeepLabV3+ are also taken into account in the experiment to figure out if U-Net models are capable of overcoming them. The effectiveness of the reviewed approaches was evaluated using seven performance metrics: accuracy, sensitivity, specifcity, precision, geometric mean, F-measure, and false-positive rate. To demonstrate the efectiveness of U-Net models, the acquired results are analyzed using the non-parametric statistical test using Wilcoxon Sign Rank Test $(WSRT)^{20}$, and are also further examined using the MCDA approach namely Technique for Order of Preference by Similarity to Ideal Solution $(TOPSIS)^{21}$ $(TOPSIS)^{21}$ $(TOPSIS)^{21}$ to find out the best model.

Contributions of this article

The following is a summary of the contributions:

- 1. Comprehensive investigations evaluating several U-Net models in lesion detection in women breast DCE-MRI.
- 2. A set of important metrics, such as accuracy, sensitivity, specificity, precision, F_1 -score, geometric-mean, DSC, and False-Positive Rate (FPR), were utilized to analyze the results of experiments with 100 slices of women's breast DCE-MRI.

- 3. The performances of different U-Net models are compared with the V-Net, W-Net, and DeepLabV3+ in this study.
- 4. Discussion of both quantitative and qualitative outcomes using statistical and multi-criteria decision-making techniques.

Related works

So far, several mechanisms (Classical methods, Machine Learning methods, Optimization techniques, Fuzzy systems, and Hybrid systems) have been proposed for Breast MRI segmentation and lesion classifcation. Related works are discussed here.

Classical image segmentation techniques

Priya et al.²² designed a biomedical surveillance apparatus 'LabVIEW' to pinpoint the accurate location of cancerous lumps from MR and CT scan images. Recognition of cancerous cells is carried on with the help of the Watershed Algorithm. To explore the practicality of registering ways delicate to time-to-peak (T(peak)) non-uniformity in order to check cancerous cells on breast DCE-MRI, a Time-To-Peak Optimization analysis was done by Fang et al.²³, where the authors concluded distribution of T(peak) optimization can applied in judgment of diagnostic achievement, and be used as standard delicate to intra-lesion T(peak) non-uniformity. Si and Mukhopadhyay²⁴ developed a breast DCE-MRI segmentation method using modified hard-clustering with Fireworks Algorithm (FWA) for lesion detection. Denoising of the MR images is completed using an anisotropic diffusion filter, while intensity inhomogeneities (IIHs) are corrected using the max filter-based method. The authors carried out the segmentation step using a hard-clustering technique with the FWA algorithm. Also, Si and Kar²⁵ proposed a sectionalization technique using an altered hard-clustering technique with a multi-verse optimizer (MVO) for determining breast lesions in DCE-MRI. Preprocessing steps are similar to that of earlier work, then clustering technique is used for segmentation purposes. Finally, in the post-processing step, lesions are extracted. Patra et al.[26](#page-17-24) proposed a multi-level thresholding using Student Psychology-Based Optimizer (SPBO) for lesion detection and segmentation of DCE-MR images. Denoising of the breast MRIs was carried out with an Anisotropic diffusion filter followed up with Intensity Inhomogeneities correction. The processed images underwent segmentation assisted by the SPBO algorithm. It recorded an accuracy level of 99.44%, sensitivity of 96.84%, and DSC of 93.41%.

Statistical Image Segmentation Techniques

A Markov Random Field (MRF) model and dissection approach rooted in the Bayesian theory of maximizing the posterior probability given by Wu et al.^{[27](#page-17-25)} had an edge over and proved to be powerful in image accession. Furthermore, a new MRF model developed by Azmi et al[.28](#page-17-26), resolved the issue of complexity of computation by erasing out the need to use a monotonous approach and yielded Area Under the Curve (AUC) = 0.9724 and Mean Accuracy = 93.10%.

Deep learning based segmentation techniques

A comparative study of Conventional and Multi-State Cellular Neural Networks was performed by Ertas et al.[29](#page-17-27) for breast region segmentation. The results of segmentation achieved by Conventional CNNs : ($PR = 85.5 \pm 17.0\%$, TPVF = $90.4 \pm 17.8\%$ and FPVF = $2.0 \pm 2.2\%$) and Multi-State CNNs: (PR = $99.3 \pm 1.8\%$, TPVF = $99.5 \pm 1.3\%$ and FPVF = 0.1 ± 0.2 %). A novel technique (3D-LESH) was developed by Summrina et al.³⁰ to determine breast lesions in volumetric medical images. Teir results show that 3D-LESH helps in detecting various cancer stages. A CAD system designed by Rasti et al.³¹ is derived from a mixture ensemble of CNN (ME-CNN) to distinguish cancerous and non-cancerous breast tumors. In that study, the dataset consisted of T1-weighted axial image metadata over the entire breast region obtained with the image size of 512×512 pixels. The ME-CNN design had three CNN experts and a convolutional gating network which recorded a high validity of 96.39%, a responsiveness of 97.73%, and a specifcity of 94.87%. Another CNN-based method developed by Xu et al[.32](#page-17-30) was used to slice the mammary gland region in transverse fat-suppressed breast DCE-MRI. They used 50 3-D fatsuppressed DCE-MRI series of datasets. The DSC was found to be 97.44%, DDC recorded 5.11%, and the distance error (automated fragmentation comparison with manual one) was about 1.25 pixels. Zhang et al.³³ proposed an upgraded nine-layer CNN on a dataset of uni-breast mammogram images (1024 \times 1024) which resulted in the correctness of disease = 93.4%, correctness without a disease = 94.6%, exactness of 94.5%, and accuracy of 94.0%. A suitably modifed CNN to fully automatize the signifcant mammary gland cell dissection job in 3D MR data (3D coronal DCE T1-weighted images), was given by Piantadosi et al.^{[34](#page-17-32)}. Mask-guided hierarchical learning (MHL) structure was developed by Zhang et al.³⁵ for breast lump dissection via fully convolutional networks (FCN). An architecture rooted in U-net fCNN was introduced by Benjelloun et al.³⁶, who worked on the idea of an upgraded Fully CNN (fCNN) by including regular fCNN layers and ensuing up-sampling ones to surge the image size and integrating earlier feature maps for a better image representation learning. Deep Learning assisted Efficient Adaboost Algorithm (DLA-EABA) for breast tumor determination was put forward by Zheng et al.^{[37](#page-17-35)} with modern computational methods recorded a high accuracy level of 97.2%, Sensitivity 98.3%, and Specifcity 96.5%. Negi et al.[38](#page-18-0) put forward a Generative Adversarial Network (GAN) based model operating on Breast Ultrasound (BUS) Images which consists of two modules: Residual-Dilated-Attention-Gate-UNet (RDAU-NET), which serves as dissolution block, and a CNN, which acts as distinguisher. Tis hybrid model is also named as WGAN-RDA-UNET. The overall Accuracy, PR-AUC, ROC-AUC and F1-score were achieved 0.98, 0.95, 0.89, and 0.88, respectively. A novel attention-guided joint-phase-learning network for multilabel segmentation method was developed by Qiao et al.^{[39](#page-18-1)}, which produced a Dice Coefficient of 0.83. Zhang et al.^{[40](#page-18-2)} developed a deep learning approach using Mask Regional-CNN for automatic determination of breast lesions in DCE-MRI datasets (non-fat-sat sequence and fat-sat sequence). The tumor is segmented using a fuzzy c-means clustering algorithm and then Mask R-CNN is implemented. Their results show high accuracy in detecting and localizing the tumor.

An ailing administered deep learning technique for breast MRIs was evaluated in a study given by Liu et al.^{[41](#page-18-3)}. The evaluation was done in the absence of pixel-level segmentation in order to achieve higher specificity in breast lesion classification. The model achieved an AUC of 0.92 (SD \pm 0.03), accuracy of 94.2% (SD \pm 3.4), specificity of 95.3% (SD \pm 3.3), and sensitivity of 74.4% (SD \pm 8.5). Huo et al.^{[42](#page-18-4)} presented a study to achieve higher accuracy and efficiency in the segmentation of whole breast in 3-D fat-suppressed DCE-MRI by introducing an adaptable deep learning framework using nnU-Net. Results show its robustness in achieving higher accuracy and prove to be a potential asset in clinical workfow in quantifying breast cancer risk. To surmount the weakness of regular imaging techniques used in machine learning-based approaches, such as limited data size and less information to feed them, Venkata and Lingamgunta^{[43](#page-18-5)} introduced a CNN (CNN (LeNet-5)) based diagnosis of breast using Zenker moments which achieved 88.2% sensitivity and 76.92% accurateness, 83.3% sensitivity and 62.5% malignant growth accuracy. Jaglan et al.⁴⁴ developed a one-ordered algorithm to distinguish breast lesions (normal/ abnormal), which consists of an integrated fning technique for de-noising, breast boundary region extraction via selection of nipple and mid-sternum points, and followed by morphological operations and hole flling. For classification, an SVM was implemented in their study. The proposed method recorded an accuracy of 93.7%, sensitivity of 95.6%, and specifcity of 87.2%.

Kim et al.⁴⁵ developed an edge extraction algorithm (eLFA algorithm) over ultrasonic breast images which implements a cRNN-based learning design to distinguish breast lumps among others. The proposed model advances in most scenarios as an unnecessary specifc set of limitations is avoided in detecting line segments and calculates threshold values itself to determine precise line segments masterfully, thereby achieving the highest accuracy of 99.75%. Lv et al[.46](#page-18-8) developed a dynamic mode-based self-supervised dual attention deep clustering network (DADCN) in order to attain detailed segmentation of breast intra-tumor heterogeneity region individually. The graph attention network learns and combines particular representations with features taken out from deep CNN. Soleimani et al[.47](#page-18-9) developed an algorithm that makes capital of Dijkstra's process, and allows it to track the boundary between the pectoralis muscle and breast tissue as well as between the breast and its surroundings. By evaluating the proposed method, the authors reported its robustness and accuracy in nature.

Methods

DCE‑MRI dataset

Total 100 T2-Weighted Sagittal fat-suppressed DCE-MRI 2D slices of 20 women patients were collected from public dataset of The Cancer Genome Atlas Breast Invasive Carcinoma (TCGA-BRCA)^{[48,](#page-18-10)49} in "The Cancer Imaging Archive (TCIA), USA" (<https://www.cancerimagingarchive.net>). The size of all MRI slices is 256 \times 256. The ground truths are prepared using manual segmentation by the expert radiologist, which is considered as the gold standard⁵⁰. The lesion pixels are assigned the true value and all other pixels are assigned false in the ground truth images.

Overview of the work

In this work, lesion segmentation in breast DCE-MRI using V-Net, W-Net, and ten U-Net based deep learning models is proposed. The proposed method has the following steps.

- Step 1: training all the proposed models for lesion segmentation.
- Step 2: testing of the trained models for the test data and the generation of segmentation outputs.
- Step 3: Localize the lesion in MRI.
- The overview of the proposed method is provided in Fig. [1](#page-4-0).

Deep learning for segmentation

Deep learning models are being widely used in image segmentation as they are robust for performing such tasks. In image segmentation, a digital image is broken down into various segments and the targeted region is extracted. Segmentation tasks can be subdivided into two categories: semantic segmentation and instance segmentation. Semantic segmentation involves assigning a class label for every pixel in the given image. In our study, each pixel is divided into two classes. For each pixel, deep learning models need to determine if it forms a lesion on the breast tissue. In our study, V-Net, W-Net, U-Net and its nine variants are included to achieve the objectives.

U‑net for segmentation

In image segmentation based on deep learning approaches U-Net is considered to be an efficient model that works well even with a limited number of training samples. The 'U' shaped architecture of U-Net is symmetrical and has two major sections, which are the contracting network and expanding network. Down-sampling and up-sampling of the image are performed by these networks respectively. The contracting network is constituted by the general convolutional process whereas the expanding network is constituted by transposed 2D convolutional layers. Both down-sampling and up-sampling occur in a layer-wise manner. Tis architecture does not contain any fully connected layers.

The architecture of the classical U-Net model is presented in Fig. [2](#page-4-1) and the interested readers are directed to respective articles of the diferent U-Net models for the network architecture diagrams. . Each of the contracting path and expanding path contains four sections. To link these two paths, there is another section of convolutional layers in the middle. In the contracting path, each section contains three padded 3×3 convolution layers, that maintain the input dimensions. Out of these layers, the middle layer contains twice the number of feature channels than the other two layers. In each section, batch normalization layer followed by a rectifed linear unit is added after each of the three convolution layers. For down-sampling in the end of each section, a 2×2 max

Figure 1. Overview of our proposed method.

Figure 2. Architecture of U-Net. Each box in the diagram represents a multi-channel feature map.

pooling operation is performed with stride 2. In such down-sampling activity, the feature channels get doubled. The transitional section that connects the contracting path and the expanding path contains similar convolution layers. Rectified linear units are added after this section. The sections in both the contracting path and expanding path follow the same feature channel increment pattern. The main purpose of the expanding path is upsampling. At each section of the expanding path, the feature map gets upsampled. This is followed by a 2×2 convolution layer for reducing the number of feature channels by half. Then, a concatenation operation is performed between feature channels and the corresponding feature map from the contracting path. For maintaining the number of feature channels that have exactly the same format as that of the sections in the contracting path, the concatenation operation is followed by three 3×3 padded convolution layers. Each 3×3 convolution layer in each

section of expanding path is followed by batch normalization and a rectified linear unit. A 1×1 convolutional layer is added at the end to decide the class for each feature vector. The complete architecture has a total of 32 convolutional layers. U-Net's architecture, which includes a contracting path for capturing context and a symmetric expanding path for accurate localization, is what gives it its competitive edge. With this layout, accurate segmentation outcomes are possible even with small training data.

We have implemented several other models: ResUNet, Dense UNet Attention UNet, UNet++, MultiResUNet, RAUNet, Inception U-Net, U-Net GAN. Te architectures of these models are based on the derivation of the U-Net for multiple purposes.

ResUNet refers to Deep Residual U-Net. It is a semantic segmentation model based on U-Net and inspired by the idea of deep residual learning. This model was proposed for road area extraction. There are two major advantages of this model. First, the training process of deeper models requires more resources; therefore, residual units used in ResUNet ease the training process. Second, the networks can be designed with fewer parameters, which helps maintain improved performance due to the rich skip connections. Tis also facilitates information propagation.

A densely connected convolutional network strengthens the utilization of features and provides improved segmentation results even with a limited number of training samples. We combine the U-Net network and densely connected convolutional network (commonly called Dense UNet). Between each pair of convolutional layers there are some extra concatenation layers compared to U-Net and other traditional convolutional networks. In Dense UNet, each layer can acquire the feature maps of all its previous layers as inputs while its feature maps can be passed to all successive layers, and without increasing the size of datasets a higher segmentation accuracy can be achieved.

DUNet (Deformable U-Net) was initially proposed for retinal vessel segmentation. In DUNet, some of the convolutional layers of the basic U-Net are replaced with deformable convolution blocks. Upsampling operators are used to increase the output resolution. DUNet performs quite well in precise localization by combining low-level feature maps with high-level ones.

In Attention U-Net, attention gates are attached on top of the standard U-Net. It aims at increasing segmentation accuracy even with lesser training data. Attention gates reduce the computational resources wasted on irrelevant activations by highlighting only the relevant activations during training. This model was first applied to medical image segmentation which eliminates the necessity of applying an external object localisation model.

UNet++ was proposed for image segmentation in the feld of medical science. It is encoder-decoder based network containing encoder and decoder sub-networks. The two sub-networks are connected by a series of nested, dense skip pathways. The re-designed skipping pathways also help to reduce the semantic gap between the feature maps of the encoder and decoder sub-networks.

In MultiResUNet, each pair of convolutional layer pairs in the basic U-Net is replaced with a MultiRes block. This configuration basically is derived from factorizing 5×5 and 7×7 convolution operations to 3×3 ones and reuses them to obtain results from 3×3 , 5×5 and $\overline{7} \times 7$ convolution operations simultaneously. To regulate the semantic distance between Encoder and Decoder networks residual path is added. To proportionate the anticipated gap between two corresponding layers additional convolutions are added along the shortcut path.

RAUNet was initially proposed for the semantic segmentation of Cataract Surgical Instruments. The basic architecture of RAUNet is similar to that of U-Net. By combining both low and high-level feature maps, RAUNet extracts contextual information. To adaptively change the attention-aware features attention residual modules are integrated.

Inception U-Net is inspired by Inception net. Inception modules are introduced into the original U-Net architecture. Tis architecture is also encoder-decoder based. Inception modules are used because of the fact that convolution layers of diferent dimensions can correlate with diferent spatial features present in the same feature map. Inception U-Net follows U-net like architecture. However, here inception modules are used in place of the regular stack of convolution layers as in U-Net.

U-Net GAN uses a segmentation network as the discriminator. Tis segmentation network predicts two classes. In doing so, the discriminator gives the generator region-specifc feedback.

Applications of all the models are mentioned in Table [1](#page-6-0).

V‑net for segmentation

V-Net is a CNN-based deep learning model widely used in the feld of medical image segmentation. Tis model was frst used to detect prostate segments from MRI volumes.

The V-Net network consists of two paths, a compression path and an expansion path. The compression path compresses the signal, while the expansion path decompresses it to its original size. At each stage of the compression path, a residual function is learnt and a convolution operation is performed. At each stage of the expansion path, a deconvolution operation is performed. Similar to the compression path, residual function is learnt at each stage. Horizontal connections help to forward the features learnt in the early stages of the compression path to the expansion path.

W‑net for segmentation

W-Net is an unsupervised image segmentation model consisting of an encoder network and a decoder network. Both the encoder and decoder networks are concatenated together to form an auto-encoder.

W-Net is a W-shaped architecture consisting of 18 modules. Each module contains two 3×3 convolutional layers, one Rectified Linear Unit (ReLU), and batch normalization. The whole network contains 46 convolutional layers. The initial nine modules combined form the encoder unit that facilities the image segmentation, while the decoder unit consists of the later nine modules that form the reconstructed images.

Table 1. Diferent U-Net models and their applications.

Results

In this paper, lesion detection using V-Net, W-Net, and ten U-Net based deep learning models is proposed. A summary of the parameters used in the models and hyperparameters for each model and the experimental environment of Google Colaboratory^{[62](#page-18-13)} is presented in Tables [2](#page-6-1) and 3, respectively.

We compare the effectiveness of V-Net, W-Net, and U-Net based deep learning models on the basis of evaluation metrics: Accuracy, Precision, Sensitivity, F-measure, Specifcity, Geometric Mean (G-mean), DSC, and false positive rate (FPR)^{[63](#page-18-14)}. Let TP, FP, TN, and FN be the True Positive rate, False Positive rate, True Negative rate, and False Negative rate, respectively. Various performance measures used in our study can be calculated as follows:

$$
Accuracy = (TP + TN)/(TP + FN + TN + FP)
$$
\n(1)

Table 2. Summary of all the parameters and hyperparameters used for each model.

Table 3. Experimental environment.

Scientifc Reports | (2023) 13:22555 | https://doi.org/10.1038/s41598-023-48553-z

$$
Precision = TP/(TP + FP)
$$
\n(2)

$$
Sensitivity(recall) = TP/(TP + FN)
$$
\n(3)

$$
F-measure = \frac{2 \times recall \times precision}{recall + precision}
$$
 (4)

$$
Specificity = TN/(TN + FP)
$$
\n(5)

$$
G-mean = \sqrt{Sensitivity \times Specificity}
$$
 (6)

For signifcant evaluation of the lesion segmentation performance, another widely used performance metric known as DSC is used. The formula for calculating DSC is:

$$
DSC(A, B) = \frac{2|A \cap B|}{|A| \cup |B|}
$$
\n⁽⁷⁾

where A and B are representing binary masks for the segmented lesion and ground truth, respectively. DSC indicates the overlapping ratio between the segmented lesion and the ground truth. A higher DSC value indicates better lesion segmentation performance.

The problem in hand in this current work is to classify the pixels of breast MR images into lesions (cancerous tissues) and non-lesions (non-cancerous or healthy tissues). In the ground truth images, the pixels belonging to lesions are labeled as '1' whereas pixels belonging to non-lesions are labeled as '0'. Similarly, in the resultant images during testing of the models, the lesional pixels are assigned to '1' and non-lesional pixels are assigned to '0'. Te loss function used for each model is binary cross-entropy loss. Te equation for the binary cross-entropy loss can be written as follows:

$$
L(Y_i, \hat{Y}_i) = -\frac{1}{N} \sum_{i=1}^{N} Y_i \log(\hat{Y}_i) + (1 - Y_i) \log(1 - \hat{Y}_i)
$$

Here, the number of training data points is defined by N, Y_i is the ground truth label, and \hat{Y}_i is the label predicted by the model.

The optimizer used is Adam. Adam optimizer updates the weight parameters such that loss function gets minimized. Keras's default learning rate (0.01) is used in training. In addition, batch size of 60, 20 and 20 is used for training, validation, and testing, respectively. Each of the models is trained for diferent epochs based on hyperparameter tuning implemented in KerasTuner. Other hyperparameters are selected based on the Random Search⁶⁴ method to achieve the best possible outcome from each trained model. Table [2](#page-6-1) gives a summary of hyperparameters used for the models. All the models are trained on Tesla T4 GPU, with CUDA Version 11.2 and a GPU size of 15109 MiB. GPU confgurations are mentioned in Table [3](#page-6-2).

Afer training each model to the desired number of epochs, all 20 images from the test set are evaluated. For analysis, the mean and standard deviation of diferent performance measures over those 20 results are taken into consideration.

Quantitative results

The performance measures for each model are briefly presented in Table [4](#page-8-0). The box plots (models versus different performance measures) of diferent models over 20 MR images are shown in Figs. [1](#page-4-0), [2,](#page-4-1) [3](#page-9-0), [4](#page-9-1), [5,](#page-9-2) [6](#page-10-0), [7](#page-10-1) and [8](#page-10-2) in order to provide a better understanding of the results.

In Table [4](#page-8-0), it is observed that RAUNet has achieved a higher mean classifcation accuracy than rest of the models. The mean accuracy of RAUNet is 99.76% with standard deviation of 0.00228, whereas mean accuracies the other models namely, U-Net, ResUNet, Dense UNet, DUNet, Attention U-Net, UNet++, MultiResUNet, Inception U-Net, U-Net GAN, V-Net, and W-Net are 99.71%, 99.62%, 99.74%, 99.35%, 99.72%, 99.53%, 99.30%, 99.68%, 99.45, 99.58%, and 99.37%, respectively.

The average precision value of the ResUNet model is 99.56%, which is the highest among all the models. Sensitivity is considered as one of the key measures in image segmentation. The mean sensitivity value of RAUNet is 85.04%, whereas the mean sensitivity values of U-Net, ResUNet, Dense UNet, DUNet, Attention U-Net, UNet++, MultiResUNet, Inception U-Net, U-Net GAN, V-Net, and W-Net are 71.99%, 62.26%, 78.93%, 63.22%, 75.92%, 73.88%, 72.27%, 74.90%, 52.46%, 73.46%, and 79.04% respectively. These values are relatively low. Though mean specifcity value obtained from all the models are very high, ResUNet shows the highest mean specifcity value of 99.99% with a standard deviation of 0.00005.

The F-measure summarizes precision and sensitivity into a single measure. The mean F-measure for the RAUNet model is 85.04% which is higher than F-measure values of U-Net, ResUNet, Dense UNet, DUNet, Attention U-Net, UNet++, MultiResUNet, Inception U-Net, U-Net GAN, V-Net, and W-Net. Since it is difficult to get a detailed view of the performance of the segmentation task from either precision or sensitivity, the usage of F-measure becomes benefcial in this regard. In our study the higher value of F-measure in the case if the RAUNet model indicates that this model is doing a very well job in lesion segmentation.

The RAUNet model has an excellent mean G-mean score of 90.97%, which is also higher than that of U-Net, ResUNet, Dense UNet, DUNet, Attention U-Net, UNet++, MultiResUNet, Inception U-Net, U-Net GAN, V-Net,

Table 4. Mean and standard deviation (in parenthesis) of performance measure values (in %) for all the models for the test set. Bold-faced results indicate better.

Figure 3. The box plot (models versus accuracy) of comprehensive classification performance of different models over 20 MR Images.

Figure 4. The box plot (models versus precision) of comprehensive classification performance of different models over 20 MR Images.

Figure 5. The box plot (models versus sensitivity) of comprehensive classification performance of different models over 20 MR Images.

and W-Net. Since the proportion of healthy tissues is very high compared to lesions in breast MR images, it implies high-class imbalance problem may be present. A high mean G-mean score indicates that the model is capable of solving this problem.

The mean DSC value of RAUNet is 85.04% which is higher than other models. DSC value reflects the extent of overlap of the result with the ground truths of the segmented lesions. A higher DSC value means better performance of the model.

FPR is a measure of how many samples are wrongly identifed as the positive out of all the samples. FPR does not show any responsive nature towards changes in data distribution and, therefore, it can be used even with imbalanced data. The mean FPR of the ResUNet model is 0.0032% which is much lower than that of U-Net, Dense UNet, DUNet, Attention U-Net, UNet++, MultiResUNet, RAUNet, Inception U-Net, U-Net GAN, V-Net, and W-Net. The lower value of FPR in the model of ResUNet indicates that wrongly identified negative samples are minimum in images.

Figure 6. The box plot (models versus f-measure) of comprehensive classification performance of different models over 20 MR Images.

Figure 7. The box plot (models versus specificity) of comprehensive classification performance of different models over 20 MR Images.

Figure 8. The box plot (models versus g-mean) of comprehensive classification performance of different models over 20 MR Images.

The robustness of the models is one of the most essential aspects of the segmentation task. A robust model can be expected to perform well irrespective of fuctuations in the data. It is measured on the basis of the standard deviation of diferent performance measures. Stronger robustness is indicated by lower standard deviation. From the standard deviation results of all the models in Table [3,](#page-6-2) it is difficult to find the most robust model as no model has the lowest standard deviations for all the measurements. Therefore, some statistical analysis methods are adopted to fnd the most robust model.

A boxplot gives a good graphical interpretation of the diversity of values. Using boxplots, visual comparison of performance measures of diferent models can be easily examined. It is observed from Fig. [3](#page-9-0) that the RAUNet has the highest median accuracy of classifcation compared to the other models. It also indicates that, in comparison to other models, the minimum and maximum accuracy values in the case of RAUNet is comparatively higher with the least diference between both the values. In Fig. [4,](#page-9-1) it can be observed that ResUNet has a higher median precision score than that of other models. Similarly, Fig. [5](#page-9-2), shows that RAUNet has a higher median sensitivity

score than the other models. Fig. [7](#page-10-1) displays that ResUNet has a slightly higher median specifcity score compared to other models. RAUNet has the highest median G-Mean, F-Measure, and DSC value, as shown in, Figs. [8](#page-10-2), [6](#page-10-0) and [9](#page-11-0). From Fig. [10,](#page-11-1) it is observed that ResUNet has the minimum median FPR value. All together, as shown in Figs. [3,](#page-9-0) [4,](#page-9-1) [5](#page-9-2), [6,](#page-10-0) [7](#page-10-1), [8,](#page-10-2) [9](#page-11-0) and [10](#page-11-1), it is clear that no model has the highest median value for all the measurements.

Statistical analysis using Wilcoxon Signed Rank Test

The models' effectiveness is assessed using a non-parametric test. The best models based on results in Table [4](#page-8-0) are compared against the other models pair-wise using the Wilcoxon Signed Rank Test (WSRT) with signifcance level (α) through IBM SPSS Statistics 23⁶⁵. The test results based on the accuracy, precision, sensitivity, F-measure, specifcity, G-mean, DSC, and FPR are respectively described in Tables [5](#page-11-2), [6,](#page-12-0) [7,](#page-12-1) [8](#page-12-2), [9](#page-12-3), [10](#page-12-4), [11](#page-13-0) and [12](#page-13-1). From the WSRT results based on accuracy, sensitivity, F-measure, G-mean, and DSC, it is observed that the RAUNet model statistically outperforms other models except MultiResUNet for accuracy, VNet for sensitivity, VNet and Attention U-Net for G-Mean. On the other hand, the ResUNet model statistically outperforms all other models based on precision, specifcity, and FPR.

Figure 9. The box plot (models versus dsc) of comprehensive classification performance of different models over 20 MR Images.

Figure 10. The box plot (models versus fpr) of comprehensive classification performance of different models over 20 MR Images.

Pair	<i>p</i> -value	h	Pair	p-value	h
RAUNet vs. UNet	0.003182		RAUNet vs. MultiResUNet	0.061953	Ω
RAUNet vs. ResUNet	0.000103		RAUNet vs. Inception U-Net	0.001161	
RAUNet vs. Dense UNet	0.040136		RAUNet vs. U-Net GAN	0.008962	
RAUNet vs. DUNet	0.000681		RAUNet vs. V-Net	0.000140	$\mathbf{1}$
RAUNet vs. Attention U-Net	0.013733		RAUNet vs. W-Net	0.000103	
RAUNet vs. U-Net $++$	0.000163		RAUNet vs. DeepLabV3+	0.000089	

Table 5. WSRT results on accuracy.

Table 6. WSRT results on precision.

Table 7. WSRT results on sensitivity.

Table 8. WSRT results on F-measure.

Table 9. WSRT results on specifcity.

Table 10. WSRT results on G-Mean.

Table 11. WSRT results on DSC.

Table 12. WSRT results on FPR.

Multi‑criteria decision analysis (MCDA)

Decision-making with multi-objectives is a difficult task and prone to errors. Multi-criteria decision analysis^{[21](#page-17-19),[66](#page-18-28)} solves the issues of multi objectives up to a large extent and it has been widely used in operations research and management science. MCDA aims to provide the best reasonable solution based on the conditions provided. In practice, problems may be conficting, and any solution may not meet all the criteria simultaneously. From the performance measures in Table [4](#page-8-0) and their statistical analysis in Tables [5](#page-11-2), [6](#page-12-0), [7,](#page-12-1) [8](#page-12-2), [9](#page-12-3), [10](#page-12-4), [11](#page-13-0) and [12,](#page-13-1) it is observed that no single model could achieve the best results for all the performance metrics. Therefore, it is difficult to select the best one. In order to fnd out the best model, we have conducted a multi-criteria decision analysis. In our study, a prominent MCDA method namely Technique for Order of Preference by Similarity to Ideal Solution (TOPSIS) with Information Entropy Weighting Methodology⁶⁷ is used for performance analysis. At present, MCDA is widely used in performance assessment of methods^{26,[68,](#page-18-30)[69](#page-18-31)}. For our study, accuracy, precision, sensitivity, F-measure, specificity, G-mean, DSC, and FPR are used as multiple criteria. There is a conflict between FPR and other criteria. Lower values of FPR indicate better results, which is completely opposite to other criteria. MCDA ranks based on the TOPSIS technique are shown in Table [13.](#page-14-0) From Table [13](#page-14-0), it is observed that the ResUNet model has the top rank followed by U-Net. W-Net has the lowest rank.

Visual results

A test dataset consisting of 20 images has been used for the validation of the models. Due to space constraints, out of the 20 test results, only two images belonging to two patients are presented. The segmented results from each model for patient #1 and patient #2 are displayed in Figs. [11](#page-14-1) and [12](#page-15-0) respectively. Figures [13](#page-15-1) and [14](#page-16-0) show the localized lesions for patients 1 and 2 respectively in a comparative view.

For patient #1, Fig. [11,](#page-14-1) shows U-Net segmented result in Fig. [11](#page-14-1)c and ground truth in Fig. [11](#page-14-1)b are almost similar. The ground truth image and the U-Net segmented image contain 1071 and 935 numbers of lesion pixels, respectively, while the count of overlapping lesion pixels of both images is 931. Tis implies that U-Net is performing well in fully segmenting the lesions. Some parts of the lesions are missing from the image segmented by the ResUNet model as shown in Fig. [11d](#page-14-1). Some dark spots can be visible inside the large lesion in the ResUNet segmented image. As shown in Fig. [11](#page-14-1)e, lesions are well segmented from the Dense UNet segmented image. This model performs well in the detection of the lesions. On the contrary, DUNet did not perform well in segmenting the lesions, as displayed in Fig. [11f](#page-14-1). In addition, Fig. [11](#page-14-1)g–k, demonstrates that Attention U-Net, UNet++, MultiResUNet, RAUNet and Inception U-Net segmented images are mostly similar to the ground truth. Hence, these fve models have good performance in the segmentation of lesions. Finally, from U-Net GAN, and V-Net segmented images in Fig. [11l](#page-14-1),m in comparison with the ground truth image, we fnd their performance satisfactory, while W-Net segmented image in Fig. [11](#page-14-1)n, the performance is not satisfactory.

From Fig. [12,](#page-15-0) much similarity can be observed between the U-Net segmented result in Fig. [12c](#page-15-0) and the ground truth in Fig. [12b](#page-15-0) for patient #2. The ground truth image and the U-Net segmented image contain 516 and 474 numbers of lesion pixels, respectively, while the count of overlapping lesion pixels of both images is 467. The U-Net model is performing well in segmenting the lesions in the image. When we examined ResUNet and Dense UNet segmented images in Fig. [12d](#page-15-0),e, we found that ResUNet and Dense UNet also perform well in segmenting the lesions. In DUNet segmented image in Fig. [12](#page-15-0)f, though lesions are quite well segmented, some healthy tissues are detected as lesions. DUNet fails to properly segment the lesions in the image. Fig. [12](#page-15-0)g, supports that, Attention U-Net has good performance in fully segmenting lesions. From the image segmented by UNet++ as

Table 13. Multi-criteria decision analysis ranks based on TOPSIS technique Information Entropy Weighting Methodology.

Figure 11. Results of diferent models for the data of Patient #1. (**a**) Original MR image. (**b**) Ground truth of MR image. (**c**–**n**) Segmented images using diferent models.

shown in Fig. [12h](#page-15-0), we found that some healthy tissues, which are being detected as lesion. From MultiResUNet, RAUNet, Inception U-Net, and U-Net GAN segmented images in Fig. [12i](#page-15-0)–l, it has been noticed that MultiResU-Net, RAUNet, Inception U-Net and U-Net GAN are properly segmenting the lesions. These four models have achieved decent performance in the detection of lesions. From Fig. [12m](#page-15-0),n, it is observed that the performance of V-Net in segmenting the lesion is quite decent, whereas W-Net completely failed to segment the lesion.

A separate comparative study has been conducted with the increased number of MR images having similar characteristics to the dataset used in the frst experiment, i.e., a total of 200 MR images is used to test the model. In this experiment, ResUNet achieves 99.23% accuracy, 99.34% precision, 62.10% sensitivity, 72.74% F-measure, 99.65% specifcity, 76.08% G-mean, 72.78% DSC, and 0.0035% FPR, which are the best values, and ResUNet retains the frst rank when evaluated using TOPSIS.

Computational complexity

The optimum number of epochs for each model is calculated by hyperparameter tuning using KerasTuner. The computational time of U-Net, ResUNet, Dense UNet, DUNet, Attention U-Net, UNet++, MultiResUNet, RAU-Net, Inception U-Net, U-Net GAN, V-Net, and W-Net took are 1406 s, 503 s, 819 s, 502 s, 936 s, 631 s, 1212 s, 1286 s, 2060 s, 326 s, 1025s, and 621s respectively. They all are in the range of 326-1406 s

U-Net GAN runs faster than all the models. Based on Multi-Criteria Decision Analysis, ResUNet is evaluated as the best model with relatively lower training time compared to U-Net. The experimental results revealed

Figure 12. Results of diferent models for the data of Patient #2. (**a**) Original MR image. (**b**) Ground truth of MR image. (**c**–**n**) Segmented images using diferent models.

DeepLabV3Plus

Figure 13. Localized lesions (bright colored spot) in MR images for Patient #1. (**a**) U-Net. (**b**) ResUnet. (**c**) DenseUnet. (**d**) DUNet. (**e**) Attention U-Net. (**f**) UNet++. (**g**) MultiResUnet. (**h**) RAUNet. (**i**) Inception U-Net. (**j**) U-Net GAN. (**k**) V-Net. (**l**) W-Net (**k**) DeepLabV3Plus.

that, although models are effective in lesion segmentation, efficiency is an issue that may affect the effectiveness of these models.

Discussion

In this current study, we investigated in-depth and analyzed the performance of several U-Net models in women's breast lesion detection using DCE-MRI. Furthermore, the performances of UNet models are also compared with other state-of-the-art deep learning models such as V-Net, W-Net, and DeepLabV3+ models to test their competitiveness. The quantitative results are analyzed using Box-Plots and the non-parametric statistical test method WSRT. Finally, the MCDA method TOPSIS is used to select the best models from thirteen deep learning models and ResUNet achieves the frst rank whereas DUNet holds the last rank among the UNet models. The performance of V-Net is superior to DUNet. The ranks of ResNet, U-Net, MultiResUNet, Attention U-Net, Dense UNet, RAUNet, Inception U-Net, U-Net GAN, and UNet++ are higher than that of V-Net, W-Net, and

Figure 14. Localized lesions (bright colored spot) in MR images for Patient #2. (**a**) U-Net. (**b**) ResUnet. (**c**) DenseUnet. (**d**) DUNet. (**e**) Attention U-Net. (**f**) UNet++. (**g**) MultiResUnet. (**h**) RAUNet. (**i**) Inception U-Net. (**j**) U-Net GAN. (**k**) V-Net. (**l**) W-Net. (**k**) DeepLabV3Plus.

DeepLabV3+. From this observation, it can be concluded that all the U-Net models except DUNet are more efective than V-Net, W-Net, and DeepLabV3+ models in women's breast lesion detection using DCE-MRI. From the qualitative (visual) results in Figs. [11](#page-14-1) and [12,](#page-15-0) it is also observed that all the U-Net models and V-Net can segment the lesions in breast DCE-MRIs. It is worth noticing that W-Net segments the lesions in the MRI of patient-1 (Fig. [11](#page-14-1)n) whereas it cannot do the same for the patient-2 (Fig. [12](#page-15-0)n). DeepLabV3+ is unable to segment the lesions for both patients. Overall, ResNet performs better than all the other models in women's breast lesion segmentation using DCE-MRI and that observation is supported by both quantitative results analyzed by statistical test WSRT and MCDM ranking, and visual analysis of segmented and localized lesion images. ResUNet is based on the ResNet architecture. The use of skip connections helps to deal with the problem of diminishing gradients and allows the training of complex models, which in turn contributes to faster convergence to the solution⁷⁰. The U-Net model works very well with a small number of annotated images¹⁸. So it can perform well even on a small dataset. In the MultiResUNet model, convolutional layers between the encoder and the decoder reduce the semantic gap, and the residual connections make the model easy to train^{[56](#page-18-20)}. In our study, these are the three best-performing models. The encoding and decoding units in the case of DUNet are deformable convolutional blocks. Also, DUNet performed well in retinal vessel segmentation, which was trained on a large patch dataset^{[53](#page-18-17)}.

Apart from the success of most of the U-Net models, there are also some limitations. From the visual results, it is observed that along with the segmentation of lesions, some healthy tissues are also segmented as lesions that are not expected in breast lesion detection. It is also noticed that some lesions are not segmented. This may have occurred due to overftting of the models during the training process and the models learn the noise and intensity inhomogeneities (IIH) present in the MRIs. Overftting occurs due to model complexity and the presence of noise and IIH in MRIs afects the segmentation performance. In this study, the hyper-parameters (e.g., learning rate) are set empirically. The gradient-based optimization algorithm called Adam is used to optimize the network's parameters and gradient-based algorithms generally get stuck in the local minima of the error surface. The breast MRI segmentation problem is a class imbalance problem where the number of pixels corresponding to healthy tissues is very large compared to that of lesions and lesions have very few representations compared to healthy tissues in the training process. The future works can be as follows: (1) model's *hyper-parameters* tuning using Random Search technique⁶⁴ to improve training performance, (2) Instead of gradient-based learning algorithm, metaheuristics can be used in training of the models to overcome the local minima and class imbalance problems[71](#page-18-33), (3) training and evaluating the models on heterogeneous datasets because there are various changes in images because of sensors and other elements since the nature of data varies from hardware to hardware, and (4) use of suitable denoising and intensity inhomogeneities correction techniques as the preprocessing steps in the proposed segmentation model to enhance the segmentation performance.

Data availability

The data-set generated and/or analysed during the current study are available in the The Cancer Genome Atlas Breast Invasive Carcinoma Collection (TCGA-BRCA) repository, [https://wiki.cancerimagingarchive.net/pages/](https://wiki.cancerimagingarchive.net/pages/viewpage.action?pageId=3539225) [viewpage.action?pageId=3539225.](https://wiki.cancerimagingarchive.net/pages/viewpage.action?pageId=3539225)

Received: 31 March 2023; Accepted: 28 November 2023 Published online: 18 December 2023

References

- 1. Breast cancer statistics and resources (2021).
- 2. Boukerroui, D., Basset, O., Guerin, N. & Baskurt, A. Multiresolution texture based adaptive clustering algorithm for breast lesion segmentation. *Eur. J. Ultrasound* **8**, 135–144 (1998).
- 3. Boumaraf, S., Liu, X., Ferkous, C. & Ma, X. A new computer-aided diagnosis system with modifed genetic feature selection for bi-rads classifcation of breast masses in mammograms. *BioMed Res. Int.* **2020**, 7695207 (2020).
- 4. Behrens, S. *et al.* Computer assistance for mr based diagnosis of breast cancer: Present and future challenges. *Comput. Med. Imaging Graphics.* **31**(4–5), 236–47 (2007).
- 5. Abeytunge, S. *et al.* Evaluation of breast tissue with confocal strip-mosaicking microscopy: A test approach emulating pathologylike examination. *J. Biomed. Opt.* **22**, 34002.<https://doi.org/10.1117/1.JBO.22.3.034002>(2017).
- 6. Samantaray, L., Hembram, S. & Panda, R. A new harris hawks-cuckoo search optimizer for multilevel thresholding of thermogram images. *Int. Inf. Eng. Technol. Assoc*. <https://doi.org/10.18280/ria.340503> (2020).
- 7. Schneider, M. & Yafe, M. Better detection: Improving our chances. in *Digital Mammography* (2000).
- 8. Li, H. Computerized radiographic mass detection II. Decision support by featured database visualization and modular neural networks. *IEEE Trans. Med. Imaging.* **20**, 302–313 (2001).
- 9. Leichter, I. *et al.* Optimizing parameters for computer-aided diagnosis of microcalcifcations at mammography. *Acad. Radiol.* **7**, 406–412 (2000).
- 10. Mohanty, A., Senapati, M. & Lenka, S. Retracted article: An improved data mining technique for classifcation and detection of breast cancer from mammograms. *Neural Comput. Appl.* **22**, 303–310 (2013).
- 11. Tang, J. E. A. Computer-aided detection and diagnosis of breast cancer with mammography: Recent advances. *IEEE Trans. Inf. Technol. Biomed.* **13**, 236–251 (2009).
- 12. Horsch, A., Hapfelmeier, A. & Elter, M. Needs assessment for next generation computer-aided mammography reference image databases and evaluation studies. *Int. J. Computer Assisted Radiol. Surg.* **6**, 749 (2011).
- 13. van Ginneken, B. E. A. Comparing and combining algorithms for computer-aided detection of pulmonary nodules in computed tomography scans: The anode09 study. *Med. Image Anal.* 14, 707-722 (2010).
- 14. Doi, K. Diagnostic imaging over the last 50 years: Research and development in medical imaging science and technology. *Phys. Med. Biol.* **51**, 749 (2006).
- 15. Zheng, Y., Wei, B., Liu, H., Xiao, R. & Gee, J. Measuring sparse temporal-variation for accurate registration of dynamic contrastenhanced breast MR images. *Comput. Med. Imaging Graph.* **46**, 73–80 (2015).
- 16. Yassin, N. I., Omran, S., Houby, E. E. & Allam, H. Machine learning techniques for breast cancer computer aided diagnosis using diferent image modalities: A systematic review. *Comput. Methods Programs Biomed.* **156**, 25–45 (2018).
- 17. Lundervold, A. An overview of deep learning in medical imaging focusing on MRI. *Zeitschrif fur medizinische Physik* **29**(2), 102–127 (2019).
- 18. Ronneberger, O., Fischer, P. & Brox, T. U-net: Convolutional networks for biomedical image segmentation. in *Medical Image Computing and Computer-Assisted Intervention—MICCAI 2015*, vol. 9351 of *Lecture Notes in Computer Science* (eds Navab, N., Hornegger, J., Wells, W. & Frangi, A.) (Springer, 2015).
- 19. Piantadosi, G., Marrone, S., Galli, A., Sansone, M. & Sansone, C. DCE-MRI breast lesions segmentation with a 3TP U-net deep convolutional neural network. *2019 IEEE 32nd Int. Symp. on Comput. Med. Syst.* 628–633 (2019).
- 20. Derrac, J., Garcia, S., Molina, D. & Herrera, F. A practical tutorial on the use of nonparametric statistical tests as a methodology for comparing evolutionary and swarm intelligence algorithms. *Swarm Evol. Comput.* **1**, 3–18 (2001).
- 21. Brown, S., Tauler, R. & Walczak, B. *Comprehensive Chemometrics-Chemical and Biochemical Data Analysis* 2nd edn. (Elsevier, 2020).
- 22. Priya M., H. & Krishnaveni, S. LabVIEW implemented breast cancer detection using watershed algorithm. in *2020 5th International Conference on Communication and Electronics Systems (ICCES)*, 1096–1100.<https://doi.org/10.1109/ICCES48766.2020.9137945> (2020).
- 23. Liu, F., Kornecki, A., Shmuilovich, O. & Gelman, N. Optimization of time-to-peak analysis for diferentiating malignant and benign breast lesions with dynamic contrast-enhanced MRI. *Acad. Radiol.* **18**(6), 694–704 (2011).
- 24. Si, T. & Mukhopadhyay, A. Breast DCE-MRI segmentation for lesion detection using clustering with fireworks algorithm. Algo*rithms Intell. Syst. Appl. Artif. Intell. Eng*. https://doi.org/10.1007/978-981-33-4604-8_2 (2021).
- 25. Kar, B. & Si, T. Breast DCE-MRI segmentation for lesion detection using clustering with multi-verse optimization algorithm. *Sof Comput. Teor. Appl. Adv. Intell. Syst. Comput.* **1381**, 265-278. https://doi.org/10.1007/978-981-16-1696-9_25 (2021).
- 26. Patra, D. K., Si, T., Mondal, S. & Mukherjee, P. Breast DCE-MRI segmentation for lesion detection by multi-level thresholding using student psychological based optimization. *Biomed. Signal Process. Control.* **69**, 102925. [https://doi.org/10.1016/j.bspc.2021.](https://doi.org/10.1016/j.bspc.2021.102925) [102925](https://doi.org/10.1016/j.bspc.2021.102925) (2021).
- 27. Wu, Q., Salganicoff, M., Krishnan, A., Fussell, D. S. & Markey, M. K. Interactive lesion segmentation on dynamic contrast enhanced breast MRI using a Markov model. in *Medical Imaging 2006: Image Processing*, vol. 6144, 1487–1494. International Society for Optics and Photonics (eds Reinhardt, J. M. & Pluim, J. P. W.) (SPIE, 2006).
- 28. Azmi, R. & Norozi, N. A new markov random feld segmentation method for breast lesion segmentation in MR images. *J. Med. Signals Sensors* **1**, 156–164 (2011).
- 29. Ertas, G., Demirgunes, D. D. & Erogul, O. Conventional and multi-state cellular neural networks in segmenting breast region from MR images: Performance comparison. *2012 Int. Symp. on Innov. Intell. Syst. Appl.* 1–5 (2012).
- 30. Wajid, S. K., Hussain, A. & Huang, K. Tree-dimensional local energy-based shape histogram (3D-LESH): A novel feature extraction technique. *Expert. Syst. Appl.* **112**, 388–400 (2018).
- 31. Rasti, R., Teshnehlab, M. & Phung, S. L. Breast cancer diagnosis in DCE-MRI using mixture ensemble of convolutional neural networks. *Pattern Recognit.* **72**, 381–390 (2017).
- 32. Xu, X. *et al.* Breast region segmentation being convolutional neural network in dynamic contrast enhanced MRI. in *2018 40th Annu. Int. Conf. IEEE Eng. Medicine Biol. Soc. (EMBC)* 750–753 (2018).
- 33. Zhang, Y., Pan, C., Chen, X. & Wang, F. Abnormal breast identifcation by nine-layer convolutional neural network with parametric rectifed linear unit and rank-based stochastic pooling. *J. Comput. Sci.* **27**, 57–68 (2018).
- 34. Piantadosi, G., Sansone, M. & Sansone, C. Breast segmentation in MRI via U-net deep convolutional neural networks. in *2018 24th Int. Conf. on Pattern Recognit. (ICPR)* 3917–3922 (2018).
- 35. Zhang, J., Saha, A., Zhu, Z. & Mazurowski, M. Hierarchical convolutional neural networks for segmentation of breast tumors in MRI with application to radiogenomics. *IEEE Trans. Med. Imaging* **38**, 435–447 (2019).
- 36. Benjelloun, M., Adoui, M., Larhmam, M. A. & Mahmoudi, S. Automated breast tumor segmentation in DCE-MRI using deep learning. in *2018 4th Int. Conf. on Cloud Comput. Technol. Appl. (Cloudtech)* 1–6 (2018).
- 37. Zheng, J. *et al.* Deep learning assisted efcient adaboost algorithm for breast cancer detection and early diagnosis. *IEEE Access* **8**, 96946–96954 (2020).
- 38. Negi, A., Raj, A., Nersisson, R., Zhuang, Z. & Murugappan, M. RDA-UNET-WGAN: An accurate breast ultrasound lesion segmentation using wasserstein generative adversarial networks. *Arab. J. Sci. Eng.* **45**, 1–12 (2020).
- 39. Qiao, M. *et al.* Tree-dimensional breast tumor segmentation on DCE-MRI with a multilabel attention-guided joint-phase-learning network. *Comput. Med. Imaging Graph.* **90**, 101909 (2021).
- 40. Zhang, Y. *et al.* Automatic detection and segmentation of breast cancer on MRI using mask R-CNN trained on non-fat-sat images and tested on fat-sat images. *Acad. Radiol*. **29**, S135–S144 (2020).
- 41. Liu, M. Z. *et al.* Weakly supervised deep learning approach to breast MRI assessment. *Acad. Radiol.* **29**, S166–S172 (2021).
- 42. Huo, L. *et al.* Segmentation of whole breast and fbroglandular tissue using nnU-Net in dynamic contrast enhanced MR images. *Magn. Resonance Imaging*. **82**, 31–41 (2021).
- 43. Devarakonda Venkata, M. & Lingamgunta, S. A convolution neural network based MRI breast mass diagnosis using zernike moments. *Mater. Today Proc.*<https://doi.org/10.1016/j.matpr.2021.06.133> *(2021).*
- 44. Jaglan, P., Dass, R. & Duhan, M. An automatic and efficient technique for tumor location identification and classification through breast MR images. *Expert. Syst. Appl.* **185**, 115580.<https://doi.org/10.1016/j.eswa.2021.115580> (2021).
- 45. Kim, C.-M., Park, R. C. & Hong, E. J. Breast mass classifcation using eLFA algorithm based on CRNN deep learning model. *IEEE Access* **8**, 197312–197323. <https://doi.org/10.1109/ACCESS.2020.3034914>(2020).
- 46. Lv, T., Pan, X. & Li, L. DCE-MRI based breast intratumor heterogeneity analysis via dual attention deep clustering network and its application in molecular typing. in *2020 IEEE International Conference on Bioinformatics and Biomedicine (BIBM)*, 681–686. <https://doi.org/10.1109/BIBM49941.2020.9313272> (2020).
- 47. Soleimani, H. & Michailovich, O. V. Automatic breast tissue segmentation in MRI scans. in *2020 IEEE International Conference on Systems, Man, and Cybernetics (SMC)*, 1572–1577.<https://doi.org/10.1109/SMC42975.2020.9282847>(2020).
- 48. Lingle, W. *et al.* Radiology data from the cancer genome atlas breast invasive carcinoma collection [tcga-brca] (2007).
- 49. Clark, K. *et al. Te cancer imaging archive (TCIA): Maintaining and operating a public information repository*. (2013).
- 50. ME, G. M. & Subashini, M. M. Medical imaging with intelligent systems: A review. in (ed Sangaiah, A. K.) *Deep Learning and Parallel Computing Environment for Bioengineering Systems*, chap. 4, 53–73.<https://doi.org/10.1016/B978-0-12-816718-2.00011-7> (Academic Press, 2019).
- 51. Diakogiannis, F. I., Waldner, F., Caccetta, P. & Wu, C. Resunet-a: a deep learning framework for semantic segmentation of remotely sensed data (2020). [arXiv:1904.00592](http://arxiv.org/abs/1904.00592).
- 52. Zeng, Y., Chen, X., Zhang, Y., Bai, L. & Han, J. Dense-U-Net: densely connected convolutional network for semantic segmentation with a small number of samples. in (eds Li, C., Yu, H., Pan, Z. & Pu, Y.) *Tenth International Conference on Graphics and Image Processing (ICGIP 2018)*, vol. 11069, 665 – 670. <https://doi.org/10.1117/12.2524406>. International Society for Optics and Photonics (SPIE, 2019).
- 53. Jin, Q. *et al.* Dunet: A deformable network for retinal vessel segmentation. *Knowledge-Based Syst.* **178**, 149–162. [https://doi.org/](https://doi.org/10.1016/j.knosys.2019.04.025) [10.1016/j.knosys.2019.04.025](https://doi.org/10.1016/j.knosys.2019.04.025) (2019).
- 54. Oktay, O. *et al.* Attention u-net: Learning where to look for the pancreas (2018). [arXiv:1804.03999.](http://arxiv.org/abs/1804.03999)
- 55. Zhou, Z., Siddiquee, M. M. R., Tajbakhsh, N. & Liang, J. Unet++: A nested u-net architecture for medical image segmentation (2018). [arXiv:1807.10165.](http://arxiv.org/abs/1807.10165)
- 56. Ibtehaz, N. & Rahman, M. S. Multiresunet: Rethinking the u-net architecture for multimodal biomedical image segmentation. *Neural Netw.* **121**, 74–87.<https://doi.org/10.1016/j.neunet.2019.08.025> (2020).
- 57. Ni, Z.-L. *et al.* Raunet: Residual attention u-net for semantic segmentation of cataract surgical instruments (2019). [arXiv:1909.](http://arxiv.org/abs/1909.10360) [10360.](http://arxiv.org/abs/1909.10360)
- 58. Li, H., Li, A. & Wang, M. A novel end-to-end brain tumor segmentation method using improved fully convolutional networks. *Comput. Biol. Med.*<https://doi.org/10.1016/j.compbiomed.2019.03.014> *(2019).*
- 59. Schönfeld, E., Schiele, B. & Khoreva, A. A u-net based discriminator for generative adversarial networks (2021). [arXiv:2002.12655](http://arxiv.org/abs/2002.12655).
- 60. Milletari, F., Navab, N. & Ahmadi, S.-A. V-net: Fully convolutional neural networks for volumetric medical image segmentation. <https://doi.org/10.48550/ARXIV.1606.04797> (2016).
- 61. Xia, X. & Kulis, B. W-net: A deep model for fully unsupervised image segmentation.<https://doi.org/10.48550/ARXIV.1711.08506> (2017).
- 62. Google Colaboratory.<http://colab.google>.
- 63. Tarwat, A. Classifcation assessment methods. *Appl. Comput. Informatics* **17**, 168–192.<https://doi.org/10.1016/j.aci.2018.08.003> (2018).
- 64. Bergstra, J. & Bengio, Y. Random search for hyper-parameter optimization. *J. Mach. Learn. Res.* **13**, 281–305 (2012).
- 65. IBM SPSS Statistics 23. [http://www.ibm.com/products/spss-statistics.](http://www.ibm.com/products/spss-statistics)
- 66. Mishra, P., Biancolillo, A., Roger, J. M., Marini, F. & Rutledge, D. N. New data preprocessing trends based on ensemble of multiple preprocessing techniques. *TrAC Trends Anal. Chem.* **132**, 116045. <https://doi.org/10.1016/j.trac.2020.116045>(2020).
- 67. Ameri Sianaki, O., Sherkat Masoum, M. & Potdar, V. A decision support algorithm for assessing the engagement of a demand response program in the industrial sector of the smart grid. *Comput. & Ind. Eng.* **115**, 123–137. [https://doi.org/10.1016/j.cie.2017.](https://doi.org/10.1016/j.cie.2017.10.016) [10.016](https://doi.org/10.1016/j.cie.2017.10.016) (2018).
- 68. Si, T., Miranda, P., Galdino, J. & Nascimento, A. Grammar-based automatic programming for medical data classifcation: An experimental study. *Artif. Intell. Rev.*<https://doi.org/10.1007/s10462-020-09949-9> *(2021).*
- 69. Bagchi, J. & Si, T. Artifcial neural network training using marine predators algorithm for medical data classifcation. in (eds Tiwari, R., Mishra, A., Yadav, N. & Pavone, M.) *Proceedings of International Conference on Computational Intelligence*, Algorithms for Intelligent Systems (Springer, 2022).
- 70. Siddique, N., Paheding, S., Elkin, C. P. & Devabhaktuni, V. U-net and its variants for medical image segmentation: A review of theory and applications. *IEEE Access* **9**, 82031–82057. <https://doi.org/10.1109/ACCESS.2021.3086020>(2021).
- 71. Kaveh, M. & Mesgari, M. Application of meta-heuristic algorithms for training neural networks and deep learning architectures: A comprehensive review. *Neural Process. Lett.* **55**, 4519–4622 (2023).

Acknowledgements

Z.Z. was partially supported by the Cancer Prevention and Research Institute of Texas (CPRIT RP170668 and RP180734) (to Z.Z.). Publication costs were funded by Z.Z.'s Professorship Fund. The funder had no role in the study design, data collection and analysis, decision to publish, or preparation of the manuscript.

Author contributions

S.S., D.D. and T.S. performed the experiments. K.B., S.M., and U.M. has done the validation and assessment of the models and Z.Z. supervised the work and also helped in manuscript writing.

Competing interests

The authors declare no competing interests.

Additional information

Correspondence and requests for materials should be addressed to Z.Z.

Reprints and permissions information is available at [www.nature.com/reprints.](www.nature.com/reprints)

Publisher's note Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Open Access Tis article is licensed under a Creative Commons Attribution 4.0 International $\overline{ }$ $\overline{ }$ \odot License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. Te images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit<http://creativecommons.org/licenses/by/4.0/>.

 $© The Author(s) 2023$