Model Fooling Threats Against Medical Imaging*

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Abstract Automatic medical image diagnosis tools are vulnerable to modern model fooling technologies. Because medical imaging is a way of determining the health status of a person, the threats could have grave consequences. These threats are not only dangerous to the individual but also threaten the patients' trust in modern diagnosis methods and in the healthcare sector as a whole. As recent diagnosis tools are based on artificial intelligence and machine learning, they can be exploited using attack technologies such as image perturbations, adversarial patches, adversarial images, one-pixel attacks, and training process tampering. These methods take advantage of the non-robust nature of many machine learning the probability of cancerous cell growth in tissue samples. In this study, we review the current state of these attacks and discuss their effect on medical imaging. By comparing the known attack methods and their use against medical imaging, we conclude with an evaluation of their potential use against medical imaging.

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1 Introduction

The goal of the research is to examine the literature related to potential model fooling attacks against medical imaging, with digital pathology as the main interest. In the modern digitalised world, Artificial Intelligence (AI) based solutions are utilised extensively in everyday life. For example, paper [38] introduces an AI-based healthcare assistant. Heart functioning is analysed and predicted with neural networks by using electrocardiogram (ECG) data in the studies [40, 41] and similarly, electroencephalogram (EEG) data is analysed by AI for detecting brain tumors [33]. Syam and Marapareddy used deep neural networks for network intrusion detection, heart disease prediction and for skin cancer classification [52].

The usage of sub-disciplines of AI, Machine learning (ML) and Deep Learning (DL) based solutions is rapidly increasing in the medical imaging for prediction and decision making by itemizing and labeling disease patterns from image samples [25]. The large amounts of available information makes the medical domain very interesting for researchers so that new applications can be developed [45]. The tremendous development of medical imaging has produced advances in diagnostics and prediction of diseases [13, 3]. The benefit achieved by the DL in the analysis of modern medical big data is the capability for algorithmic realisation of the various associations and capability to combine learned lines or edges of low level to the higher-level shapes [21].

The vast development of machine learning has produced several modern examples of applying ML/DL for the medical imaging as computer-aided diagnosis (CAD) tools. Comprehensive review for ML in medicine is presented by authors of paper [39]. Hussein et al. studied lung and pancreatic tumor characterization with DL whereas Lu et al. utilised ensemble learning with data mining for predicting recurrent ovarian cancer. Among others, during this year, utilisation of ML/DL for medical image classification and detection is studied for example with brain tumors in [43, 49, 42] and breast cancer in [11, 44, 37]. It should also be noticed that developing AI for healthcare is a highly technical subject but in addition with usage of AI for healthcare there are ethical, legal and social challenges involved such as 'Data ownership, confidentiality and consent' or 'Medical moral and professional responsibility' [9].

Modern networked and digitalized cyber domain is an extremely complex entity that comprises unpredictable phenomena. A classical example of that complexity is a cyber attack against an electricity company, which may endanger the patient safety of the hospital. Finland's cyber security strategy [46] classifies healthcare as an area vulnerable to cyber security issues and states that these issues will be more important in the future. As known, there are several cyber attacks executed globally against healthcare infrastructure, and healthcare infrastructure is seen as valuable target for cyber attacks or an intrusion. The International Criminal Police Organization (INTERPOL) states that cyber attacks' target is shifting towards governments and critical health infrastructure during the ongoing COVID-19 pandemic [20].

As can be seen, ML/DL applications are widely applied in the medical imaging and simultaneously, the overall medical cyber domain is realised as a potential target for the cyber attacks. In this regard, our study focuses on the model fooling threats against medical imaging. During this study following threat categories were identified: (i) adversarial images, (ii) adversarial patches, (iii) one-pixel attacks, (iv) training process tampering and (v) generating fake data.

We investigated literature related to the possible cybersecurity threat vectors using using a scoping review method. According to Munn et al. scoping review is a suitable method for the search for scientific gaps in the research area, or building the knowledge base or the synthesis of literature to confirm the research results [32]. In this paper, the point of scoping review is to seek support from previous research for the findings of this research, thus building a stronger knowledge base for the phenomenon. In the scoping review, we used Google Scholar and IEEE databases. Searches were performed by using the following search parameters: fooling neural networks, adversarial attack / adversarial example and medical imaging. Studies in English related to the medical domain were selected. Furthermore, studies with actual applications of the attacks were included. In addition, some essential methodology studies are mentioned.

This article is an extension of a short survey originally presented in the Second International Scientific Conference "*Digital Transformation, Cyber Security and Resilience*" (DIGILIENCE 2020) and published in the special conference issue of Information & Security: An International Journal [48]. This research is expanded from the original as follows: we have identified more publications that are essential related to the topic and presented them in a new manner. In addition, we have restructured this paper to better reflect the contents of the identified research literature.

The paper is organised as follows: Fooling neural networks is introduced in section 2. The specific categories of attacks are discussed in section 3 and its subsections. The research is concluded with the found future research topics in section 4.

2 Fooling Deep Neural Networks in Medical Imaging

Deep learning tries to combine simple concepts into a representation of the actual object. This is conducted by creating an artificial neural network of interconnected nodes [17]. The complex nature of these networks makes them susceptible to unexpected attacks, which force the network to output completely reverse results that are unlike the expected outcome. A reverse result in medical imaging could be harmful to the patient.

Adversarial attacks against deep learning image classifiers are plentiful. In a white-box attack, the attacker knows the internal workings of the classifiers. This is usually useful when using the neural network gradient as a way of finding adversarial examples. On the other hand, black-box attacks are performed against a system that has only its image input and classification result exposed to the attacker. The attack methods use optimization to find examples that produce the most diverging classification scores [56]. Computer vision is especially affected by these threats because deep neural networks are the most prominent method. Akhtar and Mian

estimate that the Carlini & Wagner [8] and Universal perturbations [30] are the strongest methods. Both are white-box attacks, so they need the complete knowledge of the inner workings of the target classifier [2]. Furthermore, Afifi et al. demonstrate that simple color constancy errors can change the classification of a natural image [1].

Since many methods use a gradient as the guiding principle for the optimization, gradient masking and obfuscation could help to defend against these attacks. This would mislead the attacks or make the attack optimization very difficult to achieve. Another defence method is the use of robust optimization. Robust classifiers are less likely to behave in an unexpected manner, such as falling for an adversarial image. This could be achieved, e.g., with adversarial retraining. The third defence could be adversarial example detection before the input images are fed to the real classifier [56, 27]. Tizhoosh and Pantanowitz mention adversarial attacks as one of the challenges facing digital pathology. They raise the question whether minimal artifacts could reduce the reliability of neural network classifiers. This might be caused by the old problem of overfitting in artificial intelligence [54]. Akhtar and Mian propose three ways of defending against adversarial attacks. Firstly, modified training during learning or modified input during testing can be used. Secondly, they suggest modifying deep neural networks and their architecture. Thirdly, for unseen examples, an external model could be used to act as a network add-on [2]. The point of intervention and defence against these attacks is also a problem to be solved, which will probably need regulatory best practices since the problem resembles that of trying to counteract ever developing hacking attempts [14]. A recent survey by Apostolidis and Papakostas on adversarial attacks against medical image analysis discusses the robustness of deep neural networks. It identifies many image modalities that have been attacked: X-ray images, magnetic resonance imaging (MRI), computer tomography scans (CT), retinal images, histology and skin. In addition to the modalities, the survey lists attacks, their target models, detection methods and defences. The authors emphasize the need for robust models in automated medical imaging [4].

3 Attack Types

Based on the literature introduced in this study, Figure 1 shows the most obvious attack vectors against medical imaging neural networks. The two proposed attack vectors are changing the training process to create a faulty AI model and modifying the input images, so that the classification fails even with a correctly working AI model.

There are several ways to attack against medical imaging. The main methods can be categorized as (i) adversarial images, (ii) adversarial patches, (iii) one-pixel attack, (iv) training process tampering and (v) generating fake data. Table 1 shows the identified attack methods and their use against medical imaging. The following

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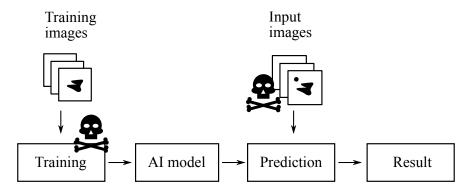


Fig. 1 The most prominent attack vectors described in literature. Tampering with training compromises the automated detection pipeline from the beginning. Modifying input images is perhaps the easier attack method and compromises the results of automated detection.

subsections discuss each of these methods in more detail, introducing the methods themselves, and discussing their applications.

Table 1 Adversarial methods against artificial neural networks, and their implementations in themedical domain. The *References* column shows general references about the methods, while theMedical domain column shows applications.

Method	References	Medical domain
Adversarial images	[34], [31], [6], [28], [12], [19]	[35], [53], [15], [26]
Adversarial patches	[7]	[15]
One-pixel attack	[51], [50], [24], [55]	[36], [47], [23], [22]
Training tampering	[18], [57]	
Generating fake data		[29], [10], [5]

3.1 Adversarial Images

Adversarial images are images that are somehow changed by adding perturbation to create a misclassified image. As shown by Nguyen et al., it is possible to produce images that are unrecognizable to humans, but that are classified with 99.99% confidence by deep neural networks. Firstly, their adversarial examples include pictures that resemble noise generated by an evolutionary algorithm using direct encoding. Secondly, their other adversarial examples resemble wave patterns and lattices, which have been created by an evolutionary algorithm using indirect encoding. Their evolu-

tionary optimization uses the classifying deep neural network as the fitness function, which makes the approach a black-box method. [34]

Moossavi-Dezfooli et al. present the DeepFool algorithm that finds perturbations to deceive deep neural networks. They use a gradient descent algorithm to find those perturbations. The combination of an image and the perturbation is falsely classified as representing something that it does not. [31] Athalye et al. raise the question that viewpoint shifts, camera noise, and transformations can make adversarial examples less effective. They created a 3D-printed turtle that is classified as rifle from images taken of it in the physical world. The optimization process takes into account the expectation of transformation, which creates more robust adversarial examples. [6]

Some other examples of adversarial images include those generated using adversarial noise [28], using a generative approach to fool black-box classifiers [12] and gradient shielding to identify sensitive regions where attacks could be executed [19].

Medical images have been used as targets for these kinds of adversarial images. Paschali et al. studied neural network performance under extreme inputs such as noise, outliers, and ambiguous data. They used fast gradient sign, DeepFool and saliency map attacks to create the adversarial images. They performed the attacks on skin lesion images and whole brain imaging [35]. Taghanaki et al. used three types of adversarial attacks: gradient-based, score-based and decision-based. These added perturbations to X-ray images producing images that look quite natural in some cases [53]. Finlayson et al. used projected gradient descent to create visually unnoticeable perturbations against fundoscopy, chest X-ray, and dermoscopy images [15].

Ma et al. created adversarial images in medical imaging domain using unnoticeable perturbations. They go on to claim that medical images can be more vulnerable than natural images in this context. Firstly, they suggest that medical images have larger high attention regions, which draw unnecessary attention from the neural network. Secondly, modern neural networks are designed for natural images, causing them to overparametrize for medical images. Furthermore, a simple adversarial image detector classifier is sufficient to protect the actual classifier from most of the attacks. [26].

3.2 Adversarial Patches

Adversarial patches can be applied onto images to output any target class. These patches can be natural, meaning a cut-and-pasted part of an existing image, or generated using optimization, resulting in wild-looking but successful patches when applied. According to Brown et al., even small patches can shift the focus of the classifier to the patch and change the classification of the scene. Suitable patches are found with similar optimization as with adversarial images. [7]

There have been examples of adversarial patches used against medical imaging. Finlayson et al. demonstrated that this method works against fundoscopy, chest Xray, and dermoscopy images. Furthermore, they tested natural patches, patches built on the victim model and patches built on another independent model later used as attacks against the victim model. [15]

3.3 One-pixel Attacks

One-pixel attack means that the alteration of color values of a single pixel will cause misclassification. Su et al. have shown that this extremely limited attack is successful against natural images. They use differential evolution optimization against the blackbox classifier to find successful one-pixel examples [51]. Furthermore, they propose a variation of the attack with multiple objectives [50]. Gilmer et al. propose that small perturbations are adversarial against machine learning models because of the high-dimensional geometry of the data manifold. [16]

Kügler et al. created simple problems about pose estimation of surgical tools in order to localize areas where one-pixel attacks were lucrative. They discovered that the vulnerable areas of the image are close to the decision boundary. [24]

Vargas et al. propose propagation maps to illustrate how much the perturbations affect neural network layers. They discovered that complex neural networks let the single pixel propagate widely causing it to create unreasonable consequences to the classification result. Attacks against pixels located near the successful attacks are also quite effective. [55]

There are not many examples of one-pixel attacks against real medical imaging data. Paul et al. attacked against the National Lung Screening Trial (NLST) dataset using a one-pixel attack. They also used fast gradient signed method (FGSM) attack, which was more successful. They applied an ensemble defence strategy to create more robust classifiers [36]. The concept of using one-pixel attacks against whole slide images was explored by Sipola and Kokkonen [47], and implemented by Korpihalkola et al. using an existing database of those images [23]. The attack was refined by optimizing the color so that the adversarial pixels would be less prominent to the human eye [22].

3.4 Training Process Tampering

A backdoored neural network has been trained with malicious training material that causes it to react in unexpected ways when given specific input. The act of infiltrating training data with malicious samples is called poisoning. Yang et al. used direct gradient method and auto-encoders to generate poisoned data for neural network training. [57] Gu et al. present this idea of including a hidden backdoor detector inside the classifier by using crafted training data. They demonstrate this threat using traffic signs, which causes the classifier to detect a stop sign as a speed limit sign. [18]

3.5 Generating Fake Data

Not all applications of adversarial methods are malicious. Generative adversarial networks can also be used for synthesizing data samples [29]. Another application is to use generative adversarial methods to inpaint medical images that contain areas of missing data [5]. Another kind of proof of the power of adversarial images is that they can fool human experts. Chuquicusma et al. have shown that images produced by generative adversarial networks can fool radiologists [10]. Even if this is not an attack against a diagnosis tool, it shows the potential of generating fake data.

4 Conclusion

Machine learning based solutions are successfully used in healthcare, especially in the medical imaging for prediction and decision making in case of a potential tumor. Medical imaging and analysis methods are not safe from model fooling attacks. Suitable research exploits have been shown to successfully fool neural network models in this domain. The most prominent methods are (i) adversarial images, (ii) adversarial patches, (iii) one-pixel attacks, (iv) training process tampering and (v) generating fake data. The first three (i)-(iii) main types of attacks against medical imaging are present in the scientific studies included in this review. In addition, generating fake data (v) for non-exploitative purposes was identified. Although it might seem that these attacks are quite elaborate, with a suitable target system and with a high value patient, an attacker could find it worthwhile to use an adversarial attack. Based on the conducted scoping review, future research could include a comprehensive systematic literature review of the phenomenon, especially for specific imaging modalities or attack methods. Further investigation needs to be focused on the deep neural network methods used in medical classifiers. The underlying causes and robustness of those networks are not yet apparent and the theoretical considerations still remain unresolved.

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