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| 2 | SAFNet: a deep spatial attention network with classifier fusion for breast |
| 3 | cancer detection |
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| 12 | Abstract: Breast cancer is a top dangerous killer for women. An accurate early diagnosis of breast cancer is the |
| 13 | primary step for treatment. A novel breast cancer detection model called SAFNet is proposed based on ultrasound |
| 14 | images and deep learning. We employ a pre-trained ResNet-18 embedded with the spatial attention mechanism as the |
| 15 | backbone model. Three randomized network models are trained for prediction in the SAFNet, which are fused by |
| 16 | majority voting to produce more accurate results. A public ultrasound image dataset is utilized to evaluate the |
| 17 | generalization ability of our SAFNet using 5-fold cross-validation. The simulation experiments reveal that the SAFNet |
| 18 | can produce higher classification results compared with four existing breast cancer classification methods. Therefore, |
| 19 | our SAFNet is an accurate tool to detect breast cancer that can be applied in clinical diagnosis. |
| 20 | 11 8 |
| 21 | Keywords: breast cancer; ultrasound image; ResNet; randomized neural network; randomized vector functional-link; |
| 22 | computer-aided diagnosis |
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| 24 | 1. Introduction |
| 25 | |
| 26 | Various factors can cause breast cancer, including radiation, family history, etc. There are millions of women |
| 27 | diagnosed with breast cancer annually, and approximately fifty percent of them would eventually lose their lives |
| 28 | because of late diagnosis [1]. Therefore, accurate detection of breast cancer at its initial stage plays a crucial part in |
| 29 | treating this deadly disease. Ultrasound imaging is one of the most prevailing imaging modalities in clinical diagnosis |
| 30 | that can generate clear images of the organs inside bodies [2]. Hence, ultrasound images (USIs) can be used for breast |
| 31 | cancer diagnosis. |
| 32 | Nevertheless, manual interpretation of the rich information in USIs suffers from low reproducibility and low |
| 33 | efficiency, so developing automatic USI analysis systems is necessary, which can make accurate predictions based on |
| 34 | the USIs. Due to the unprecedented development of deep learning and computer vision, computer-aided diagnosis |
| 35 | (CAD) has gained significant improvements, such as classification and segmentation [3-6]. Researchers and |
| 36 | practitioners have proposed a bunch of breast cancer detection models in the past decade. |

37 Rouhi, Jafari, Kasaei and Keshavarzian (2015) [7] proposed a segmentation and recognition system for breast 38 cancer detection based on mammograms. A cellular neural network was trained to segment the mammograms. Then, 39 they extracted a group of handcrafted image features and trained several classical artificial neural networks for classification. Amrane, Oukid, Gagaoua and Ensari (2018) [1] used the k-nearest neighbors (k-NN) algorithm and 40 41 naïve Bayesian classifier (NBC) to classify breast cancer from healthy samples using clinical data. They discovered 42 that the k-NN performed better than the NBC. Gao, Wu, Li, Zheng, Ruan, Shang and Patel (2018) [8] presented a 43 CNN-based model for breast cancer classification. They designed a shallow CNN model to learn latent representations 44 from the medical images. Aslan, Celik, Sabanci and Durdu (2018) [9] employed four different classification models 45 to diagnose breast cancer with blood statistics, including k-NN, extreme learning machine (ELM), artificial neural 46 network (ANN), and support vector machine (SVM). They also utilized an optimization method to find the optimal 47 hyper-parameters in the four classification models. Dai, Chen, Zhu and Zhang (2018) [10] trained a random forest to diagnose breast cancer in clinical data. The Wisconsin Diagnostic Breast Cancer dataset was utilized in the 48 49 experiments to evaluate the classification ability of their method. Ghasemzadeh, Sarbazi Azad and Esmaeili (2018) 50 [11] put forward a breast cancer classification approach using mammograms. A Gabor wavelet transform was 51 employed to generate the image representations from the mammograms, and several machine learning (ML) 52 algorithms were trained for classification, including SVM, ANN, decision tree (DT), etc. A public mammogram 53 dataset (DDSM) was employed in their evaluation experiments. Gupta and Gupta (2018) [12] also trained a group of 54 heterogeneous ML classifiers to detect breast cancer using the Wisconsin Diagnostic Breast Cancer dataset. Through 55 their experiments, they found that the multi-layer perceptron was better than other classification models. Heidari, 56 Khuzani, Hollingsworth, Danala, Mirniaharikandehei, Qiu, Liu and Zheng (2018) [13] presented a locally preserving 57 projection to reduce the dimension of image feature vectors from the mammograms, which were extracted based on 58 the bilateral asymmetry of the breasts. Two classical machine learning algorithms were selected as the classifiers, 59 including SVM and k-NN. The leave-one-out validation method was employed to obtain the testing results of their model. Hussain, Aziz, Saeed, Rathore and Rafique (2018) [14] proposed to calculate a group of different features 60 61 using the mammograms based on the texture analysis, Fourier transform, scale-invariant feature transform (SIFT), etc. 62 Then, combined features were used to train an SVM and a DT using 10-fold cross-validation. Wang, Li, Wang, Jiang, 63 Yao, Zhang and Xin (2019) [15] put forward a mass detection algorithm based on the deep CNN and ELM. They 64 extracted the deep features and fused them with classical handcrafted features to form the feature vectors. Finally, an ELM was trained to distinguish the benign and malignant samples. Islam, Haque, Iqbal, Hasan, Hasan and Kabir (2020) 65 66 [16] conducted a comparison of the classification results of the ANN and SVM on the Wisconsin Breast Cancer dataset 67 for breast cancer detection. They found that the ANN outperformed the SVM marginally. Lahoura, Singh, Aggarwal, Sharma, Mohammed, Damasevicius, Kadry and Cengiz (2021) [17] provided a breast cancer classification system 68 69 based on cloud computing. A gain ratio feature selection algorithm was used to eliminate the excessive features, and 70 an ELM served as the classifier for remote breast cancer detection. The Wisconsin Diagnostic Breast Cancer was employed in their experiments. Zuluaga-Gomez, Al Masry, Benaggoune, Meraghni and Zerhouni (2021) [18] 71

72 presented a breast cancer diagnosis framework using ResNet and thermal images. The trained model can achieve good

73 classification performance in their experiments. Rehman, Zhuang, Muhamed Ali, Ibrahim and Li (2019) [19] 74 attempted to find out the correlation between microRNAs and breast cancer. They proposed to utilize a bunch of feature selection algorithms to obtain the refined microRNA set which were the most related to breast cancer. A random 75 forest and an SVM were trained for breast cancer classification with the refined microRNAs as the inputs. Singh and 76 77 Singh (2020) [20] presented a systematic review on the breast cancer diagnosis using thermography. Stark, Hart, 78 Nartowt and Deng (2019) [21] used personal healthcare data to predict breast cancer risk. The input features included 79 age, ethnicity, etc. For classification, multiple machine learning models were trained, including ANN, SVM, NBC, etc. However, their accuracies were relatively low. Tapak, Shirmohammadi-Khorram, Amini, Alafchi, Hamidi and 80 81 Poorolajal (2019) [22] implemented the survival prediction of breast cancer using clinical data and classical machine 82 learning classifiers, including SVM, ANN, random forest, etc. however, the sensitivity of their best model was only 83 36%. Zheng, Lin, Gao, Wang, He and Fan (2020) [23] developed a breast tumor classification model using a 84 convolutional neural network (CNN) and Adaboost algorithms. They gathered images of different modalities, 85 including mammograms, USIs, and magnetic resonance images (MRIs). Their model can accurately classify the samples as normal, benign, and malignant after training. Khuriwal and Mishra (2018) [24] firstly exploited watershed 86 87 segmentation to remove the background in the mammograms. Then, they extracted 12 texture and statistical features 88 from the segmented images, including entropy, energy, mean value, standard deviation, etc. Finally, an ANN model was trained to classify breast cancer from normal controls. Kadam, Jadhav and Vijayakumar (2019) [25] employed a 89 90 sparse autoencoder to identify malignant breast cancer from benign breast cancer. Mercan, Mehta, Bartlett, Shapiro, 91 Weaver and Elmore (2019) [26] proposed a breast cancer classification framework based on biopsy images. They 92 generated small patches from the images and trained a CNN model to generate patch-level tissue labels. Then, the 93 image-level tissue labels can be obtained based on the fusion of patch-level tissue labels. Finally, an SVM was trained 94 to identify the breast cancer subtypes. Turkki, Byckhov, Lundin, Isola, Nordling, Kovanen, Verrill, von Smitten, 95 Joensuu, Lundin and Linder (2019) [27] employed a CNN to generate features from the tissue microarray images. 96 Then, the principal component analysis (PCA) and Fisher vector were used for feature reduction. An SVM was trained 97 to predict the input as high breast cancer risk and low breast cancer risk. Zeebaree, Haron, Abdulazeez and Zebari 98 (2019) [28] put forward a segmentation method using USI. They separated the USIs into small patches and extracted 99 several features from the patches to train a back propagation neural network (BPNN), which was used to classify the 100 patches as region of interest (ROI) and non-ROI. Finally, a region growing algorithm was used to refine the segmented 101 results. Sharma and Mehra (2020) [29] developed a new method to recognize breast cancer in histopathological images. 102 They employed two different approaches to extract image features. One is the classical handcrafted features, including 103 Hu moment, texture analysis. The other is the features based on deep CNN models, including VGG-16, VGG-19, and 104 ResNet-50. In the classification stage, the SVM, linear discriminant analysis (LDA), random forest were trained to 105 distinguish benign cancer from malignant ones. Zuluaga-Gomez, Al Masry, Benaggoune, Meraghni and Zerhouni (2020) [30] trained a deep CNN model to detect breast cancer in thermal images. In experiments, they tested a bunch 106 107 of different backbone models. The best accuracy of their model was 92%. Mahmood, Arsalan, Owais, Lee and Park 108 (2020) [31] tried to implement breast cancer detection by mitotic cell counting. A faster region CNN (FR-CNN) was

developed to detect mitotic cells in the images based on ResNet-50 and DenseNet-201. The FR-CNN achieved an F1 score of 85.8% on a public dataset.

The above analysis shows that CAD systems for breast cancer detection using either clinical data or medical images are becoming more and more effective. However, we believe that the performance of automatic breast cancer detection classification can be further improved because most current methods are developed based on either deep CNN models or classical machine learning models with handcrafted features. This paper presented a novel and simple breast cancer detection model called spatial attention fusion network (SAFNet) based on USIs. The main contributions are three-fold:

- A spatial attention deep CNN model was designed based on the spatial attention module and the ResNet-18
 backbone, pre-trained on the ImageNet dataset, and fine-tuned with the USIs for image feature extraction.
- Three randomized neural networks (RNNs) were employed as the classifiers to prevent the overfitting
 problem because they are all shallow networks with simple structures and are easy and fast to train.
- 3) We proposed a late fusion mechanism to fuse the output labels using majority voting to stabilize the
 classification results of our breast cancer classification system.

The rest of this paper is arranged as follows. The introduction of the dataset for evaluation experiments is given in Section 2. The detailed presentation of the proposed SAFNet is demonstrated in Section 3. Section 4 presents the experiment results, and the discussion is demonstrated in Section 5. Finally, the conclusion of this study is provided in Section 6.

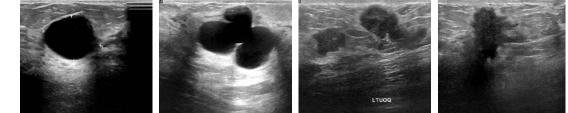
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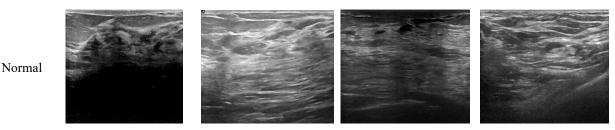
128 2. Materials

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We utilized a public USI dataset to evaluate the generalization ability of our SAFNet [32]. The USI dataset can be downloaded from Kaggle (https://www.kaggle.com/datasets/aryashah2k/breast-ultrasound-images-dataset). We finally obtained 437 benign USIs, 210 malignant USIs, and 166 normal USIs, which were approximately in size of 500×500 pixels. The ages of the women in the USIs ranged from 25 to 75. We focused on breast cancer detection in this study. We regarded both the benign and malignant samples as breast cancer USIs. Some samples in the USI dataset are shown in Figure 1.







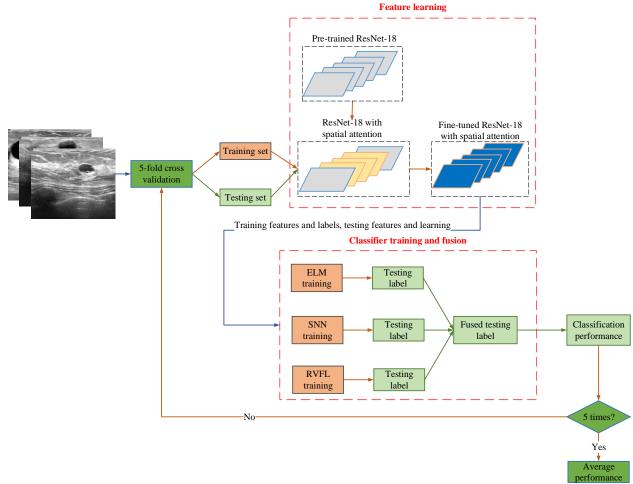
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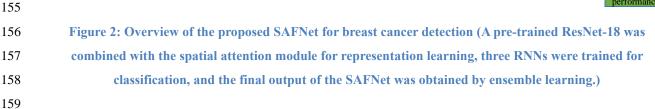
- 137 Figure 1: Samples in the USI dataset (The four USIs in the first row are breast cancer, and the four USIs in
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140 **3.** Methodology

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142 We proposed a spatial attention fusion network (SAFNet) to diagnose breast cancer in USIs. The SAFNet was 143 designed using both deep learning and classical machine learning. The overview of the SAFNet is presented in Figure 144 2. Firstly, the backbone model for the SAFNet was a pre-trained ResNet-18, which was modified with a spatial 145 attention module and fine-tuned on the USI dataset. Afterward, the fine-tuned backbone generated image 146 representations using the training and testing USIs. Afterward, three RNNs were trained using the image representations and the labels, including ELM, random vector functional-link net (RVFL), and Schmidt neural network 147 148 (SNN). Finally, the prediction labels of the SAFNet were calculated by the majority voting-based fusion of the 149 predictions from the three RNNs. The spatial attention module can improve the image representation learning ability 150 of the pre-trained ResNet-18 backbone. The RNNs were simple three-layered networks, which can effectively avoid 151 the overfitting problem. The late fusion of the RNNs was designed to handle the negative effects of randomly 152 initialized parameters in the RNNs and improve the classification performance of the SAFNet. 5-fold cross-validation 153 was employed in our evaluation experiments.





160 **3.1.** Feature learning based on spatial attention network

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During the recent ten years, deep CNN architectures can be the first choice for computer vision tasks. Among the various CNN models, ResNet can be one of the most significant inventions designed with residual connections to make it easy to approximate identity mappings [33]. The residual connection can directly link two layers that are not adjacent, which skips the hidden layers between the two layers, as is shown in Figure 3. Given X as the activation maps of the previous layer, the training target was G(X). with the residual connection, the target now becomes

167 $\mathbf{F}(\mathbf{X}) := \mathbf{G}(\mathbf{X}) - \mathbf{X}$ (1)

168 In this way, the original learning target can be expressed as

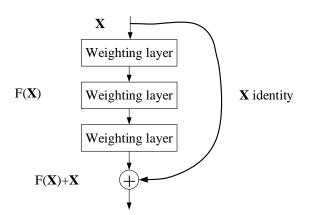
$$\mathbf{G}(\mathbf{X}) = \mathbf{F}(\mathbf{X}) + \mathbf{X} \tag{2}$$

170 The function F(X) is the residual between the training target and identity mapping. Instead of learning the target

171 mapping directly, the model was trained to approximate the residual function so that the identity mappings could be

trained more effectively.

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175 Figure 3: An example of a residual connection (A shortcut connection is added to skip the middle layers.)

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177 The attention modules in the recent networks were inspired by human vision, which often focuses on specific regions of images. Researchers discovered that the attention mechanism could be embedded in current CNN models 178 179 to boost their classification ability [34], and the implementation of attention modules can be plug-and-play [35]. Therefore, we proposed to embed the spatial attention module with the pre-trained ResNet-18 for feature learning 180 181 from the breast USIs. A detailed presentation of the spatial attention module with ResNet-18 is illustrated in Figure 4. 182 The spatial attention module was inserted between the rectified linear unit (ReLU) layer and the multiplication layer. 183 In the spatial attention module, an average pooling layer and a max pooling layer were employed to generate two sets 184 of feature maps. Then, the two groups of feature mappings were concatenated and passed to a convolution layer to 185 fuse them together. Finally, a ReLU served as the activation function. The feature mappings of the spatial attention 186 module were multiplied with the activation matrix of the top ReLU layer. We also modified the densely connected layers before the final layer of the ResNet-18 according to our breast USI dataset. The 'Fully connected layer 256' 187 188 contained 256 nodes, which served as the feature layer in this study. As the breast cancer diagnosis is a binary 189 classification problem, the dimension of the output layer was set as 2. 190 The pre-trained ResNet-18 with spatial attention mechanism was fine-tuned on the USI dataset for only four total 191 epochs before the feature extraction from the 'Fully connected layer 256'.

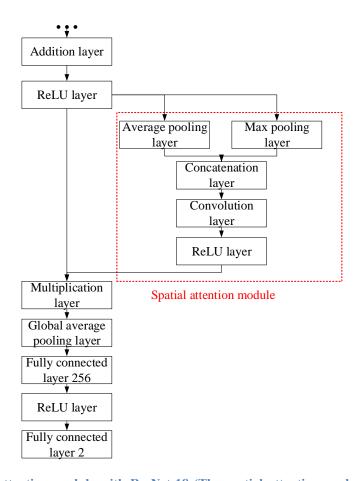


Figure 4: The spatial attention module with ResNet-18 (The spatial attention module was inserted into the
ResNet-18. In The spatial attention module, the activations were processed with average pooling and max
pooling, respectively. Then, the two activation maps were concatenated and sent into a convolution layer with
ReLU activation function.)

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199 **3.2.** Training and fusion of classifiers

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As for the classifier in our SAFNet, we utilized three RNNs, including ELM [36], random vector functional-link (RVFL) [37], and Schmidt neural network (SNN) [38]. The three RNNs are all shallow networks with merely three layers, so the overfitting problem can be alleviated effectively, which often occurs when training deep CNN architectures with a small training set. Meanwhile, the training of RNNs is considerably faster compared with BPNNs benefiting from the randomized weights and biases in the hidden layer. Though RNNs can be trained fast, their classification performance is promising. Therefore, RNNs have been employed in a variety of machine learning tasks [39-41].

The architecture of an ELM is demonstrated in Figure 5. The parameters to be trained include the hidden weights w_i , hidden biases b_i , and output weights β_i . The training algorithm of an ELM consists of only three steps, as is given in Algorithm 1. Suppose we get the training representation from the fine-tuned backbone and their labels as

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$$S_{train} = [(f_1, t_1), (f_2, t_2), (f_3, t_3), \dots, (f_N, t_N)]$$
(3)

212 where the $f_j = (f_{j1}, f_{j2}, f_{j3}, \dots, f_{jn})^T$ denotes the *j*-th image feature vector, the $\mathbf{t}_j = (t_{j1}, t_{j2}, t_{j3}, \dots, t_{jm})^T$ represents

213 the label of the *j*-th USI, and *N* stands for the entire number of training USIs. Then, the activation matrix of the hidden 214 H_{act} can be calculated as

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$$\mathbf{H}_{act} = \sum_{i=1}^{\hat{N}} g(\mathbf{w}_i \mathbf{f}_j + b_i) = \begin{bmatrix} g(\mathbf{w}_1 \mathbf{f}_1 + b_1) & \cdots & g(\mathbf{w}_{\hat{N}} \mathbf{f}_1 + b_{\hat{N}}) \\ \vdots & \ddots & \vdots \\ g(\mathbf{w}_1 \mathbf{f}_N + b_1) & \cdots & g(\mathbf{w}_{\hat{N}} \mathbf{f}_N + b_{\hat{N}}) \end{bmatrix}_{N \times \hat{N}}, j = 1, \dots, N$$
(4)

in which the g(x) denotes the activation function, and \hat{N} stands for the dimension of the hidden space. Finally, the prediction of the ELM is

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 $\mathbf{O} = \mathbf{H}_{act}\boldsymbol{\beta} \tag{5}$

(7)

where $\boldsymbol{\beta} = (\boldsymbol{\beta}_1, \boldsymbol{\beta}_2, \boldsymbol{\beta}_3, ..., \boldsymbol{\beta}_{\tilde{N}})^{\mathrm{T}}$ is the output weights, and the $\mathbf{0} = (\boldsymbol{o}_1, \boldsymbol{o}_2, \boldsymbol{o}_3, ..., \boldsymbol{o}_{\mathrm{N}})^{\mathrm{T}}$ denotes the prediction matrix of the ELM. The learning purpose is to make the predictions of the ELM equal to the ground-truth labels, so we can have

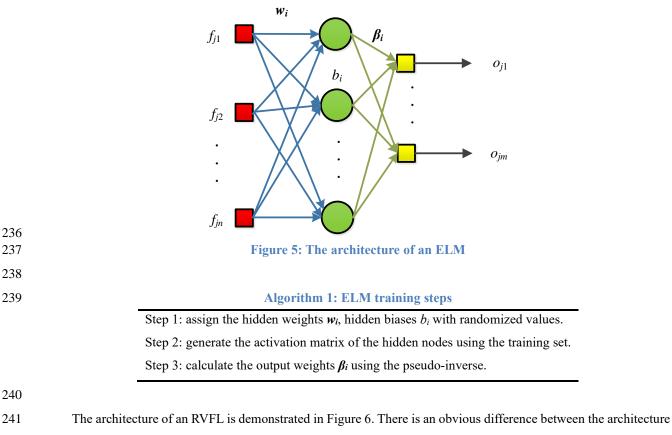
 $H_{act}\beta = T$ (6)

where $\mathbf{T} = (t_1, t_2, t_3, ..., t_N)^T$ means the ground-truth labels. In this way, the output weights $\boldsymbol{\beta}$ can be obtained based on the Moore-Penrose pseudo-inverse as

 $\beta = \mathbf{H}_{act}^{\dagger} \mathbf{T}$

where the $\mathbf{H}_{act}^{\dagger}$ denotes the Moore-Penrose pseudo-inverse of \mathbf{H}_{act} . With random initialization and pseudo-inverse, all the parameters in the ELM are decided within three steps. ELM training is much faster than vanilla BPNN, which is trained with iterations. Yet, the classification performance of the ELM is usually promising because it is likely to obtain smaller weight norms [42].

As a type of randomized neural network, ELM leverages the random projection to project the input features into the random hidden space, and only the output weights are trained. This random feature projection has also been used in deep networks in recent years [43]. ELM is a popular classification model in the last decade, which has been used in a variety of machine learning tasks, such as geography [44], big data analysis [45], clustering [46], chemistry [47], food safety [48], etc.



of RVFL and ELM. There is a shortcut connection between the input nodes and the output space [49]. In an RVFL, the input representations are randomly mapped into the hidden space (the green nodes), and the randomly mapped features are concatenated with the input feature vectors [50]. This extra shortcut connection from the input to the output layer can effectively stabilize the classification performance of the RVFL and improve the robustness of the system [51, 52].

| 248 | b_i b_i f_j f_{j1} f_{j2} f_{jn} b_i f_j | |
|-----|--|--|
| 249 | Figure 6: The structure of a RVFL | |
| 250 | | |
| 251 | The training algorithm of RVFL is summarized in Algorithm 2. The activation matrix of the hidden layer with \widehat{N} | |
| 252 | nodes can be formulated the same as equation (4). Then, the activation matrix of the hidden space is concatenated with | |
| 253 | the input features to generate the combined features F: | |
| 254 | $\mathbf{F} = concat(\mathbf{H}_{act}, \mathbf{F}_{train}) \tag{8}$ | |
| 255 | where $\mathbf{F}_{train} = (f_1, f_2, f_3, \dots, f_N)^T$ stands for the features from the feature layer in the backbone, and <i>concat</i> () | |
| 256 | is the matrix concatenation function. The training purpose is to achieve that the predictions of the RVFL are equal to | |
| 257 | the ground-truth labels: | |
| 258 | $0 = \mathbf{T} \tag{9}$ | |
| 259 | where $0 = (o_1, o_2, o_3,, o_N)^T$ is the predictions of the RVFL, and $\mathbf{T} = (t_1, t_2, t_3,, t_N)^T$ means the ground-truth | |
| 260 | labels of the training set. Therefore, we have | |
| 261 | $\mathbf{F}\boldsymbol{\beta} = \mathbf{T} \tag{10}$ | |
| 262 | Consequently, the output weights can be determined as: | |
| 263 | $\boldsymbol{\beta} = \mathbf{F}^{\dagger} \mathbf{T} \tag{11}$ | |
| 264 | where $\boldsymbol{\beta} = (\boldsymbol{\beta}_1, \boldsymbol{\beta}_2, \boldsymbol{\beta}_3, \dots, \boldsymbol{\beta}_{\widehat{N}})^{\mathrm{T}}$ is the output weights, and \mathbf{F}^{\dagger} represents the pseudo-inverse of the F. | |
| 265 | | |
| 266 | Algorithm 2: RVFL training steps | |
| _00 | Step 1: assign the hidden weights w_i , hidden biases b_i with randomized values. | |
| | Step 2: compute the activation matrix of the hidden layer using the training set. | |
| | Step 3: concatenate the input features with the activation matrix of the hidden layer. | |
| | Step 4: determine the output weights β_i using the pseudo-inverse. | |
| | | |

We also employ an SNN as the classifier in our SAFNet. SNN is a feedforward neural network proposed by Schmidt, Kraaijveld and Duin (1992) [38] thirty years ago. The structure of the SNN is almost the same as the ELM, which is shown in Figure 7. There are learnable output biases in the output layer, which is the only difference between the architecture of SNN and ELM. Therefore, the training algorithm of SNN is the same as ELM, which is illustrated in Algorithm 3. With the training features and labels, the prediction matrix of the SNN with \hat{N} nodes in the hidden layer is

 $\mathbf{0} = \left[\sum_{i=1}^{\hat{N}} g\left(\mathbf{w}_{i} \mathbf{f}_{j} + b_{i}\right)\right] \times concat(\boldsymbol{\beta}, \boldsymbol{C})$ (12)

where $C = (c_1, c_2, c_3, ..., c_m)^T$ is the output biases. The output weights and biases can be generated using close-formed solutions similar to the ELM and RVFL. In the original SNN, the standard numerical methods were employed to calculate the output weights and biases.

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| | f_{j1} f_{j2} |
|-----|--|
| 279 | f_{jn} |
| 280 | Figure 7: The structure of an SNN |
| 281 | |
| 282 | Algorithm 3: SNN training steps |
| | Step 1: assign the hidden weights w_i , hidden biases b_i with randomized values. |
| | Step 2: calculate the activation matrix of the hidden layer using the training set. |
| | Step 3: determine the output weights β_i and biases c_i using the pseudo-inverse. |
| 283 | |

w.

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Now, we have presented three classifiers for breast cancer detection. The randomly initialized parameters in the RNNs can reduce the training time of our models, but ill-conditioned input weights and biases can worsen the classification performance of our breast cancer detection model, which can cause serious results in real-world applications. Therefore, we proposed to utilize late fusion to further improve our system's generalization performance and robustness, which directly fuses the predictions of the three RNNs by majority voting. Compared with swarm optimization methods, majority voting works more efficiently. The final model with this majority voting-based fusion is termed as SAFNet. The classifiers: ELM, RVFL, and SNN are heterogeneous RNNs, so the late fusion of their
 predictions can receive diversified information so that the fused results can be better.

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293 4. Experimental results

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Our SAFNet was developed using MATLAB 2021a. The training and testing experiments are run on a laptop with CPU i7 7700HQ, 16 GB memory, and GPU GTX1060 (6GB). The trained models can be deployed in other environments with appropriate configurations.

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299 4.1. Hyper-parameter settings

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The hyper-parameters in the proposed SAFNet are illustrated in Table 1. We set the mini-batch size as only 64 according to the capability of our GPU. The learning rate was set as 1×10^{-4} , which is a conventional value. The optimizer for fine-tuning the spatial attention backbone was Adam. We fine-tuned the backbone for only four epochs in order to prevent overfitting. The dimension of the hidden space in the three RNNs was 1000, as the feature dimension was 256. The random mapping from the input space to the hidden space of high dimension is beneficial to improve the classification performance.

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Table 1: Hyper-parameters in the proposed SAFNetHyper-parameterValueMini-batch size64Learning rate 1×10^{-4} OptimizerAdamMax epochs4Number of hidden neurons in the three RNNs1000

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- 310 **4.2.** Classification performance
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312 The classification performance of the proposed SAFNet is listed in Table 2. We employed sensitivity, precision,

- 313 F1-score, and accuracy as the performance metrics in the evaluation experiments.
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Table 2: Classification results of the SAFNet (F: fold)

| | Sensitivity | Precision | F1-score | Accuracy |
|---------|-------------|-----------|----------|----------|
| F1 | 93.38% | 98.45% | 95.85% | 92.95% |
| F2 | 95.59% | 100.00% | 97.74% | 96.15% |
| F3 | 93.94% | 96.12% | 95.02% | 91.67% |
| F4 | 94.78% | 98.45% | 96.58% | 94.23% |
| F5 | 96.95% | 97.69% | 97.32% | 95.51% |
| Average | 94.93% | 98.14% | 96.50% | 94.10% |

317 **4.3.** Effects of the spatial attention mechanism

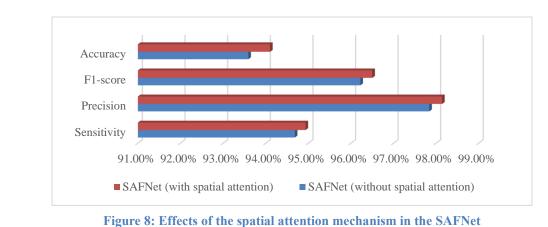
- To obtain the effects of the spatial attention mechanism, we tested the performance of the proposed SAFNet with and without the spatial attention module using 5-fold cross-validation. The backbone for the SAFNet without the spatial attention mechanism was the vanilla pre-trained ResNet-18. The results are shown in Table 3 and Figure 8.
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Table 3: Effects of the spatial attention mechanism in the SAFNet

| Model | Sensitivity | Precision | F1-score | Accuracy |
|------------------------------------|-------------|-----------|----------|----------|
| SAFNet (without spatial attention) | 94.68% | 97.84% | 96.22% | 93.59% |
| SAFNet (with spatial attention) | 94.93% | 98.14% | 96.50% | 94.10% |



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328 4.4. Effects of the late fusion

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We also tested the effects of the late fusion of classifiers using 5-fold cross-validation. The classification results of the SAFNet with the ELM classifier, RVFL classifier, SNN classifier, and the fusion of the three RNNs are given in Table 4 and Figure 9.

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Table 4: Effects of the late fusion of classifiers

| Model | Sensitivity | Precision | F1-score | Accuracy |
|-----------------------------------|-------------|-----------|----------|----------|
| SAFNet (ELM) | 94.24% | 98.30% | 96.22% | 93.59% |
| SAFNet (SNN) | 94.76% | 96.76% | 95.73% | 92.82% |
| SAFNet (RVFL) | 94.50% | 97.99% | 96.20% | 93.59% |
| SAFNet (fusion of the three RNNs) | 94.93% | 98.14% | 96.50% | 94.10% |

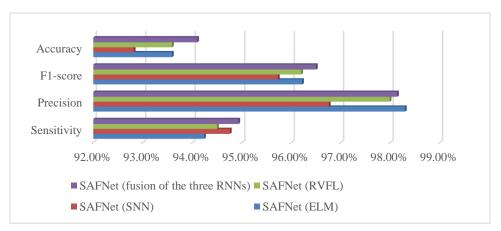


Figure 9: Effects of the late fusion of classifiers

336

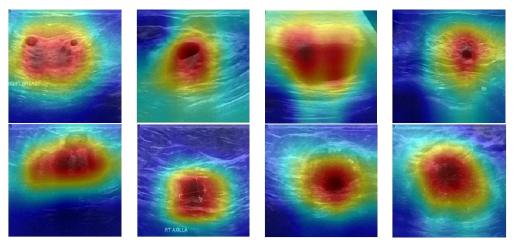
339 4.5. Visual explanation

340

We utilized the Gradient-weighted class activation mapping (Grad-CAM) [53] for a visual explanation of the proposed SAFNet. Some Grad-CAMs of the breast cancer USIs based on the SAFNet were presented in Figure 10.

343 The regions in red and orange are considered the most significant for the predictions by the SAFNet.

344



345

Figure 10: Grad-CAMs of the breast cancer USIs based on the SAFNet

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347 **4.6.** Comparison with state-of-the-art methods

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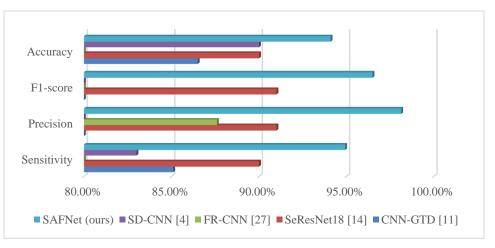
We compared our SAFNet with other existing breast cancer detection methods, and the statistics were presented in Table 5 and Figure 11. Our SAFNet produced better classification results than four existing breast cancer classification methods in terms of sensitivity, precision, F1-score, and accuracy.

353

Table 5: Comparison with other state-of-the-art breast cancer detection models

| Model | Sensitivity | Precision | F1-score | Accuracy |
|---------------------------|-------------|-----------|----------|----------|
| CNN-GTD [<u>15</u>] | 85.10% | - | - | 86.50% |
| SeResNet18 [<u>18</u>] | 90.00% | 91.00% | 91.00% | 90.00% |
| FR-CNN [<u>31</u>] | - | 87.60% | - | - |
| SD-CNN [<mark>8</mark>] | 83.00% | - | - | 90.00% |
| SAFNet (ours) | 94.93% | 98.14% | 96.50% | 94.10% |







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Figure 11: Comparison with other state-of-the-art breast cancer detection models

5. Discussion 358

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The SAFNet achieved an average sensitivity of 94.93%, an average precision of 98.14%, an average F1 score of 360 361 96.50%, and an average accuracy of 94.10% for the 5-fold cross-validation. All four metrics fluctuated around 95%, 362 which revealed the good generalization performance of the proposed SAFNet for breast cancer detection in USIs. 363 Meanwhile, the running time for the 5-fold cross-validation of the SAFNet was merely 274.83 seconds, which was 364 affordable for clinical applications.

365 We discovered that the spatial attention mechanism could boost the generalization ability of the SAFNet for 366 breast cancer detection in terms of all four metrics, although the improvement was not considerable. It can be inferred 367 from the results that the spatial attention module is an effective and simple method to boost the generalization ability 368 of the proposed model for breast cancer diagnosis.

369 The SAFNet with the late fusion of the three RNNs achieved better sensitivity, F1-score, and accuracy than the 370 SAFNets with the three individual RNNs. The precision of the SAFNet with the late fusion was 98.14%, which was 371 also close to the best precision of 98.30% in the list. Therefore, we held the view that the late fusion mechanism can 372 be beneficial for the generalization ability of the SAFNet for breast cancer detection in USIs.

373 From the Grad-CAMs, we can claim that our SAFNet can locate the potential lesion areas in the breast USIs, 374 which contributes to the promising results.

375 The potential reasons for the outstanding generalization ability of the SAFNet include that the spatial attention 376 module improves the representation learning capability of the pre-trained ResNet-18 backbone, the RNN classifiers

can avoid the overfitting problem, and the late fusion of the classifiers can eliminate the bad effects of the randomparameters in the RNNs.

379

380 6. Conclusion

381 This study put forward a novel breast cancer detection approach called SAFNet based on the ultrasound images. 382 A pre-trained ResNet-18 was embedded with the spatial attention module to serve as the backbone model in the 383 SAFNet to extract features from the breast USIs. We trained three RNNs as the classifiers and fused their predictions 384 as the final output of the SAFNet. A public breast USI dataset was utilized to evaluate the generalization ability of the 385 SAFNet based on 5-fold cross-validation. Extensive experiments were conducted, and our SAFNet outperformed four 386 existing approaches in terms of sensitivity, precision, F1 score, and accuracy. The Grad-CAMs also revealed the high 387 performance of the SAFNet to locate lesion areas. In all, our SAFNet is accurate in detecting breast cancer from USIs. For future research, we shall collect more breast USIs as the number of normal USIs in the current dataset is 388 small. In addition, we shall attempt to apply vision transformers for breast cancer detection. We will also try to harness 389 390 optimization algorithms to train the RNNs [54-57]. Moreover, image segmentation shall be studied in the future [58-391 <u>61</u>]. 392

393 Statements of ethical approval

This work does not contain any studies with human participants or animals performed by any authors.

394

Declaration of competing interests

The authors declare that they have no competing interests.

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399

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410 **References**

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412 1. Amrane, M., S. Oukid, I. Gagaoua, and T. Ensari, *Breast Cancer Classification Using Machine Learning*. 2018 Electric Electronics, Computer

413 Science, Biomedical Engineerings' Meeting (EBBT), 2018

- 414 2. Roy, S., W. Menapace, S. Oei, B. Luijten, E. Fini, C. Saltori, I. Huijben, N. Chennakeshava, F. Mento, A. Sentelli, E. Peschiera, R. Trevisan,
- 415 G. Maschietto, E. Torri, R. Inchingolo, A. Smargiassi, G. Soldati, P. Rota, A. Passerini, R.J.G. van Sloun, E. Ricci, and L. Demi, Deep Learning
- 416 for Classification and Localization of COVID-19 Markers in Point-of-Care Lung Ultrasound. IEEE Transactions on Medical Imaging, 2020.
 417 39(8): p. 2676-2687
- 418 3. Li, C., L. Hou, B.Y. Sharma, H. Li, C. Chen, Y. Li, X. Zhao, H. Huang, Z. Cai, and H. Chen, *Developing a new intelligent system for the*419 *diagnosis of tuberculous pleural effusion*. Computer methods and programs in biomedicine, 2018. 153: p. 211-225
- 4. Su, H., D. Zhao, F. Yu, A.A. Heidari, Y. Zhang, H. Chen, C. Li, J. Pan, and S. Quan, *Horizontal and vertical search artificial bee colony for image segmentation of COVID-19 X-ray images.* Computers in Biology and Medicine, 2022. 142: p. 105181
- 5. Chen, X., H. Huang, A.A. Heidari, C. Sun, Y. Lv, W. Gui, G. Liang, Z. Gu, H. Chen, C. Li, and P. Chen, *An efficient multilevel thresholding image segmentation method based on the slime mould algorithm with bee foraging mechanism: A real case with lupus nephritis images.*Computers in Biology and Medicine, 2022. 142: p. 105179
- 425 6. Zhao, X., X. Zhang, Z. Cai, X. Tian, X. Wang, Y. Huang, H. Chen, and L. Hu, *Chaos enhanced grey wolf optimization wrapped ELM for*426 *diagnosis of paraquat-poisoned patients.* Computational biology and chemistry, 2019. 78: p. 481-490
- Rouhi, R., M. Jafari, S. Kasaei, and P. Keshavarzian, *Benign and malignant breast tumors classification based on region growing and CNN segmentation*. Expert Systems with Applications, 2015. 42(3): p. 990-1002
- 429 8. Gao, F., T. Wu, J. Li, B. Zheng, L. Ruan, D. Shang, and B. Patel, *SD-CNN: A shallow-deep CNN for improved breast cancer diagnosis.*430 Computerized Medical Imaging and Graphics, 2018. **70**: p. 53-62
- 431 9. Aslan, M.F., Y. Celik, K. Sabanci, and A. Durdu, *Breast Cancer Diagnosis by Different Machine Learning Methods Using Blood Analysis Data*.
 432 International Journal of Intelligent Systems and Applications in Engineering, 2018. 6(4): p. 289–293
- 433 10. Dai, B., R.-C. Chen, S.-Z. Zhu, and W.-W. Zhang, Using Random Forest Algorithm for Breast Cancer Diagnosis, in 2018 International
 434 Symposium on Computer, Consumer and Control (IS3C). 2018. p. 449-452.
- 435 11. Ghasemzadeh, A., S. Sarbazi Azad, and E. Esmaeili, *Breast cancer detection based on Gabor-wavelet transform and machine learning methods.*436 International Journal of Machine Learning and Cybernetics, 2018. 10(7): p. 1603-1612
- 437 12. Gupta, M. and B. Gupta, *A Comparative Study of Breast Cancer Diagnosis Using Supervised Machine Learning Techniques*. Proceedings of
 438 the Second International Conference on Computing Methodologies and Communication (ICCMC 2018), 2018
- 439 13. Heidari, M., A.Z. Khuzani, A.B. Hollingsworth, G. Danala, S. Mirniaharikandehei, Y. Qiu, H. Liu, and B. Zheng, *Prediction of breast cancer*
- risk using a machine learning approach embedded with a locality preserving projection algorithm. Physics in Medicine & Biology, 2018. 63(3):
 p. 035020
- Hussain, L., W. Aziz, S. Saeed, S. Rathore, and M. Rafique, Automated Breast Cancer Detection Using Machine Learning Techniques by
 Extracting Different Feature Extracting Strategies, in 2018 17th IEEE International Conference On Trust, Security And Privacy In Computing
- 444 And Communications/ 12th IEEE International Conference On Big Data Science And Engineering (TrustCom/BigDataSE). 2018. p. 327-331.
- Wang, Z., M. Li, H. Wang, H. Jiang, Y. Yao, H. Zhang, and J. Xin, *Breast Cancer Detection Using Extreme Learning Machine Based on Feature Fusion With CNN Deep Features*. IEEE Access, 2019. 7: p. 105146-105158
- 447 16. Islam, M.M., M.R. Haque, H. Iqbal, M.M. Hasan, M. Hasan, and M.N. Kabir, *Breast Cancer Prediction: A Comparative Study Using Machine*448 *Learning Techniques.* SN Computer Science, 2020. 1(5)

- Lahoura, V., H. Singh, A. Aggarwal, B. Sharma, M.A. Mohammed, R. Damasevicius, S. Kadry, and K. Cengiz, *Cloud Computing-Based Framework for Breast Cancer Diagnosis Using Extreme Learning Machine.* Diagnostics (Basel), 2021. 11(2)
- 451 18. Zuluaga-Gomez, J., Z. Al Masry, K. Benaggoune, S. Meraghni, and N. Zerhouni, *A CNN-based methodology for breast cancer diagnosis using* 452 *thermal images.* Computer Methods in Biomechanics and Biomedical Engineering: Imaging & Visualization, 2021. 9(2): p. 131-145
- 453 19. Rehman, O., H. Zhuang, A. Muhamed Ali, A. Ibrahim, and Z. Li, *Validation of miRNAs as Breast Cancer Biomarkers with a Machine Learning* 454 *Approach.* Cancers (Basel), 2019. 11(3)
- 20. Singh, D. and A.K. Singh, *Role of image thermography in early breast cancer detection- Past, present and future.* Comput Methods Programs
 Biomed, 2020. 183: p. 105074
- 457 21. Stark, G.F., G.R. Hart, B.J. Nartowt, and J. Deng, *Predicting breast cancer risk using personal health data and machine learning models*. PLoS
 458 One, 2019. 14(12): p. e0226765
- 459 22. Tapak, L., N. Shirmohammadi-Khorram, P. Amini, B. Alafchi, O. Hamidi, and J. Poorolajal, *Prediction of survival and metastasis in breast*
- 460 *cancer patients using machine learning classifiers.* Clinical Epidemiology and Global Health, 2019. 7(3): p. 293-299
- 23. Zheng, J., D. Lin, Z. Gao, S. Wang, M. He, and J. Fan, *Deep Learning Assisted Efficient AdaBoost Algorithm for Breast Cancer Detection and Early Diagnosis.* IEEE Access, 2020. 8: p. 96946-96954
- 463 24. Khuriwal, N. and N. Mishra, *Breast Cancer Detection From Histopathological Images Using Deep Learning*. 3rd International Conference
 464 and Workshops on Recent Advances and Innovations in Engineering, 2018: p. 22-25
- 465 25. Kadam, V.J., S.M. Jadhav, and K. Vijayakumar, Breast Cancer Diagnosis Using Feature Ensemble Learning Based on Stacked Sparse
 466 Autoencoders and Softmax Regression. Journal of Medical Systems, 2019. 43(8): p. 263
- 467 26. Mercan, E., S. Mehta, J. Bartlett, L.G. Shapiro, D.L. Weaver, and J.G. Elmore, *Assessment of Machine Learning of Breast Pathology Structures* 468 *for Automated Differentiation of Breast Cancer and High-Risk Proliferative Lesions.* JAMA Network Open, 2019. 2(8): p. e198777
- 27. Turkki, R., D. Byckhov, M. Lundin, J. Isola, S. Nordling, P.E. Kovanen, C. Verrill, K. von Smitten, H. Joensuu, J. Lundin, and N. Linder, Breast
- 470 *cancer outcome prediction with tumour tissue images and machine learning.* Breast Cancer Research and Treatment, 2019. **177**(1): p. 41-52
- 28. Zeebaree, D.Q., H. Haron, A.M. Abdulazeez, and D.A. Zebari, *Machine learning and Region Growing for Breast Cancer Segmentation*. 2019
 International Conference on Advanced Science and Engineering (ICOASE), 2019: p. 88-93
- 473 29. Sharma, S. and R. Mehra, Conventional Machine Learning and Deep Learning Approach for Multi-Classification of Breast Cancer
 474 Histopathology Images-a Comparative Insight. Journal of Digital Imaging, 2020. 33(3): p. 632-654
- 30. Zuluaga-Gomez, J., Z. Al Masry, K. Benaggoune, S. Meraghni, and N. Zerhouni, *A CNN-based methodology for breast cancer diagnosis using thermal images.* Computer Methods in Biomechanics and Biomedical Engineering: Imaging & Visualization, 2020. 9(2): p. 131-145
- 477 31. Mahmood, T., M. Arsalan, M. Owais, M.B. Lee, and K.R. Park, *Artificial Intelligence-Based Mitosis Detection in Breast Cancer* 478 *Histopathology Images Using Faster R-CNN and Deep CNNs.* Journal of Clinical Medicine, 2020. 9(3)
- 479 32. Al-Dhabyani, W., M. Gomaa, H. Khaled, and A. Fahmy, Dataset of breast ultrasound images. Data Brief, 2020. 28: p. 104863
- 480 33. He, K., X. Zhang, S. Ren, and J. Sun, Deep Residual Learning for Image Recognition, in The IEEE Conference on Computer Vision and Pattern
 481 Recognition (CVPR). 2016, IEEE: Las Vegas, NV, USA. p. 770-778.
- 482 34. Woo, S., J. Park, J.-Y. Lee, and I.S. Kweon, CBAM: Convolutional Block Attention Module. ECCV, 2018
- 35. Xiao, Y., H. Yin, S.H. Wang, and Y.D. Zhang, *TReC: Transferred ResNet and CBAM for Detecting Brain Diseases*. Frontiers in
 Neuroinformatics, 2021. 15: p. 781551

- 485 36. Guang-Bin, H., Z. Qin-Yu, and S. Chee-Kheong, *Extreme learning machine: Theory and applications*. Neurocomputing, 2006. 70(1-3): p. 489501
- 487 37. Pao, Y.H., G.H. Park, and D.J. Sobajic, *Learning and generalization characteristics of random vector functional-link net*. Neurocomputing,
 488 1994. 6: p. 163-180
- 38. Schmidt, W.F., M.A. Kraaijveld, and R.P.W. Duin, Feedforward neural networks with random weights, in Proceedings., 11th IAPR
 International Conference on Pattern Recognition. Vol.II. Conference B: Pattern Recognition Methodology and Systems. 1992. p. 1-4.
- 491 39. Vuković, N., M. Petrović, and Z. Miljković, A comprehensive experimental evaluation of orthogonal polynomial expanded random vector
 492 *functional link neural networks for regression*. Applied Soft Computing, 2018. **70**: p. 1083-1096
- 493 40. Zhang, Y., J. Wu, Z. Cai, B. Du, and P.S. Yu, *An unsupervised parameter learning model for RVFL neural network*. Neural Networks, 2019.
 494 112: p. 85-97
- 495 41. Wong, P.K., W. Huang, C.M. Vong, and Z. Yang, *Adaptive neural tracking control for automotive engine idle speed regulation using extreme*496 *learning machine.* Neural Computing and Applications, 2019
- 497 42. Bartlett, P.L., The Sample Complexity of Pattern Classification with Neural Networks: The Size of the Weights is More Important than the Size

498 of the Network. IEEE TRANSACTIONS ON INFORMATION THEORY, 1998. 44(2): p. 525-536

- 43. Baek, S., M. Song, J. Jang, G. Kim, and S.B. Paik, *Face detection in untrained deep neural networks*. Nature Communications, 2021. 12(1):
 p. 7328
- 44. Huang, F., K. Yin, J. Huang, L. Gui, and P. Wang, *Landslide susceptibility mapping based on self-organizing-map network and extreme learning machine.* Engineering Geology, 2017. 223: p. 11-22
- 503 45. Deng, W.Y., Z. Bai, G.B. Huang, and Q.H. Zheng, A Fast SVD-Hidden-nodes based Extreme Learning Machine for Large-Scale Data Analytics.
 504 Neural Networks, 2016. 77: p. 14-28
- 505 46. He, Q., X. Jin, C. Du, F. Zhuang, and Z. Shi, Clustering in extreme learning machine feature space. Neurocomputing, 2014. 128: p. 88-95
- 47. Bian, X., C. Zhang, X. Tan, M. Dymek, Y. Guo, L. Lin, B. Cheng, and X. Hu, *A boosting extreme learning machine for near-infrared spectral*
- 507 quantitative analysis of diesel fuel and edible blend oil samples. Analytical Methods, 2017. 9(20): p. 2983-2989
- 48. Zhu, Z., X. Zhu, F. Kong, and W. Guo, *A rapid method on identifying disqualified raw goat's milk based on total bacterial count by using dielectric spectra*. Journal of Food Engineering, 2018. 239: p. 40-51
- 510 49. Suganthan, P.N., Letter: On non-iterative learning algorithms with closed-form solution. Applied Soft Computing, 2018. 70: p. 1078-1082
- 50. Bisoi, R., P.K. Dash, and S.P. Mishra, *Modes decomposition method in fusion with robust random vector functional link network for crude oil price forecasting*. Applied Soft Computing, 2019. 80: p. 475-493
- 51. Zhang, L. and P.N. Suganthan, *Visual Tracking With Convolutional Random Vector Functional Link Network*. IEEE Transactions on Cybernetics,
 2017. 47(10): p. 3243-3253
- 515 52. Ren, Y., P.N. Suganthan, N. Srikanth, and G. Amaratunga, *Random vector functional link network for short-term electricity load demand* 516 *forecasting*. Information Sciences, 2016. **367-368**: p. 1078-1093
- 517 53. Selvaraju, R.R., M. Cogswell, A. Das, R. Vedantam, D. Parikh, and D. Batra, Grad-CAM: Visual Explanations from Deep Networks via
- 518 Gradient-Based Localization. International Journal of Computer Vision, 2017. 128: p. 336-359

- 54. Xia, J., Z. Wang, D. Yang, R. Li, G. Liang, H. Chen, A.A. Heidari, H. Turabieh, M. Mafarja, and Z. Pan, *Performance optimization of support* 520 *vector machine with oppositional grasshopper optimization for acute appendicitis diagnosis.* Computers in Biology and Medicine, 2022: p.
- 521 105206
- 522 55. Xia, J., D. Yang, H. Zhou, Y. Chen, H. Zhang, T. Liu, A.A. Heidari, H. Chen, and Z. Pan, *Evolving kernel extreme learning machine for medical*523 *diagnosis via a disperse foraging sine cosine algorithm.* Computers in Biology and Medicine, 2022. 141: p. 105137
- 524 56. Hu, J., z. Han, A.A. Heidari, Y. Shou, H. Ye, L. Wang, X. Huang, H. Chen, Y. Chen, and P. Wu, *Detection of COVID-19 severity using blood* 525 *gas analysis parameters and Harris hawks optimized extreme learning machine.* Computers in Biology and Medicine, 2022. **142**: p. 105166
- 526 57. Hu, J., Y. Liu, A.A. Heidari, Y. Bano, A. Ibrohimov, G. Liang, H. Chen, X. Chen, A. Zaguia, and H. Turabieh, *An effective model for predicting* 527 *serum albumin level in hemodialysis patients.* Computers in Biology and Medicine, 2022. 140: p. 105054
- 528 58. Zhang, Q., Z. Wang, A.A. Heidari, W. Gui, Q. Shao, H. Chen, A. Zaguia, H. Turabieh, and M. Chen, *Gaussian Barebone Salp Swarm Algorithm* 529 with Stochastic Fractal Search for medical image segmentation: A COVID-19 case study. Computers in Biology and Medicine, 2021. 139: p.
- 530 104941
- 531 59. Liu, L., D. Zhao, F. Yu, A.A. Heidari, J. Ru, H. Chen, M. Mafarja, H. Turabieh, and Z. Pan, *Performance optimization of differential evolution* 532 *with slime mould algorithm for multilevel breast cancer image segmentation.* Computers in Biology and Medicine, 2021. 138: p. 104910
- 60. Liu, L., D. Zhao, F. Yu, A.A. Heidari, C. Li, J. Ouyang, H. Chen, M. Mafarja, H. Turabieh, and J. Pan, *Ant colony optimization with Cauchy and greedy Levy mutations for multilevel COVID 19 X-ray image segmentation.* Computers in Biology and Medicine, 2021. 136: p. 104609
- 535 61. Yu, M., M. Han, X. Li, X. Wei, H. Jiang, H. Chen, and R. Yu, *Adaptive soft erasure with edge self-attention for weakly supervised semantic* 536 segmentation: *Thyroid ultrasound image case study*. Computers in Biology and Medicine, 2022. 144: p. 105347