# Revisiting Hidden Representations in Transfer Learning for Medical Imaging

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### **Abstract**

While a key component to the success of deep learning is the availability of massive amounts of training data, medical image datasets are often limited in diversity and size. Transfer learning has the potential to bridge the gap between related yet different domains. For medical applications, however, it remains unclear whether it is more beneficial to pre-train on natural or medical images. We aim to shed light on this problem by comparing initialization on ImageNet and RadImageNet on seven medical classification tasks. We investigate their learned representations with Canonical Correlation Analysis (CCA) and compare the predictions of the different models. Our results show that, contrary to intuition, ImageNet and RadImageNet converge to distinct intermediate representations, and that these representations are even more dissimilar after fine-tuning. Despite these distinct representations, the predictions of the models remain similar. Our findings challenge the notion that transfer learning is effective due to the reuse of general features in the early layers of a convolutional neural network and show that weight similarity before and after fine-tuning is negatively related to performance gains.

#### 1 Introduction

Transfer learning has become an increasingly popular approach in medical imaging, as it offers a solution to the challenge of training models with limited dataset sizes. The ability to leverage knowledge from pretrained models has proven to be beneficial in various medical imaging applications (Raghu et al., 2019; Cheplygina et al., 2019; Shin et al., 2016). Despite its widespread use, the precise effects of transfer learning on medical image classification are still heavily understudied.

While pre-training on ImageNet has become a common practice in medical image classification, there have been growing concerns within the medical imaging community regarding its suitability for medical imaging tasks. Medical images differ from natural images in several ways, including local texture variations as an indication of pathology rather than a clear global subject present in natural images (Raghu et al., 2019). Additionally, medical datasets are smaller in size, have fewer classes, have higher resolution compared to ImageNet, and go beyond 2D. These differences between natural and medical image datasets have led to the argument that ImageNet may not be the optimal solution for pre-training in medical imaging due to the well-known performance degradation effect caused by domain shift. This has led to increased efforts to explore alternative solutions for pre-training, such as using existing medical datasets (El-Nouby et al., 2021), their alterations (Kataoka et al., 2020; Asano et al., 2020), and creating new medical image datasets specifically designed for pre-training, such as RadImageNet (Mei et al., 2022).

Recent studies have challenged the conventional wisdom that the source dataset used for pre-training must be closely related to the target task in order to achieve good performance. Evidence has emerged suggesting that the source dataset may not have a significant impact on the performance of the target task and we can pre-train on any real large-scale diverse data (Gavrikov & Keuper, 2022a;b). Further, more evidence suggests that ImageNet leads to the best transfer performance in terms of accuracy, as ImageNet not only boosts the performance (Ke et al., 2021) but also is a better source than medical image datasets (Brandt

et al., 2021). This is likely due to the focus on texture in ImageNet models (Geirhos et al., 2022), which has been hypothesized to be an important cue for medical image classification.

While RadImageNet has demonstrated ImageNet-level accuracy (Mei et al., 2022) on radiology image classification, it remains uncertain whether it leads to improved representations when applied to medical target datasets. Furthermore, it is essential to understand the broader implications of source datasets beyond their effect on target task performance, in order to enable practitioners to make more informed decisions when selecting a source dataset.

In light of the ongoing debate on the choice of source dataset for medical pre-training, we set out to investigate this with a series of systematic experiments on the difference of representations learned from natural (ImageNet) and medical (RadImageNet) source datasets on a range of (seven) medical targets. Our main contributions are:

- We extend the work presented in Mei et al. (2022) by doing a reproducibility study of four of their seven experiments (derived from three small medical targets: breast, thyroid, and knee datasets) and adding four additional medical imaging target datasets. Contrary to the findings in (Mei et al., 2022), we observe that in most cases, models pre-trained on ImageNet tend to perform better than those trained on RadImageNet. However, it is important to note that this discrepancy does not necessarily indicate the superiority of one source dataset over the other. Rather, it emphasizes the sensitivity of transfer performance to the choice of model architecture and hyperparameters.
- We investigate the learned intermediate representations of the models pre-trained on ImageNet and RadImageNet using Canonical Correlation Analysis (CCA) (Raghu et al., 2017; 2019). Our results reveal that the networks converge to distinct intermediate representations, and that these representations are even more dissimilar after fine-tuning.
- We show that although models pre-trained on ImageNet and RadImageNet learn different intermediate representations, they still produce similar predictions after fine-tuning.
- Our findings demonstrate limited feature reuse even in the first two layers of a convolutional neural network (CNN), contrary to findings by Raghu et al. (2019). We discover a negative relationship between model similarity (before and after fine-tuning) and the improvement in performance across all layers.

Our code and experiments are available on Github<sup>1</sup>.

## 2 Related work

## 2.1 Pre-training on different data

Transfer performance degradation due to distribution shift is a known problem. This issue is particularly relevant in scenarios where the availability of large-scale in-domain supervised data for pre-training is limited. In light of this, a line of research has emerged that challenges the common practice of pre-training on ImageNet and instead explores pre-training on different in-domain source datasets:

Training on target data. El-Nouby et al. (2021) have shown the potential of using denoising autoencoders for pre-training on target data in a self-supervised manner. Although this approach has yielded promising results, the experiments were conducted using natural image target datasets. Even the smallest target dataset consisted of 8,000 images, making it unclear how well this approach would translate to the domain of medical imaging where the availability of large-scale target data is often limited.

Synthetic data has been shown to be a viable alternative to real-world data for pre-training, particularly in domains where labeled data is scarce. Kataoka et al. (2020) have explored pre-training on gray-scale automatically generated fractals with labels and showed that this approach can generate unlimited amounts

<sup>1</sup>https://anonymous.4open.science/r/86380762B567/

of synthetic labeled images, although it does not surpass the performance of pre-training on ImageNet in all cases.

**Data augmentation** is often used to increase dataset size, but Asano et al. (2020) showed that it can also be used to scale a single image for self-supervised pre-training, though this approach still falls short of using real diverse data. Even with millions of unlabeled images, it cannot fully bridge the gap between fully-supervised and self-supervised pre-training for deeper layers of a CNN.

Although aforementioned work in general computer vision has demonstrated the potential of synthetic and augmented data, the importance of large-scale labeled source datasets remains strong, particularly ImageNet in medical imaging (Brandt et al., 2021; Wen et al., 2021) despite its out-of-domain nature for medical targets. Recently, Mei et al. (2022) have demonstrated that RadImageNet, a large-scale dataset of radiology images similar in size to ImageNet, outperforms ImageNet on radiology target datasets. To further these findings, we investigate the potential of pre-training on the RadImageNet on a range of modalities that were not included in Mei et al. (2022) experiments, such as X-rays, dermoscopic images, and histopathological scans.

#### 2.2 Effects of transfer learning

Transfer learning is a useful method for studying representations and generalization in deep neural networks. Yosinski et al. (2014) defined the generality of features learned by a convolutional layer based on their transferability between tasks. They analyzed representations learned in ImageNet models and found that the early layers form general features resembling Gabor filters and color blobs, while deeper layers become more task-specific. More recently, Raghu et al. (2019) studied feature reuse in medical imaging using transfer learning and found that this reuse is limited to the lowest two convolutional layers. Besides feature reuse, they demonstrated that the scaling of pre-trained weights can result in significant improvement in convergence speed.

Instead of investigating the transferability of weights at different layers, Gavrikov & Keuper (2022b) investigated the distributions of convolution filters learned by computer vision models and found that they only exhibit minor variations across various tasks, image domains, and datasets. They noted that models based on the same architecture tend to learn similar distributions when compared to each other, but differ significantly when compared to other architectures. The authors also discovered that medical imaging models do not learn fundamentally different filter distributions compared to models for other image domains. Based on these findings, they concluded that medical imaging models can be pre-trained with diverse image data from any domain.

Our work extends previous studies on the effects of pre-training by characterizing the representations learned from ImageNet and RadImageNet and investigating the implications of the source dataset on the learned representations. Our results further challenge the notion of the generality of early convolutional layers, showing that feature reuse and target task performance are negatively related in all convolutional layers. Our results provide additional evidence that the source domain may not be of high importance for pre-training medical imaging models, as we observe that even though ImageNet and RadImageNet pre-trained models converge to distinct hidden representations, their predictions are still similar.

#### 3 Method

We outline our overall method in Figure 1. We fine-tune publicly available pre-trained ImageNet and RadImageNet weights on medical target datasets and quantify the model similarity by comparing the network activations over a sample of images from the target datasets using two similarity measures, Canonical Correlation Analysis (CCA, Section 3.1) and prediction similarity (Section 3.2).

## 3.1 Canonical Correlation Analysis

CCA (Hotelling, 1936) which is a statistical method used to analyze the relationship between two sets of variables.

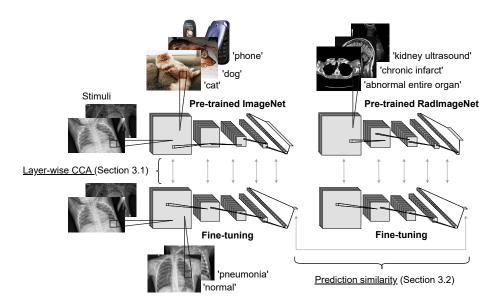


Figure 1: Overview of the experimental setup. Publicly available pre-trained ImageNet and RadImageNet weights are fine-tuned on medical targets. Model similarity is evaluated by comparing network activations over sampled stimuli images from target datasets using both CCA (described in Section 3.1) and prediction similarity (Section 3.2).

Let **X** be a dataset  $\mathbf{x}_1, \mathbf{x}_2, ..., \mathbf{x}_n$  of n data points, all consisting of p variables, and  $\mathbf{Y}$  – a dataset of n data points  $\mathbf{y}_1, \mathbf{y}_2, ..., \mathbf{y}_n$  and q variables. CCA seeks to find the transformation matrices **A** and **B** that linearly combine the initial variables p and q in the datasets **X** and **Y** into  $\min(p, q)$  canonical variables  $\mathbf{X}\mathbf{a}^i$  and  $\mathbf{Y}\mathbf{b}^i$  such that the correlation between these canonical variables is maximized:

$$\begin{aligned} \mathbf{a}^i, \mathbf{b}^i &= \operatorname*{argmax}_{\mathbf{a}^i, \mathbf{b}^i} \operatorname{corr}(\mathbf{X} \mathbf{a}^i, \mathbf{Y} \mathbf{b}^i) \\ \operatorname{subject to} \ \forall_{j < i} \ \mathbf{X} \mathbf{a}^i \perp \mathbf{X} \mathbf{a}^j \\ \forall_{j < i} \ \mathbf{Y} \mathbf{b}^i \perp \mathbf{Y} \mathbf{b}^j \end{aligned}$$

The restrictions ensure that the canonical variables are orthogonal. This can be solved by defining substitutions  $\bar{\mathbf{A}} = \mathbf{\Sigma}_{V}^{1/2} \mathbf{A}$  and  $\bar{\mathbf{B}} = \mathbf{\Sigma}_{V}^{1/2} \mathbf{B}$  obtaining:

$$\begin{split} \bar{\mathbf{A}}, \bar{\mathbf{B}} &= \operatorname*{argmax}_{\bar{\mathbf{A}}, \bar{\mathbf{B}}} \operatorname{tr}(\bar{\mathbf{A}}^{\top} \boldsymbol{\Sigma}_{X}^{-1/2} \boldsymbol{\Sigma}_{XY} \boldsymbol{\Sigma}_{Y}^{-1/2} \bar{\mathbf{B}}) \\ & \text{subject to } \bar{\mathbf{A}}^{\top} \bar{\mathbf{A}} = \mathbf{I} \\ \bar{\mathbf{B}}^{\top} \bar{\mathbf{B}} &= \mathbf{I} \end{split}$$

Because of the orthogonality constraints, the solution is found by decomposing  $\Sigma_X^{-1/2}\Sigma_{XY}\Sigma_Y^{-1/2}$  into left and right singular vectors using singular value decomposition.

Layer-wise model similarity. Raghu et al. (2017) proposed the use of CCA for comparing representations learned by neural networks. CCA's invariance to linear combinations makes it suitable for comparing the representations learned by different models as the layer weights in neural networks are combined before being passed on (Morcos et al., 2018).

Raghu et al. (2019) used CCA to examine representations in medical imaging models. In order to maintain consistency with Raghu et al. (2019) results, we adopt the same approach of applying CCA to CNNs and use an open source CCA implementation by Raghu et al. (2017) available on GitHub<sup>2</sup>.

In our case, X and Y are same-level layer activation vectors over n stimuli images sampled from a target dataset, in two models with different initializations. We extract these intermediate representations and use them as input to CCA to project the representations onto a common space, where the correlation between the projections is maximized. This common space can be thought of as a shared representation that captures the common patterns of activity across the compared networks. Then, layer-wise similarity at layer L is the average of the correlations between the canonical variables:

$$\rho_L = \sum_{i=1}^p \operatorname{corr}(\mathbf{X}\mathbf{a}^i, \mathbf{Y}\mathbf{b}^i) \tag{1}$$

Intermediate representations extracted from CNNs are of shape  $(n, h_L, w_L, p_L)$ , where  $h_L$ ,  $w_L$  are the layer spatial dimensions and  $p_L$  is the number of channels in the layer. These representations are reshaped into **X** and **Y** matrices of shape  $(n \times h_L \times w_L, p_L)$ . As CCA is sensitive to the shape of the input matrices and the shapes vary across the layers within a network, we sample n,  $p_L$  and then calculate layer similarity  $\rho_L$ . This is repeated five times and the final layer similarity is obtained by averaging layer similarities  $\rho_L$ .

#### 3.2 Prediction similarity

We calculate prediction similarity as described in Mania et al. (2019). A model mistake is defined as  $q_f(x,y) = \mathbf{1}_{f(x)\neq y}$ , where x is an image from the test set, y is its label and f is a network fine-tuned on the target training set. Then the prediction similarity of two networks  $f_{\text{ImageNet}}$  and  $f_{\text{RadImageNet}}$ , fine-tuned on the same target dataset, is:

$$\mathbb{P}(q_{f_{\text{ImageNet}}}(x, y) = q_{f_{\text{RadImageNet}}}(x, y)) \tag{2}$$

Therefore, the prediction similarity is the probability that two networks will make the same errors. To gauge the prediction similarities between ImageNet and RadImageNet models, we compare them to the prediction similarity of two classifiers with the same accuracy as ImageNet and RadImageNet models but otherwise random predictions. If the mistakes made by two models with accuracy  $a_1$  and  $a_2$  are independent, the similarity of their predictions is equal to  $a_1a_2 + (1 - a_1)(1 - a_2)$ .

## 4 Experimental setup

#### 4.1 Datasets

**Source.** We use publicly available pre-trained ImageNet (Deng et al., 2009) and RadImageNet (Mei et al., 2022) weights as source tasks in our experiments.

**Target.** We investigate transferability to several medical target datasets. In particular, to five radiology RadImageNet in-domain datasets, and two out-of-domain datasets in the fields of dermatology and microscopy. A representative image from each dataset can be seen in Figure 2.

- 1) Chest. Chest X-rays (Kermany et al., 2018) dataset contains chest X-ray images from pediatric patients aged one five years old, labeled by expert physicians with binary labels of 'normal' or 'pneumonia'. The dataset has 5,856 images, with 1,583 labeled as 'normal' and 4,273 labeled as 'pneumonia'. The image size varies, with dimensions ranging from  $72 \times 72$  to  $2,916 \times 2,583$  pixels.
- 2) Breast. Breast ultrasound (Al-Dhabyani et al., 2020) dataset is collected for the detection of breast cancer. The images have a range of sizes, from  $190 \times 335$  to  $1,048 \times 578$  pixels. The dataset is divided into

<sup>&</sup>lt;sup>2</sup>https://github.com/google/svcca

Figure 2: Example images of (a) chest, (b) breast, (c) thyroid, (d) mammograms, (e) knee, (f) pcam-small, and (g) ISIC datasets.

three classes: normal, benign, and malignant images. However, following Mei et al. (2022), we use a binary classification of 'benign' and 'malignant' for our analysis.

- 3) Thyroid. The Digital Database of Thyroid Ultrasound Images (DDTI) (Pedraza et al., 2015) contains 480 images of size 569 × 360 pixels, extracted from thyroid ultrasound videos. The images have been annotated by radiologists into five categories. Following Mei et al. (2022)'s study, these categories were transformed into binary labels: 'normal' for categories (1) normal thyroid, (2) benign and (3) no suspicious ultrasound (US) feature, and 'malignant' for categories (4a) one suspicious US feature, (4b) two suspicious US features, (4c) three or four suspicious US features and (5) five suspicious features.
- 4) Mammograms. Curated Breast Imaging Subset of Digital Database for Screening Mammography (CBIS-DDSM) (Sawyer-Lee et al., 2016; Lee et al., 2017; Clark et al., 2013) is a dataset that targets breast cancer detection. It contains scanned film mammograms with pathologically confirmed labels: 'benign' (2,111 images) or 'malignant' (1,457 images) with image sizes ranging from 1,846 × 4,006 to 5,431 × 6,871 pixels.
- 5) Knee. MRNet (Bien et al., 2018) is a collection of 3D knee MRI scans. The labels for the dataset were obtained through manual extraction from clinical reports. Following Mei et al. (2022)'s study, we use extracted 2D sagittal views (1 to 3 samples per scan) amounting to a total of 4,235 'normal', 569 'ACL' (anterior cruciate ligament), and 418 'meniscal tear' images, all of size 256 × 256 pixels.
- 6) PCam-small. PatchCamelyon (Veeling et al., 2018) is a metastatic tissue classification dataset consisting of 237,680 colored patches extracted from histopathological scans of lymph node sections. The images are labeled as 'positive' or 'negative' based on the presence of metastatic tissue. To simulate a realistic target dataset size, a random subset of 10,000 images was created, with 5,026 positive and 4,974 negative samples.
- 7) ISIC. ISIC 2018 Challenge Task 3: Lesion Diagnosis (Codella et al., 2019; Tschandl et al., 2018) a dermoscopic lesion image dataset released for the task of skin lesion classification. The dataset comprises images of  $600 \times 450$  pixels, which are split into seven disease categories. The dataset is unbalanced, with the class 'melanocytic nevus' having the most samples at 6,705, and the class 'dermatofibroma' having the least number of samples at 115.

Due to memory constraints, we reduced the original image sizes for most of the target datasets using interpolation without image cropping. Table 1 provides details of the image sizes and number of images used for fine-tuning on each target dataset. As we used publicly available pre-trained weights images were preprocessed to align with the pre-trained weights. As per the approach in Mei et al. (2022), we normalized the images with respect to the ImageNet dataset.

## 4.2 Fine-tuning

We select ResNet50 (He et al., 2016) as the standard model architecture for our experiments. This architecture is widely adopted in the field of medical imaging and has been demonstrated to be a strong performer in various image classification tasks.

We fine-tuned pre-trained networks using an average pooling layer, a dropout layer with a probability of 0.5, and the Adam optimizer with an initial learning rate of 1e-05. Due to memory limitations, the batch size was set to 32 for the datasets with the image size  $224 \times 224$ , and to 128 for the rest. The models were trained for a maximum of 200 epochs, with early stopping after 3 epochs of no decrease in validation loss. This was done to prevent overfitting and ensure that the models generalize well to unseen data.

Table 1: Target datasets with number of images, number of classes and image size used to fine-tune the pre-trained ImageNet and RadImageNet weights.

| Dataset    | Size      | Classes | Image size       |
|------------|-----------|---------|------------------|
| Chest      | 5,856     | 2       | $112 \times 112$ |
| Breast     | 780       | 2       | $224 \times 224$ |
| Thyroid    | 480       | 2       | $224 \times 224$ |
| Mammograms | $3,\!568$ | 2       | $224 \times 224$ |
| PCam-small | 10,000    | 2       | $96 \times 96$   |
| ISIC       | 10,015    | 7       | $112 \times 112$ |

Table 2: Mean AUC  $\pm$  std (both  $\times 100$ ) after fine-tuning on target datasets. Underlined is the highest mean AUC per dataset.

|                | ImageNet                  |                | RadIamgeNet                |                 | Random init     |
|----------------|---------------------------|----------------|----------------------------|-----------------|-----------------|
| Target dataset | No Freeze                 | Freeze         | No Freeze                  | Freeze          | No Freeze       |
| Thyroid        | $52.3 \pm 8.9$            | $58.5 \pm 5.1$ | $50.2 \pm 4.1$             | $49.4 \pm 5.9$  | $52.9 \pm 5.1$  |
| Breast         | $90.5 \pm 3.8$            | $89.8 \pm 3.0$ | $88.2 \pm 4.2$             | $72.4 \pm 22.1$ | $51.2 \pm 10.0$ |
| Chest          | $98.7 \pm 0.4$            | $98.6 \pm 0.6$ | $98.8 \pm 0.2$             | $98.7 \pm 0.3$  | $82.5 \pm 1.0$  |
| Mammograms     | $63.4 \pm 4.3$            | $68.8 \pm 2.0$ | $\overline{62.0 \pm 12.2}$ | $66.2 \pm 6.3$  | $49.6 \pm 3.7$  |
| Knee           | $91.5 \pm 1.1$            | $89.0 \pm 3.1$ | $89.3 \pm 6.3$             | $63.8 \pm 5.7$  | $68.3 \pm 11.0$ |
| ISIC           | $\overline{94.2 \pm 1.3}$ | $93.1 \pm 2.3$ | $90.8 \pm 0.8$             | $90.6 \pm 0.7$  | $84.0 \pm 2.3$  |
| Pcam-small     | $93.8 \pm 1.1$            | $93.2 \pm 2.5$ | $87.1 \pm 2.3$             | $86.0 \pm 1.9$  | $73.4 \pm 8.2$  |

In addition to full fine-tuning, we used a freezing strategy where we froze all the pre-trained weights to train the classification layer first and then fine-tuned the whole network with the same hyperparameters as above.

Models were implemented using Keras (Chollet et al., 2015) library and fine-tuned on 3 NVIDIA GeForce RTX 2070 GPU cards.

## 4.3 Evaluation

We fine-tune the pre-trained networks on each target dataset using five-fold cross-validation approach. To ensure patient-independent validation where patient information is available (chest, thyroid, mammograms, knee), the target data is split such that the same patient is only present in either the training or validation split. We evaluate fine-tuned network performance using AUC (area under the receiver operating characteristic curve), while a one-against-all AUC, weighted by class priors, is used for multi-class problems. Model similarity is evaluated using CCA and prediction similarity as described in Section 3.

## 5 Results

We carried out a series of experiments to evaluate the effect of pre-training on ImageNet and RadImageNet on model accuracy and learned representations after fine-tuning on medical targets. In the following section, we present the results of our experiments and provide a thorough analysis of the findings. The results offer new insights into the effects of transfer learning and provide a foundation for future research in this field.

#### 5.1 ImageNet tends to outperform RadImageNet

We show the AUC performances in Table 2. Overall, the unfrozen ImageNet leads to the highest AUCs in four out of the seven datasets, followed by the frozen ImageNet in two datasets. Only Chest reaches the highest performance with frozen RadImageNet weight, though we note that all performances were close for

this dataset. In general, we found that the performance of ImageNet and RadImageNet was comparable for radiology targets. However, when evaluating on the ISIC and pcam-small datasets, ImageNet produced considerably better results. This discrepancy is likely due to the fact that RadImageNet consists of grayscale images, which might limit its performance in dermatology and microscopy images despite its medical nature.

Compared to Mei et al. (2022), we get similar ImageNet AUC for knee and breast, but much lower RadImageNet AUC. We note that Mei et al. (2022) evaluated RadImageNet performance using a subset of 349 images from the thyroid dataset, compared to 480 images available and split the classification of ACL and meniscal tear into two separate tasks for the knee dataset. Furthermore, the results reported by Mei et al. (2022) were aggregated over four different model architectures and multiple hyperparameter (freezing and learning rate) combinations, making it difficult to directly compare our ResNet50 results. This highlights the sensitivity of transfer performance to the choice of model architecture and hyperparameters.

## 5.2 Layer-wise representations become more different after fine-tuning

In Figure 3 we show layer-wise ImageNet and RadImageNet CCA similarity to themselves after fine-tuning, ImageNet<sup>FT</sup> and RadImageNet<sup>FT</sup> respectively (Figure 3a), as well as layer-wise ImageNet and RadImageNet CCA similarity before and after fine-tuning (Figure 3b). ImageNet weights change less during fine-tuning (orange line on the left). The two networks converge to distinct solutions after fine-tuning (green line on the right), even more distinct than before fine-tuning, and their similarity is significantly lower when compared to the similarity of two random initializations. Here we only provide results on knee, however we observed similar patterns for the other target datasets (Appendix A).

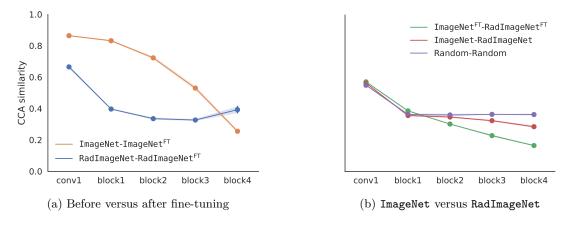


Figure 3: Layer-wise CCA similarity (Equation 1) of (a) a network to itself before and after fine-tuning on knee and (b) ImageNet to RadImageNet. ImageNet-ImageNet<sup>FT</sup> similarity (orange line) is higher (the weights change less after fine-tuning) than RadImageNet-RadImageNet<sup>FT</sup> similarity (blue line). ImageNet<sup>FT</sup> and RadImageNet<sup>FT</sup> are highly dissimilar (green line), even more dissimilar than before fine-tuning (red line). They are also more dissimilar than two randomly initialized networks (purple line). Error bars present mean  $\pm$  std over five-fold cross-validation.

For experiments where pre-trained weights were initially frozen and fine-tuned after training the classification layer, we observed essentially similar trends of weight similarity as shown in Figure 3, but with higher variability between the folds of the cross-validation.

The results of the layer-wise CCA similarity analysis reveal that ImageNet and RadImageNet converge to distinct solutions after fine-tuning on the same target dataset, to the extent that they become even more dissimilar than before fine-tuning. This outcome contradicts our expectation that the representations of the two networks would become more similar after training on the same target dataset. This discrepancy may be due to memorization of the target dataset by one or both of the networks, as suggested by the findings of Morcos et al. (2018). They found that networks trained to classify randomized labels, hence memorizing the data, tend to converge to more distinct solutions compared to networks that generalize to unseen data.

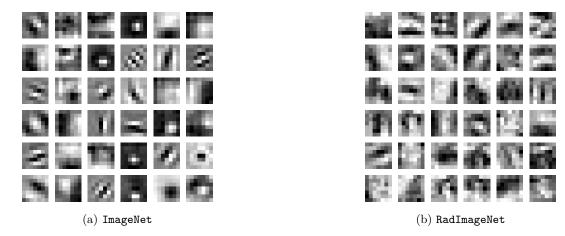


Figure 4: First 36 conv1 filters of ResNet50 pre-trained on (a) ImageNet and (b) RadImageNet. Observe that the filters in ImageNet have a more pronounced resemblance to Gabor filters.

The stability of early layer weights during fine-tuning is often attributed to their capture of general features, such as edge detectors (Raghu et al., 2019; Yosinski et al., 2014), which are necessary regardless of the target domain and task. Our results are consistent with this hypothesis, in that we observe that early layers change less compared to later layers (Figure 3a). However, we also found an interesting and contradictory result, in that the early layers of ImageNet and RadImageNet are as similar as two randomly initialized layers. This observation is inconsistent with the intuition that early layers capture general features, as we would expect them to be more similar than randomly initialized layers if they indeed capture general features.

When we examine the first convolutional layer filters pre-trained on ImageNet and RadImageNet (Figure 4), we observe that the filters in ImageNet more closely resemble Gabor filters, while those in RadImageNet are more fuzzy. This is expected, as natural images often contain regular structures, such as 90 degree angles and edges, that are typically less prominent in some of radiology images, resulting in less distinct edges in the filters learned from radiology images. Interestingly, these different first layer features in both ImageNet and RadImageNet, without changing significantly during fine-tuning (Figure 3), lead to comparable performance in most cases (Table 2).

#### 5.3 Networks make similar mistakes after fine-tuning

In order to further understand the similarity between ImageNet and RadImageNet pre-trained networks, we compared their predictions after fine-tuning on medical targets, as shown in Figure 5. Despite the networks converging to different hidden representations, as evidenced in Figure 3, their predictions were found to be more similar than expected for independent predictions. The fact that both ImageNet and RadImageNet were fine-tuned on the same target dataset may explain this observation. However, it also suggests the possibility that the networks are learning similar misleading cues in the data, resulting in similar misclassifications.

#### 5.4 Higher weight similarity associated with less AUC improvement

Our results challenge the commonly held belief that the benefits of transfer learning in deep neural networks come from feature reuse in the early layers. Figure 6 shows that the gain in AUC due to pre-training is negatively related to the weight CCA similarity between the pre-trained and fine-tuned networks. This means that models that reused the pre-trained weights without making changes during fine-tuning performed worse, as opposed to those that adapted the weights. This negative relationship was observed across all layers and was particularly pronounced in the RadImageNet pre-trained networks.

Our findings align with recent results in the Natural Language Processing field which demonstrate that the benefits of pre-training are not related to knowledge transfer (Krishna et al., 2021). Additionally, our results complement the findings of Raghu et al. (2019) who showed that there are feature-independent benefits of

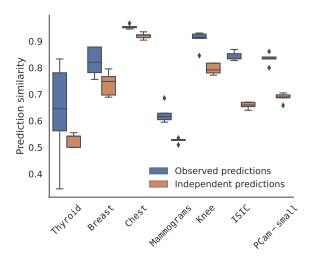


Figure 5: Prediction similarity (Equation 2) between ImageNet<sup>FT</sup> and RadImageNet<sup>FT</sup> (blue box plot) compared to prediction similarity of two networks that would make independent mistakes (orange box plot). ImageNet<sup>FT</sup> and RadImageNet<sup>FT</sup> predictions are more correlated than expected for independent predictions on average across all target datasets, with notable variation observed for thyroid dataset.

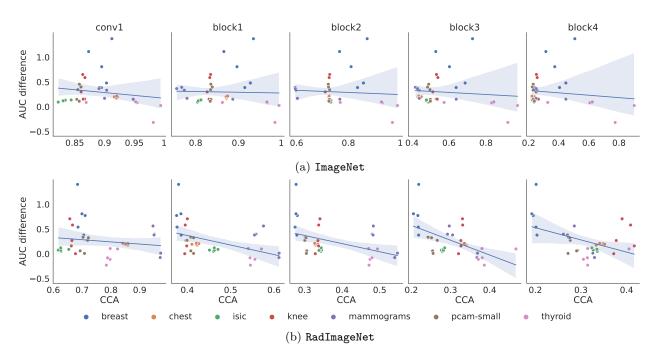


Figure 6: AUC pre-training gains over random initialization for seven target datasets vs CCA similarity before and after fine-tuning on those targets, for (a) ImageNet and (b) RadImageNet, with 95% confidence interval. Higher CCA similarity after fine-tuning is associated with lower AUC gains, observed across all layers with a more pronounced effect on RadImageNet. Note that the scaling on the x-axes are different in each plot for visibility, and for RadImageNet the CCA similarity is lower overall.

using pre-trained weights, such as better scaling compared to random initialization. These results highlight the complex nature of transfer learning and the need for further investigation into the underlying mechanisms that drive its performance benefits.

## 6 Discussion

In our experiments ImageNet generally outperformed RadImageNet on medical target datasets, in contrast to the earlier results reported in Mei et al. (2022). However, this highlights the sensitivity of source dataset transfer performance to the model architecture and hyperparameters, rather than the inherent superiority of one source dataset over the other. While our study did not comprehensively analyze this sensitivity, interested readers can refer to related studies, such as Raghu et al. (2019) and Wen et al. (2021), which offer valuable insights into the impact of model architecture and hyperparameters on transfer performance.

Contrary to our preconceptions about transfer learning, our analysis with CCA found that the models converged to distinct intermediate representations and that these representations are even more dissimilar after fine-tuning on the same target dataset. Despite distinct intermediate representations, model predictions on an instance level exhibit a significant degree of similarity. This extends Mania et al. (2019) findings which showed that ImageNet models are similar in their predictions even with different architectures. We extend upon the findings of Raghu et al. (2019), who demonstrated limited feature reuse in the lowest two layers of a CNN. We discover that even the features in the first two layers are not reused to the same extent as previously believed.

While our results are similar across seven different target datasets, these tasks are not fully representative of the medical imaging field as a whole. Researchers have hypothesized that for 3D target tasks such as CT or MRI, 3D pre-training might be a better alternative to 2D pre-training (Wong et al., 2018; Brandt et al., 2021), and studies have shown that incorporating information from the third dimension might be beneficial for performance (Yang et al., 2021; Taleb et al., 2020). However, RadImageNet only has 2D images, even though some of the original images are 3D. From our results, we are unable to conclude that ImageNet weights are generally more suitable for medical target tasks.

Another limitation is in only using AUC as the metric for evaluating performance. Although this is a commonly used metric for classification tasks in medical imaging, there might be nuances across applications where it could be important to consider alternative metrics, such as calibration (Reinke et al., 2021).

Our results suggest that the implications of using ImageNet in medical image classification go beyond performance alone. In particular, there might be an issue with the memorization of spurious patterns in the data. This can potentially have consequences with respect to algorithmic bias and fairness. For example, see Gichoya et al. (2022) where ImageNet pre-trained networks memorize patient race. Memorization also makes a network more vulnerable to adversarial attacks (Bortsova et al., 2021). It would be interesting to further explore these properties in future research.

While RadImageNet's comparable size to ImageNet allowed for a unique opportunity to compare natural and medical source datasets, it is important to note that the two datasets have significant differences beyond their domains. For instance, RadImageNet has differences in color, number of classes, and diversity in data due to its limited number of patients. To further explore the impact of these differences in greater detail, future research could consider including Ecoset (Mehrer et al., 2021), a natural image dataset with 565 basic-level categories selected to better reflect the human perceptual and cognitive experience.

#### 7 Conclusions

Transfer learning is a key strategy to leverage knowledge from the models pre-trained on large-scale datasets to deal with the challenge of small medical datasets. In this study, we investigated the transferability of two different domain sources (natural: ImageNet and medical: RadImageNet) to several target medical image classification tasks. Our results show that pre-training on ImageNet outperformed RadImageNet in most cases. Furthermore, we delved deeper into the learned representations after fine-tuning by using CCA and comparing the similarity of predictions. Although the models converged to distinct representations, we found they made similar predictions. Finally, we observed limited reuse of features, even in the early convolutional layers, after fine-tuning.

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# A Layer-wise CCA similarity

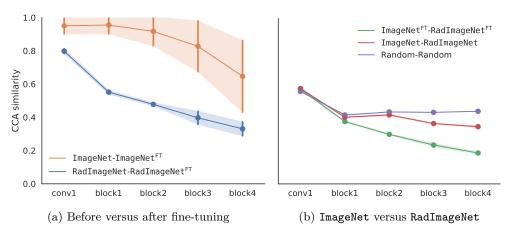


Figure 7: Layer-wise CCA similarity of networks fine-tuned on thyroid.

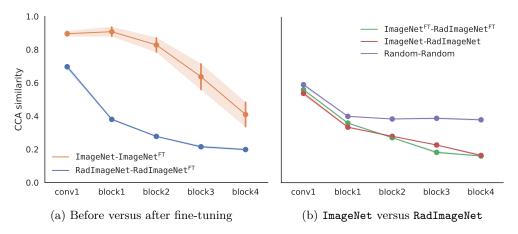


Figure 8: Layer-wise CCA similarity of networks fine-tuned on breast.

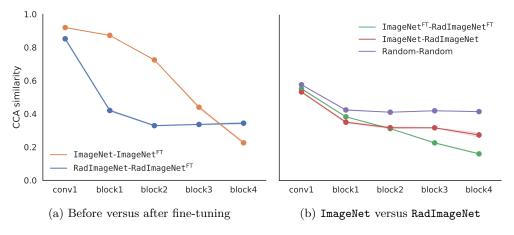


Figure 9: Layer-wise CCA similarity of networks fine-tuned on chest.

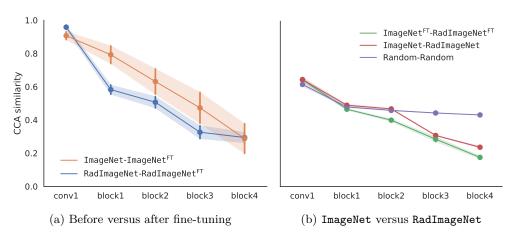


Figure 10: Layer-wise CCA similarity of networks fine-tuned on mammograms.

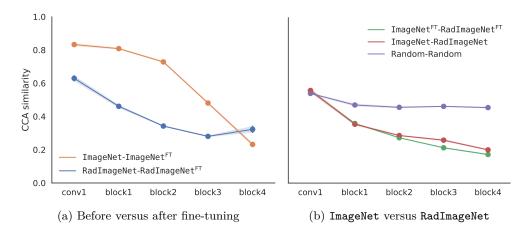


Figure 11: Layer-wise CCA similarity of networks fine-tuned on isic.

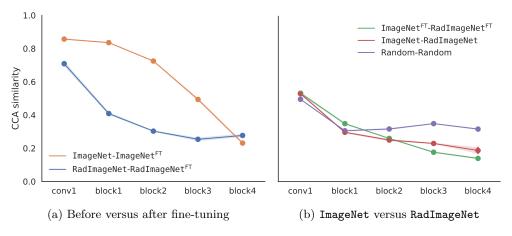


Figure 12: Layer-wise CCA similarity of networks fine-tuned on pcam-small.