Using Generative Adversarial Networks and Transfer Learning for Breast Cancer Detection by Convolutional Neural Networks

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ABSTRACT

In the U.S., breast cancer is diagnosed in about 12% of women during their lifetime and it is the second leading reason for women's death. Since early diagnosis could improve treatment outcomes and longer survival times for breast cancer patients, it is significant to develop breast cancer detection techniques. The Convolutional Neural Network (CNN) can extract features from images automatically and then perform classification. To train the CNN from scratch, however, requires a large number of labeled images, which is infeasible for some kinds of medical image data such as mammographic tumor images. In this paper, we proposed two solutions to the lack of training images. 1)To generate synthetic mammographic images for training by the Generative Adversarial Network (GAN). Adding GAN generated images made to train CNN from scratch successful and adding more GAN images improved CNN's validation accuracy to at most (best) 98.85%. 2)To apply transfer learning in CNN. We used the pre-trained VGG-16 model to extract features from input mammograms and used these features to train a Neural Network (NN)-classifier. The stable average validation accuracy converged at about 91.48% for classifying abnormal vs. normal cases in the DDSM database. Then, we combined the two deep-learning based technologies together. That is to apply GAN for image augmentation and transfer learning in CNN for breast cancer detection. To the training set including real and GAN augmented images, although transfer learning model did not perform better than the CNN, the speed of training transfer learning model was about 10 times faster than CNN training. Adding GAN images can help training avoid over-fitting and image augmentation by GAN is necessary to train CNN classifiers from scratch. On the other hand, transfer learning is necessary to be applied for training on pure real images. To apply GAN to augment training images for training CNN classifier obtained the best classification performance.

Keywords: breast mass classification, deep learning, convolutional neural networks, generative adversarial networks, transfer learning, mammogram, image augmentation, computer-aided diagnosis

1. INTRODUCTION

Breast cancer is the second leading cause of death among U.S women and will be diagnosed in about 12% of them ^{1,2}. The commonly used mammographic detection based on computer-aided detection (CAD) methods can improve treatment outcomes for breast cancer and increase survival times for the patients ³. These traditional CAD tools, however, have a variety of drawbacks because they rely on manually designed features. For example, hand-crafted features tend to be domain-specific, and the process of feature design can be tedious, difficult, and non-generalizable ⁴. In recent years, developments in machine learning have provided alternative methods for feature extraction; one is to learn features from whole images directly through a Convolutional Neural Network (CNN) ^{5,6}. Usually, training the CNN from scratch requires a large number of labeled images ⁷; for example, the AlexNet (a classical CNN model) was trained by using about 1.2 million labeled images ⁸. For some kinds of medical image data such as mammographic tumor images, however, to obtain a sufficient number of images to train a CNN classifier is difficult because the true positives are scarce in the datasets and expert labeling is expensive ⁹.

The shortcomings of an insufficient number of images to train a classifier are well-known^{8,10}, so it is worthwhile to apply **image augmentation** to create new training images and thus to improve the performance of a CNN classifier. Like CNN, the Generative Adversarial Network (GAN) is a state-of-the-art neural network-based learning technique in the field of deep learning ¹¹ introduced by Goodfellow *et al.* in 2014 ¹². Many novel applications in the field of image processing has been provided via GAN, for example, image translation ^{13,14}, object detection ¹⁵, super-resolution ¹⁶ and image blending ¹⁷. Also for the medical imaging, various GAN are also developed recently such as GANCS ¹⁸ for MRI reconstruction, SegAN ¹⁹, DI2IN ²⁰ and SCAN ²¹ for medical images. These generated images are not exactly like the original ones but could

keep the essential features, structures or patterns of the objects in original images. Therefore, GAN is a good candidate as such image augmentation method for augmenting the training dataset. We name the original images **ORG images** and the augmented images generated from GAN **GAN images** in the rest of this paper.

Another solution to deal with the lack of training images is to reuse a pre-trained CNN model that has been trained with very large image datasets from other fields as the feature extractor and re-train (fine-tune) such a model using a limited number of labeled medical images ²². This approach is also called **transfer learning**, which has been successfully applied to various computer vision questions ^{23–25}. In fact, some results of transfer learning are counterintuitive: previous studies for the pulmonary embolism and melanocytic lesion detection ^{22,26} show that the features (connection weights in the CNN) learned from natural images could be transferred to medical images, even if the target images greatly differ from the pre-trained source images.

Previous studies have applied various machine learning methods for breast cancer/tumor detection using mammograms ²⁷. The Digital Database for Screening Mammography (DDSM) ²⁸ are the most commonly used public mammogram databases. Some studies used the traditional automatic feature extraction (not manual extraction) techniques, such as Gabor filter, fractional Fourier transform and Gray Level Co-Occurrence Matrix (GLCM), to obtain features and then applied SVM or other classifier to do classification ^{29–33}. Neural networks were also used as classifiers ^{34,35}. And some studies applied CNN to generate features from mammographic images ^{36–39}. Some of these studies used pre-trained CNN as applications of transfer learning. In our study, we have tested both GAN for image augmentation and transfer learning to improve the performance of CNN classifier to breast cancer detection in mammograms. Specifically, we tested three training strategies on DDSM: 1) trained a CNN from scratch; 2) applied the pre-trained VGG-16 model ⁴⁰ to extract features from input images and used these features to train a Neural Network (NN)-classifier; 3) added GAN images in training set and repeated experiments in (1) and (2).

2. MATERIALS AND METHODS

2.1 The Mammogram Databases and Image Pre-processing

Mammography is the process of using low-energy X-rays to examine the human breast for diagnosis and screening. There are two main angles to get the X-ray images: the cranio-caudal (CC) view and the mediolateral-oblique (MLO) view. The goal of mammography is the early detection of breast cancer ⁴¹, typically through detection of masses or abnormal regions from the formed X-ray images. Usually, such abnormal regions are spotted by doctors or expert radiologists. In this study, we used mammogram from the Digital Database for Screening Mammography (DDSM) ²⁸. The DDSM is a widely used mammographic images resource by the U.S. Mammographic Image Analysis Research Community. It is a collaborative effort between Massachusetts General Hospital, Sandia National Laboratories and the University of South Florida Computer Science and Engineering Department. The DDSM database contains approximately 2,620 cases in total: 695 normal cases, 1925 abnormal cases (914 malignant\cancers cases, 870 benign cases and 141 benign without callback) with locations and boundaries of abnormalities. Each case includes four images representing the left and right breasts in CC and MLO views.

We downloaded all mammographic images from DDSM's official website (http://marathon.csee.usf.edu/Mammography /Database.html). Since images in DDSM are compressed in LJPEG format, to decompress and convert these images, we used the DDSM Utility ⁴². We converted all images in DDSM to PNG format. DDSM describes the location and boundary of actual abnormality by chain-codes, which are recorded in OVERLAY files for each breast image containing abnormalities. The DDSM Utility also provides the tool to read boundary data and display them for each image having abnormalities. Since the DDSM Utility tools run on MATLAB, we implemented all pre-processing tasks in MATLAB. We used the regions of interest of images (ROIs) instead of entire images to train our neural-network models. These ROIs are cropped rectangle-shape images and obtained by:

- For **abnormal ROIs** from images containing abnormalities, they are the minimum rectangle-shape areas surrounding the whole given ground truth boundaries.
- For **normal ROIs**, they are also rectangle-shape images and their sizes are approximately the average size of abnormal ROIs. In DDSM, the average size of abnormal ROIs is 506.02×503.90 pixels, so the cropping size for normal ROIs was chosen to be 505×505 pixels. Their locations are selected randomly on normal breast areas. In this study, we cropped only one ROI from an entire normal breast image.

The sizes of abnormal ROIs vary with abnormality boundaries. Since the neural-network models require all input images to be one specific size and the usual inputs for CNN are RGB images (images in DDSM are grayscale), we resized the ROIs by resampling and made them to RGB (3-layer cubes) by duplication (Fig. 1). These images cropped from mammogram are **ORG ROIs**.

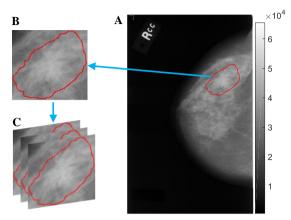


Fig. 1. (A) A mammographic image from DDSM rendered in grayscale; (B) Cropped ROI by the given truth abnormality boundary; (C) Convert Grey to RGB image by duplication.

2.2 GAN Image Augmentation

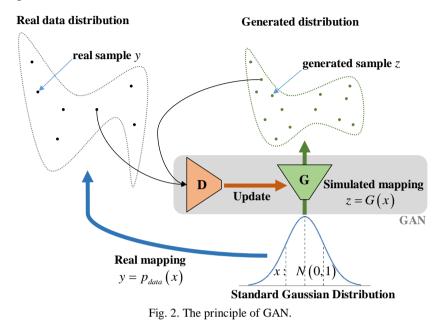
The GAN is a neural-network-based generative model that learns the probability distribution of real data and creates simulated data samples with a similar distribution (Fig. 2). Formally, in *d* -dimension space, for $x \in \mathbb{R}^d$, $y = p_{data}(x)$ is a mapping from *x* to real data *y*. We create a neural network called the **generator** *G* to simulate this mapping. If sample *y* comes from p_{data} , it is a real one; and sample *z* comes from *G*, it is a synthetic one. Another neural network **discriminator** *D* is used to detect whether a sample is real or synthetic. Ideally, D(y)=1; D(z)=0. The two neural networks *G* and *D* compose the GAN. We can find *G* and *D* by solving the two-player minimax game ¹², with value function V(G,D):

$$\min_{G} \max_{D} V(G, D) = \mathbb{E} \Big[\log D \Big(p_{data} (x) \Big) \Big] + \mathbb{E} \Big[\log \Big(1 - D \big(G (x) \big) \Big) \Big]$$

This min-max problem has a global optimum (Nash equilibrium) solution for $G(x) = p_{data}(x)$. That is the goal to find the distribution of real data. At equilibrium, discriminator D can no longer distinguish the real from the synthetic sample, where D(y) = D(z) = 0.5. Synthetic samples can be generated from G by changing the input x. In this study, the input x for G we used was a noise vector having 100 elements from a Gaussian distribution : N(0,1). The key point of a well-trained GAN is that it could generate seemingly real-like data samples by giving noise vectors. To train a GAN, we used limited number of real samples. Ideally, GAN could generate unlimited different synthetic samples.

To implement GAN, we built the generator and discriminator neural networks. The details about their structures show in

Table 1. The generator consists of four up-sampling layers to double the size of image and five convolutional layers. The activation function for each layer is the ReLU function ⁴³ except the last one for output, which is tanh function. The function of generator is to transform a 100-length vector to a 320x320x3 image. The input of discriminator is a 320x320x3 image and its output is a value between 0 and 1, which '0' stands for the synthetic image and '1' for the real one. Like a typical CNN, the discriminator has four convolutional layers with max-pooling layers and one FC layer. The activation function for each convolutional layer is also the ReLU function and the last one for output is sigmoid function, which maps the output value to the range of [0, 1].



The notation Conv_3-32 means there are 32 convolutional neurons (units) and the filter size in each unit is 3×3 -pixel (height × width) in this layer. MaxPool_2 means a max-pooling layer with size of filters is 2×2 -pixel window, stride 2. And FC_n means a fully-connected layer having *n* units. Dropout layer ⁴⁴ randomly set a fraction rate of input units to 0 for the next layer at every updating during training; it could help the networks avoid overfitting. Our training optimizer is Nadam ⁴⁵ using default parameters (except the learning rate changed to 1e-4), the loss function is Binary Cross Entropy, the updating metric is Accuracy, the batch size is 30 and the number of total epochs is set to be 1e+5.

The training methods of GAN are:

- Step 1: Randomly initialize all weights for both networks.
- Step 2: Input a batch of 100-length noise vectors to generator to obtain synthetic images.
- Step 3: To train the discriminator by a batch of synthetic images labeled '0' and real images labeled '1'.
- Step 4: To train the generator: input a batch of 100-length noise vectors to generator to obtain synthetic images and label them as '1'. Then, input these synthetic images to discriminator to obtain the predicted labels. The differences between predicted labels and '1' will be the loss for updating the generator. It is noteworthy that in this step, only the weights in generator are changed; weights in discriminator are fixed.
- Step 5: Repeat Step 2 to Step 4 until all real images have been used once, that counts one **epoch**. When the number of epochs reaches a certain value, training stops.

Actually, for the Step 5, the ideal situation to stop training is when the classification accuracy of discriminator converges to 50%. It means the discriminator no longer can distinguish the real images and synthetic images generated from a well-trained generator. The discriminator plays a role as an assistant in GAN. After training, we will use the generator neural networks to generate synthetic images for usage next.

Generator			
Layer	Shape		
input: 100-length vector	100		
FC_(256x20x20) + ReLU	102400		
Reshape to 20x20x256	20x20x256		
Normalization + Up-sampling	40x40x256		
Conv_3-256 + ReLU	40x40x256		
Normalization + Up-sampling	80x80x256		
Conv_3-128 + ReLU	80x80x128		
Normalization + Up-sampling	160x160x128		
Conv_3-64 + ReLU	160x160x64		
Normalization + Up-sampling	320x320x64		
Conv_3-32+ ReLU	320x320x32		
Normalization + Conv_3-3+ ReLU	320x320x3		
output (tanh): [-1, 1]	320x320x3		

Discriminator			
Layer	Shape		
input: RGB image	320x320x3		
Conv_3-32 + ReLU	320x320x32		
MaxPooling_2 + Dropout (0.25)	160x160x32		
Conv_3-64 + ReLU	160x160x64		
MaxPooling_2 + Dropout (0.25)	80x80x64		
Conv_3-128 + ReLU	80x80x128		
MaxPooling_2 + Dropout (0.25)	40x40x128		
Conv_3-256 + ReLU	40x40x256		
MaxPooling_2 + Dropout (0.25)	20x20x256		
Flatten	102400		
FC_1	1		
output (sigmoid): [0, 1]	1		

2.3 To Train the CNN from Scratch

Actually, a CNN was designed as the discriminator in GAN and its function is to distinguish real and synthetic mammographic ROIs. We also built a CNN to classify abnormal ROIs and normal ROIs. As shown in Table 2, this CNN classifier consists of three convolutional layers with max-pooling layers and two FC layers. The activation function for each layer is the ReLU function except the last one for output. The output layer uses a sigmoid function, which maps the output value to the range of [0, 1]. Its input is the image in size 320×320-pixel. Since the sigmoid function was used in the output layer, the predicted outcome from the CNN classifier is a value between 0 and 1. By default, the classification threshold is 0.5, meaning that if the value is less than 0.5 it will be considered as "0" (normal), otherwise it will be considered as "1" (abnormal). The optimizer for training is Nadam using default parameters ⁴⁵ (except the learning rate changed to 1e-4), the loss function is Binary Cross Entropy, the updating metric is Accuracy, the batch size is 26 and the number of total epochs is set to be 750. To train this CNN classifier from scratch, we used the labeled ROIs of abnormal and normal mammographic images.

CNN classifier			
Layer	Shape		
input: RGB image	320x320x3		
Conv_3-32 + ReLU	320x320x32		
MaxPooling _2	160x160x32		
Conv_3-32 + ReLU	160x160x32		
MaxPooling _2	80x80x32		
Conv_3-64 + ReLU	80x80x64		
MaxPooling _2	40x40x64		
Flatten	102400		
FC_64 + ReLU + Dropout (0.5)	64		
FC_1	1		
output (sigmoid): [0, 1]	1		

Table 2. The architecture of CNN classifier.

2.4 Transfer Learning: Features Extraction by Pre-trained VGG-16 network

The structure of CNN in transfer learning was combined the 13 convolutional layers in pre-trained VGG-16 model ⁴⁰ with a simple FC layer (Table 3).

CNN classifier with Transfer Learning				
	Layer			
	input: RGB image			
		Conv_3-64 + ReLU		
	Conv block 1	Conv_3-64 + ReLU		
_		MaxPool_2		
		Conv_3-128 + ReLU		
	Conv block 2	Conv_3-128 + ReLU		
_		MaxPool_2		
		Conv_3-256 + ReLU		
	Conv block 3	Conv_3-256 + ReLU		
VGG-16		Conv_3-256 + ReLU		
100-10		MaxPool_2		
		Conv_3-512 + ReLU		
	Conv block 4	Conv_3-512 + ReLU		
		Conv_3-512 + ReLU		
_		MaxPool_2		
	Conv block 5	Conv_3-512 + ReLU		
		Conv_3-512 + ReLU		
		Conv_3-512 + ReLU		
		MaxPool_2		
$FC_{256} + ReLU$ (with Dropout = 0.5)				
output (sigmoid): [0, 1]				

Table 3.	CNN architecture	for transfer	learning
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As shown in Table 3, all the weights in five convolutional blocks (the blue background layers) were imported from the pre-trained VGG-16 model and not changed (or called weights frozen) during the training of this CNN. Only weights in the FC layer were randomly initialized and updated by training. Thus, such training process can be seen as that the VGG-16 extracts features from input image and then these features were used to train a FC NN-classifier.

3. EXPERIMENT AND RESULTS

Our implementation of neural networks was on the Keras API backend on TensorFlow ⁴⁶. The development environment for Python was Anaconda3.

3.1 Experiment Plan

Table 4. Notations for data.

Set name	Notation for element	Meaning
ORG ROIs	O_{abnorm} / O_{norm}	Real abnormal/normal ROI
GAN ROIs	$G_{_{abnorm}}/G_{_{norm}}$	Synthetic abnormal/normal ROI by GAN

In this study, we collected 1300 original (real) abnormal ROIs (O_{abnorm} , 'O' for original) and 1300 original normal ROIs (O_{norm}) in total. After taking off 10% for validation, there are 1170 O_{abnorm} and 1170 O_{norm} . We firstly did the data augmentation to 1170 O_{abnorm} and 1170 O_{norm} by GAN. We used the 1170 O_{abnorm} and 1170 O_{norm} to train two generators respectively: GAN_{abnorm} and GAN_{norm} for generating GAN ROIs. As shown in Fig. 3 (GAN box), during the training process, the generator G provided synthetic ROIs to discriminator $D \cdot D$ was trained to distinguish the real from the synthetic ROIs by real and synthetic ROIs. And once synthetic ROIs were distinguished, D gave feedback loss to G for G's updating. Then G will generate synthetic ROIs more like the real ones. By inputting noise vectors to GAN_{abnorm} and GAN_{norm}. Fig. 4 shows some synthetic abnormal ROIs (G_{abnorm}) generated from GAN_{abnorm}

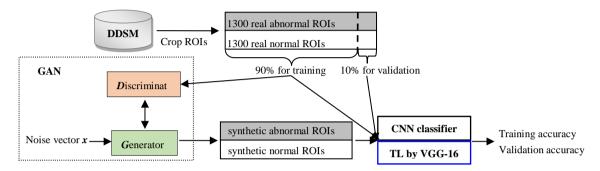


Fig. 3. The flowchart of our experiment plan. CNN classifiers were trained by data including ORG and GAN ROIs. Validation data for the classifier were ORG ROIs that were never used for training. The ORG and GAN ROIs were also used to Transfer Learning by pre-trained VGG-16 model.

Table	5.	Training	plans.
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Set#	Dataset for training	Validation	Classifier Model	
1	$\frac{1170 O_{abuorm}}{0000} \text{ labeled '1'}$ $\frac{1170 O_{norm}}{00000} \text{ labeled '0'}$			
2	$1170 G_{abnorm} \text{ labeled '1'}$ $1170 G_{norm} \text{ labeled '0'}$	$130 O_{abnom}$ labeled '1'	CNN in Table 2	TL model in Table 3
3	$1170 O_{abnorm} + 1170 G_{abnorm} \text{ labeled '1'}$ $1170 O_{norm} + 1170 G_{norm} \text{ labeled '0'}$	130 O _{norm} labeled '0'	CIVIN III TADIE 2	
4	$1170 O_{abnorm} + 2340 G_{abnorm} \text{ labeled '1'}$ $1170 O_{norm} + 2340 G_{norm} \text{ labeled '0'}$			

We repeated training the **CNN classifier** and the **transfer learning (TL) model** from scratch using different datasets of labeled ROIs shown in Table 5. During the training, there was no any data augmentation applied. In each set, the number of abnormal ROIs and normal ROIs is equal. We used 130 O_{abnorm} and 130 O_{norm} that were never used in the training process as validation data to evaluate those CNN classifiers. We generated 1170 G_{abnorm} and 1170 G_{norm} from GAN for training Set 2 and combined with 2340 ORG ROIs for Set 3. In the Set 3, the number of ORG ROIs and GAN ROIs are equal. In addition, we generated double number of GAN ROIs as ORG ROIs and put them together in Set 4.

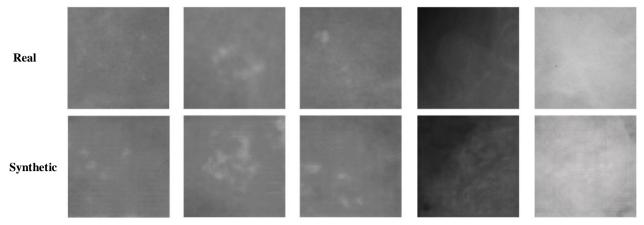


Fig. 4. (Top row) Real abnormal ROIs; (Bottom row) synthetic abnormal ROIs generated from GAN.

3.2 Classification Results

Specifically, to train GAN, we used 1170 O_{abnorm} to obtain the generator GAN_{*abnorm*}, and used 1170 O_{norm} to obtain the generator GAN_{*norm*}. Fig. 4 shows some synthetic abnormal ROIs (G_{abnorm}) generated from GAN_{*abnorm*}. Then, we generated G_{abnorm} and G_{norm} by generators.

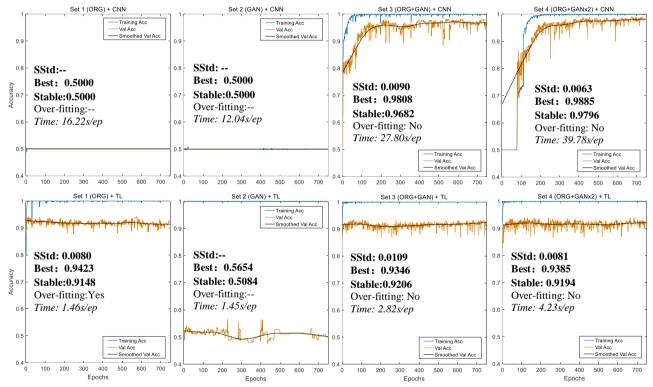


Fig. 5. Training accuracy and validation accuracy for four training datasets.

The results of training accuracy and validation accuracy after each training epoch (it is defined in 2.2, training methods, Step 5; the total epochs are 750) are shown in Fig. 5. By looking the figures, Set 3 and 4 perform well and Set 1 and 2 are worse except Set 1 using transfer learning. To analyze those results quantitatively, we show the <u>stable standard deviation</u> (SStd, which is the standard deviation of validation accuracy after 600 epochs), <u>maximum validation accuracy</u> (Best), <u>average validation accuracy after 600 epochs</u> (Stable) and <u>whether the over-fitting occurred</u>. The maximum validation

accuracy can indicate the **best performance** of the classifier, but it may be reached fortuitously. The average validation accuracy after 600 epochs can show the **stable performance** of the classifier. For a good classifier, this value will be monotone increasing and converged. And **SStd** shows how validation accuracy varies from its average after 600 epochs. The criterion for **occurrence of over-fitting** is defined by the value: average validation accuracy after 400 epochs minus (-) average validation accuracy before 400 epochs; if it is negative, then we consider that the over-fitting occurred because of the decreasing of validation accuracy during training.

Since the maximum validation accuracy may be fortuitous, the stable performance during training is more reliable to evaluate a classifier. The results in Fig. 5 demonstrate that:

- Pure ORG ROIs or GAN ROIs cannot train the CNN classifier successfully. To train CNN from scratch, adding GAN ROIs made the CNN classifier training successful. Additionally, by comparing the two results of Set 3 and Set 4, more added GAN ROIs improved CNN's performance.
- By comparing the two results of Set 1, transfer learning (TL) model successfully improved the accuracy of classification a lot. But to compare TL used to Set 1 and Set 2, pure GAN ROIs also cannot train the transfer learning model successfully.
- To transfer learning model, by examining Set 1, 3 and 4, adding GAN ROIs did not improve validation accuracies very much, however, prevented the training from over-fitting.
- By comparing the results of CNN and TL for Set 3 and 4, adding GAN ROIs have more benefit to improved CNN's performance remarkably than to the TL. TL has the advantage on speed TL was running about 10 times faster than CNN.

Overall, to train TL by only ORG ROIs, the validation accuracy is as good as training by adding GAN ROIs. But adding GAN ROIs can help avoid over-fitting. Image augmentation by GAN is necessary to train CNN classifiers from scratch. On the other hand, TL is necessary to be applied for training on pure ORG ROIs. To apply GAN to augment training images for training CNN classifier obtained the best classification performance. Then, to decrease the time cost of training, TL could be also applied to the augmented dataset.

4. DISCUSSION

As we discussed in Section 2.2, the ideally theoretical outcome of GAN is $G(x) = p_{data}(x)$. If so, the performance of CNN classifier trained by GAN ROIs will be as good as by ORG ROIs. Our results, however, show that GAN did not correspond with theoretical expectations. Opposite to ORG ROIs, pure GAN ROIs cannot train the transfer learning model successfully by comparing TL used to Set 1 and Set 2. The problem could be found by looking the synthetic images (Fig. 4): they have clear artificial flavors. One possible reason is that GAN adds some features or information not belonging to real images. Those new features disturb classifiers to detect abnormal features in real images. The possible solution is to change the architecture of generator or/and discriminator in GAN. In this paper, the architecture we used is DCGAN ⁴⁷. Recently, there are about 500 architectures of GAN ⁴⁸. We believe that some of them can achieve a better performance to train CNN from scratch.

We reviewed several recent studies highly related to ours. These studies (Table 6) applied transfer learning in CNN to detect breast cancer/abnormality based on mammogram. By comparison with these studies, we used many more mammographic images for training and testing the CNN classifiers and a distinct pre-trained model. The main difference is about the classifier and image augmentation by GAN. Our one-FC layer NN-classifier has simpler architecture and could be integrated with pre-trained convolutional layers as one complete CNN. The <u>stable classification accuracies</u> of our proposed model for abnormal vs normal cases on mammograms are competitive to other studies.

Since the GAN were introduced, it has been widely used in many image processing applications ¹¹. In medical imaging, many applications of GAN are segmentation ^{19,21,49–52}. And some studies are about medical image simulation/synthesis ^{53–57}. Image synthesis is a specialty or advantage of GAN, hence, it is apt to apply GAN as an image augmentation method ⁵⁸ for training classifiers and improving their detection performances. As far as we aware there is no study about using GAN as data augmentation method on mammogram to train CNN classifier or transfer learning model for breast cancer detection. Therefore, our study fills this gap.

Since the DDSM provides truth labels for benign and malignant tumors, in future works, we could also do classification for benign and malignant ROIs instead of abnormal and normal ROIs. We could try to recognize the abnormal areas in whole mammographic images. By using the RCNN ⁵⁹, we could recognize the abnormalities on mammographic images and draw boundaries (or rectangle region proposals) on such areas automatically. These regions do not have to be very high accuracy because they just provide another kind of reference for doctors to make decisions. We could use other pre-trained models and compare to their performances. In the research field of deep learning, VGG-16 appeared early but its depth (total number of layers is 23) is relatively shallow compared to new models, such as InceptionV3 (159 layers) ⁶⁰, ResNet50 (168 layers) ⁶¹ and InceptionResNetV2 (572 layers) ⁶². It will be interesting to see performances of breast cancer detection by using very deep CNNs. And we may also examine performances of other architectures of GAN in terms of image augmentation.

Main method	# of images	Accuracy %
Pre-trained CNN on LSVRC datasets & Fine-tuning + Two-step decision ³⁷	600	(Ben-Mal) 96.7
Pre-trained CNN with hand crafted features + RF ³⁸	410	(Ben-Mal) 91.0
Pre-trained AlexNet +Sparse MIL ³⁶	410	(Mal-nonMal) 90.0
Pre-trained VGG-16 + one FC layer by ORG ROIs (Ours)	2600	(Abnorm-Norm) 91.5
Pre-trained VGG-16 + one FC layer by ORG+GAN ROIs (Ours)	2600	(Abnorm-Norm) 92.1
CNN by ORG + <i>double</i> GAN ROIs (Ours)	2600	(Abnorm-Norm) 98.0

Table 6. Comparison of related studies.

5. CONCLUSIONS

In this paper, we proposed GAN to be used as an image augmentation method for training and to improve the performance of CNN classifiers. Our results show that, to classify the normal ROIs and abnormal (tumor) ROIs from DDSM, adding GAN generated ROIs in training data can help the classifier prevent from over-fitting and the validation accuracy using mixture ROIs reached at most (best) 98.85%. Therefore, GAN could be promising image augmentation method. To transfer learning in CNN for breast cancer detection, our results show that the pre-trained CNN model (VGG-16) can automatically extract features from mammographic images, and a good NN-classifier (achieves stable average validation accuracy about 91.48% for classifying abnormal vs. normal cases in the DDSM database) can be trained by only real ROIs. In addition, we have done the study of combining the two deep-learning-based technologies together. That is to apply GAN for image augmentation and then use transfer learning in CNN for detection. Although to train the transfer learning model by adding GAN ROIs did not perform better than to train the CNN by adding GAN ROIs, the speed of training transfer learning model was about 10 times faster than CNN training. In summary, <u>adding GAN ROIs can help training avoid over-fitting and image augmentation by GAN is necessary to train CNN classifiers from scratch. On the other hand, transfer learning is necessary to be applied for training on pure ORG ROIs. To apply GAN to augment training images for training CNN classifier obtained the best classification performance.</u>

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