#### **ORIGINAL RESEARCH**



# **A Novel Artifcial Intelligence System for the Prediction of Interstitial Lung Diseases**

**Nidhin Raju1 · D. Peter Augustine1 · J. Chandra<sup>1</sup>**

Received: 17 March 2023 / Accepted: 23 November 2023 © The Author(s), under exclusive licence to Springer Nature Singapore Pte Ltd 2024

#### **Abstract**

Interstitial lung disease (ILD) encompasses a spectrum of more than 200 fatal lung disorders afecting the interstitium, contributing to substantial mortality rates. The intricate process of diagnosing ILDs is compounded by their diverse symptomatology and resemblance to other pulmonary conditions. High-resolution computed tomography (HRCT) assumes the role of the primary diagnostic tool for ILD, playing a pivotal role in the medical landscape. In response, this study introduces a computational framework powered by artifcial intelligence (AI) to support medical professionals in the identifcation and classifcation of ILD from HRCT images. Our dataset comprises 3045 HRCT images sourced from distinct patient cases. The proposed framework presents a novel approach to predicting ILD categories using a two-tier ensemble strategy that integrates outcomes from convolutional neural networks (CNNs), transfer learning, and machine learning (ML) models. This approach outperforms existing methods when evaluated on previously unseen data. Initially, ML models, including Logistic Regression, BayesNet, Stochastic Gradient Descent (SGD), RandomForest, and J48, are deployed to detect ILD based on statistical measures derived from HRCT images. Notably, the J48 model achieves a notable accuracy of 93.08%, with the diagnostic signifcance of diagonal-wise standard deviation emphasized through feature analysis. Further refnement is achieved through the application of Marker-controlled Watershed Transformation Segmentation and Morphological Masking techniques to HRCT images, elevating accuracy to 95.73% with the J48 model. The computational framework also embraces deep learning techniques, introducing three innovative CNN models that achieve test accuracies of 94.08%, 92.04%, and 93.72%. Additionally, we evaluate fve full-training and transfer learning models (InceptionV3, VGG16, MobileNetV2, VGG19, and ResNet50), with the InceptionV3 model achieving peak accuracy at 78.41% for full training and 92.48% for transfer learning. In the concluding phase, a soft-voting ensemble mechanism amplifes training outcomes, yielding ensemble test accuracies of 76.56% for full-training models and 92.81% for transfer learning models. Notably, the ensemble comprising the three newly introduced CNN models attains the pinnacle of test accuracy at 97.42%. This research is poised to drive advancements in ILD diagnosis, presenting a resilient computational framework that enhances accuracy and ultimately betters patient outcomes within the medical domain.

**Keywords** ILD · Deep learning · Transfer learning · Multi-label classifcation · HRCT

# **Introduction**

A diverse group of acute and chronic bilateral parenchymal pulmonary disorders known as ILD contain some clinical characteristics, but their severity and prognosis vary signifcantly [\[1\]](#page-17-0). Idiopathic pulmonary fbrosis is one of the most common and serious interstitial lung disorders. It is

 $\boxtimes$  Nidhin Raju nidhin.raju@res.christuniversity.in characterized by an increase in fbrosis, decreased lung function, and eventually death [\[2](#page-17-1)]. Patients with idiopathic pulmonary fbrosis typically die 5 years after being diagnosed [[3\]](#page-17-2). Due to the need to rule out a variety of ILD, connective tissue diseases, and workplace and environmental exposures, diagnosing idiopathic pulmonary fbrosis can be challenging. Patients who are suspected of having idiopathic pulmonary fbrosis frequently undergo high-resolution CT scans, but only when the typical pattern of ILD is clearly visible can the condition be reliably identifed. Between the onset of symptoms and a diagnosis of idiopathic pulmonary fbrosis, a period of 1–2 years typically passes [\[4](#page-17-3)]. However, in

<sup>&</sup>lt;sup>1</sup> Department of Computer Science, CHRIST (Deemed to be University), Bengaluru, India

order to clarify the histopathological characteristics of ILD, many patients require an invasive surgical lung biopsy. A conclusive diagnosis may still be difficult to make despite histological evaluation due to the fact that pathologists may difer regarding histopathological fndings and that a good diagnosis may depend on individual experience. The diagnosis is more accurate when pulmonologists, radiologists, and pathologists work together; sadly, not all patients have access to information of this caliber. Patients are required to visit recognized competence-based regional centers for such in-depth assessments, which take time [\[5](#page-17-4)].

Performance on categorization tasks is crucial to the quality of medical diagnostics. Because difuse lung disease (DLD) patterns can be seen in the lung at any cross-section, high-resolution computed tomography (HRCT) images are thought to be useful for diagnosing ILD-associated DLD patterns. Unfortunately, DLDs on HRCT images display a wide range of texture pattern meanings, making it difficult to diagnose the location of ILDs. For the proper treatment of IIPs, objective diagnosis and its quality improvement are sought, because the physician's ability to diagnose has an impact on the quality of the diagnosis. A computer-aided diagnostic (CAD) system for objective diagnosis is being developed in these decades to alleviate clinician strain. Using machine learning techniques, CAD systems are designed to provide a categorization function for a second opinion [\[6](#page-17-5), [7](#page-17-6)].

By employing a variety of distinct processing layers, a subset of machine learning known as "deep learning" enables computational models to learn representations of data with multiple levels of abstraction. The most obvious diference between deep learning and traditional machine learning is that deep learning can automatically extract fully automated features and generate models that are suitable for tasks from raw data on its own, whereas traditional machine learning requires feature extraction from humans [[8\]](#page-17-7). In recent years, deep learning techniques have made progress in a number of felds, including information technology and image and speech recognition. However, deep learning applications in the medical feld are still in their infancy. The analysis of medical images and the associated patient electronic medical data is a perfect application for deep learning [[9,](#page-17-8) [10\]](#page-17-9).

The signifcance of ILD detection through deep learning within the industrial context rests upon its capacity to significantly enhance diagnosis efficiency and accuracy. In light of the escalating integration of medical imaging in clinical settings, an escalating demand arises for automated systems that can collaboratively aid radiologists and healthcare professionals in comprehending these intricate images. By harnessing deep learning algorithms, the capability to swiftly and meticulously process vast volumes of medical imaging data emerges, thereby mitigating the burden on healthcare practitioners and augmenting patient outcomes.

Moreover, the utilization of deep learning for ILD detection extends its infuence into the realms of drug development and clinical trials. Through early identifcation of ILD patients, drug developers can orchestrate more streamlined and effective assessments of potential treatments. This expedites the drug development process, ultimately translating into superior treatment options for ILD patients. The AIdriven ILD detection paradigm engenders an amalgamation of advantages: elevating diagnostic precision and efficiency, diminishing the operational load on healthcare professionals, and catalyzing the pace of drug development and clinical trials. This confuence of benefts underscores the compelling potential of AI-based ILD detection to reshape the medical landscape.

Deep learning applied to the patient's medical imaging and data mining of the patient's electronic medical record (big data) should improve patient outcomes. Cloud-based applications allow the deep learning algorithm to train continuously on data sets that are not restricted to a single institution. In order to address unmet clinical requirements, numerous organizations are currently investigating applications based on deep learning. In the feld of chest imaging, the creation and implementation of computer-aided detection (CAD) systems for the detection of nodules on chest radiographs and chest computed tomography (CT) has received a lot of attention [\[11](#page-17-10)]. Despite the fact that numerous CAD systems are in use in clinical practice, their subpar performance (frequent false-positive and false-negative cases) has prevented widespread adoption [[12\]](#page-17-11). Deep learning methods have the potential to overcome the limitations of existing CAD systems, and a number of experiments have produced encouraging outcomes. In chest imaging, disease pattern identifcation, diagnosis, and survival prediction have all been carried out with success using deep learning. There are still some reservations regarding its therapeutic application potential [[13,](#page-17-12) [14\]](#page-17-13).

The remaining sections of the article are arranged as follows. Section "[Literature Review"](#page-1-0) covers a brief review of similar existing works. Section ["Methods](#page-3-0)" deals with the methods that have been used in this work. The results and analysis of the experiments are presented in section "[Data](#page-3-1)[sets"](#page-3-1). And a brief discussion and conclusion of this work are demonstrated in sections "[Data Preprocessing](#page-3-2)" and ["Results](#page-7-0) [and analysis](#page-7-0)", respectively.

# <span id="page-1-0"></span>**Literature Review**

Classical feature extraction methods like frst order grey level statistics, grey level co-occurrence matrices (GLCM), run-length matrices (RLM), and fractal analysis were provided by the earliest CAD systems for ILDs to represent 2D texture [[15](#page-17-14)]. The adaptive multiple feature method (AMFM) was created when these features were eventually combined [[16\]](#page-17-15). Certain attempts have recently been made to apply deep learning DL techniques, particularly CNNs, following their outstanding performance in large-scale color image categorization [[17\]](#page-17-16). The CNN modes can learn features simultaneously while training an Artifcial Neural Networks (ANN) classifer by minimizing classifcation error, in contrast to other feature learning techniques that produce unsupervised data representation models. The initial experiments on lung CT scans utilized shallow architectures, despite the fact that the term "deep learning" implies the utilization of numerous successive learning layers. In [\[18\]](#page-17-17), a modified restricted Boltzmann machine (RBM) that included particular CNN features was utilized for the purpose of feature extraction and categorization of lung tissue. Weight sharing was done among the hidden neurons that were tightly connected to label (output) neurons during the supervised training of the entire network using contrastive divergence and gradient descent. A CNN was constructed entirely from scratch by the authors of [[19](#page-17-18)], consisting of one convolutional layer and three dense layers. The shallow architecture of the network, on the other hand, prevents it from utilizing deep CNNs' descriptive capabilities. The pretrained deep CNN (AlexNet) from [\[17](#page-17-16)] was used in [\[20](#page-17-19)] to classify complete lung slices after being fne-tuned using lung CT data. AlexNet was designed to identify natural color photos with an input size of  $224 \times 224$  pixels; consequently, the authors were required to scale the images and artifcially create three channels by employing various Hounsfeld unit (HU) windows. In addition, there are concerns regarding knowledge transfer due to the signifcant diferences between ordinary color images and medical images, and identifying complete slices may only provide a very rough estimation of the condition.

Anthimopoulos et al. [[21\]](#page-17-20) built and trained one of the frst CNNs to categories the most prevalent ILD patterns, reaching a classifcation performance of 85.5% and exhibiting the DL identifcation potential for lung tissue idiosyncrasy. An experienced radiology team annotated 120 HRCTs by eliminating ambiguous lung areas and the bronco-vascular tree, which were then utilized to train and test the CNN. Christodoulidis et al. [\[22](#page-17-21)] developed a CNN architecture that can extract the textural variability of ILD patterns. Using transfer learning from multiple diferent non-medical source databases, they only achieved a 2% increase in the CNN performance. One of the downsides of this study was that they used CT scans instead of HRCT scans. Kim et al. [[23\]](#page-18-0) related shallow learning (SL) to DL for pattern classifcation. The authors made an effort to use relevant data from six texture benchmark databases for the current ILD pattern categorization task. According to these studies, they suggested to select a training source dataset that is comparable to the target domain and allow the network to learn some characteristics before fne-tuning. In their investigation, they used a CNN architecture with four convolutional layers and two fully connected layers. Simply increasing the number of convolutional layers increased accuracy from 81.27 to 95.12%. Gao et al. [[20](#page-17-19)] attempted a novel method for ILD pattern categorization, because they were aware of the diffculty of manually identifying region of interest (ROI) for automated pulmonary computer-aided diagnosis (CAD) systems. They demonstrated a more autonomous, grayscalebased holistic image identifcation system that was comparable to emphysema quantifcation [[24\]](#page-18-1).

A new method for creating an infnite number of arbitrary distinct ILD patterns from 2D HRCT images, which improved CNN's ability to classify patterns in lung tissue, was proposed by Bae et al. [\[25](#page-18-2)]. By providing a wide range of ILD patterns and stabilizing accuracy loss for the validation set, the program prevented overftting. The accuracy on a specifc area of interest or the entire lung was 89.5%, which was higher than the conventional CNN data augmentation rate of 82.1% and comparable to human capability. A CAD that could be easily implemented on standard computing software was created by Walsh et al. [[26\]](#page-18-3). The preprocessing of 1157 HRCT images produced up to 500 distinct 4-slice montages (concatenations) per CT scan. This resulted in a multiplied image dataset consisting of 420,096 distinct montages for the training algorithm and 40,490 for the validation set. In this study, the neural network architecture was the convolutional neural network Inception-ResNet V2.

The knowledge gathered from training samples is taken into account when calculating the weights for deep CNNs, which have recently demonstrated remarkable proficiency in a wide range of tasks. "Off-the-shelf" features that can be used for categorization may be obtained by reusing the network feature extraction capacity, according to some studies [[27,](#page-18-4) [28](#page-18-5)]. Through weight transferring methods that either freeze or precisely adjust the network's parameters, transfer learning has been achieved in other studies. In addition, the specifcity of various layering weights was investigated in [[29\]](#page-18-6). The network's frst and last layers, which are closest to the input and are typically generic, are more task-specifc. Consequently, the parameters of the frst layer of the transfer learning process can be fxed, the parameters of subsequent layers can be fne-tuned, and the network's non-transferred weights are typically initialized at random.

Transfer learning can make up for the lack of training data and improve the outcome of target tasks by utilizing knowledge gained in the source domain. Because there are typically few medical images that can be used for instruction, the advantages of transfer learning make it a popular method in the medical feld as well. Numerous researchers have attempted to employ well-known CNNs that have already been trained on ImageNet for medical image identifcation and classifcation tasks involving ultrasound, CT, and X-ray imaging [\[30](#page-18-7)[–32](#page-18-8)]. The domain dissimilarity variables still have an impact on the transferability of knowledge, despite the fact that these studies demonstrate the possibility of knowledge transfer from the domain of natural images to the domain of medical imaging. Several strategies to mitigate this efect have recently been the subject of research. Lu et al. [\[33\]](#page-18-9) introduced a restriction between the source and target classifer predictions describe a novel approach to transfer learning for acquiring useful knowledge from source data. In "multi-stage transfer learning," authors use an intermediary domain to connect the source and target domains, as described in [[34](#page-18-10), [35](#page-18-11)].

# <span id="page-3-0"></span>**Methods**

### <span id="page-3-1"></span>**Datasets**

In this proposed work, three diferent types of datasets to detect ILD were used. The primary dataset [\[36\]](#page-18-12) was made up using the HRCT images. The dataset consists of 3045 HRCT images with three-dimensional annotations of diseased lung tissue areas and diagnostic criteria for pathologically verifed ILD disorders. It includes 108 image series with almost 41 L of annotated lung tissue patterns, 128 individuals with 1 of 13 ILD histological disorders, and a thorough set of 99 medical data. It offers a.txt file with annotations and DICOM images. A statistical dataset was used to train ML models which was created by extracting features from the HRCT images.

### <span id="page-3-2"></span>**Data Preprocessing**

The initial slice thickness of HRCT images were less than 1.5 mm and MRI images varied from 5 to 10 mm. The spatial resolution ranged from 1.34 to 1.68 mm<sup>2</sup>/pixel. During the initial stage of preprocessing, each image was reduced to  $299 \times 299 \times 3$  dimensions. A dataset with 5600 images was created for the training set, 1200 images for the validation set, and 280 images for the testing in DICOM format for both HRCT and MRI images. These DICOM-format images are loaded in our work and afterwards transformed into a NumPy array for DL model training.

The DICOM images include information on the patient and the imaging methods employed. Such data from DICOM images makes it simpler for the model to simply extract additional features during training. Each DICOM image is a single fle, and the header includes all of the necessary data to identify the fle. These data are organised into four levels of hierarchy: patient, study, series, and instance. A patient is the person who is undergoing an examination. The study is an imaging procedure that is carried out at a predetermined time and day at the hospital. Each study contains numerous series. A series can represent a patient who was physically scanned multiple times throughout a study or it might represent a patient who was physically scanned just once and the data were then reconstructed in diferent ways. A three-dimensional image's instance is treated as each slice of the image. A DICOM instance is the actual DICOM fle.

#### **InceptionV3**

The InceptionV3 model, created by Google, consists of 10 blocks with a total of 312 layers. This model has 3 inception blocks, 13 convolutional layers, and 2 pooling layers. A number of  $3 \times 3$  filters with a stride of 2 PX are present in each convolution layer. The fnal layer has the same number of output nodes as categories in the dataset. Each convolution block uses ReLU as the activation function and the SoftMax layer as the classifcation layer. The inception module seeks to behave as a multi-level feature extractor by performing  $1 \times 1$ ,  $3 \times 3$ , and  $5 \times 5$  convolutions within the same network module. Inception vN, where N is the version number disclosed by Google, has replaced the moniker of this architecture's initial iteration, GoogLeNet [\[37](#page-18-13), [38](#page-18-14)].

### **MobileNetV2**

The 16-layer blocks in the MobileNetV2 model have 33 flters and 1 PX as a stride in each layer of convolution. Google has also been working on this idea. The usage of a full convolutional division, which separates the convolution into a 33-depth and a  $1 \times 1$ -pointwise convolution, is the only difference between MobileNet and other CNNs. ReLU served as the activation function in MobileNetV2, while SoftMax was employed for categorization. Two distinct block types can be found in the MobileNetV2 model. Downsizing blocks have a stride of 2, whereas leftover blocks have a stride of 1. For these two blocks, three diferent types of layers are built. The first layer with non-linearity in  $1 \times 1$  convolution is the ReLU6. Convolution based on depth was designed for the second layer. A  $1 \times 1$  convolution with no non-linearity makes up the third layer [[39,](#page-18-15) [40\]](#page-18-16).

#### **VGG16**

One of the most popular pre-trained image categorization models is the VGG-16. The Visual Graphics Group at Oxford University created it. This VGG16, which has 13 convolutional layers, 5 pooling layers, and 3 fully connected layers, was developed using ImageNet weights. It has numerous 33 filters with 1 PX as a stride on each layer of convolution. A SoftMax layer is the fnal layer used for categorization. Each block's activation function utilised the ReLU approach. The most notable aspect of VGG16 is that it constantly used the same padding and maxpool layer of a 22 flter with stride 2 and prioritised convolution layers of a  $3\times3$  filter with stride 1 over several hyper-parameters. The frst convolutional layer has 64 flters, the second has 128 flters, the third has 256 flters, the fourth and ffth have 512 flters [[41](#page-18-17), [42](#page-18-18)].

## **VGG19**

A fixed size of  $(224 \times 224)$  RGB image was given as input to this network which means that the matrix was of shape (224,224,3). The mean RGB value of each pixel, calculated throughout the whole training set, was the only preprocessing that was carried out. They were able to cover the entirety of the image by using kernels that were  $(3 \times 3)$  in size with a stride size of 1 pixel. To maintain the image's spatial resolution, spatial padding was applied. Stride 2 was used to conduct max pooling over a  $2 \times 2$  pixel window. Rectified linear unit (ReLu) was used after this to add non-linearity to the model in order to enhance classifcation accuracy and computation time. As opposed to earlier models that used tanh or sigmoid functions, this one performed far better [\[43](#page-18-19), [44](#page-18-20)].

#### **ResNet50**

The ResNet-50 model is divided into fve stages, each comprising a convolution and an identity block. Each convolution block contains three convolution layers, as do the identity blocks. Over 23 million trainable parameters are available in the ResNet-50. A convolution with 64 distinct kernels, each with a stride of size 2, and a kernel size of  $7 \times 7$  gives us 1 layer. Following that, it use a max pooling with a stride size of 2. The following convolution consists of three layers: a  $1 \times 1,64$  kernel, a  $3 \times 3,64$  kernel, and finally a  $1 \times 1,256$  kernel. These three levels are repeated a total of three times use nine layers in this phase. The kernel of  $1 \times 1,128$  is shown next, followed by the kernel of  $3 \times 3,128$ and, finally, the kernel of  $1 \times 1,512$ . It use this procedure four times for a total of 12 layers. Following that, it has a kernel of size  $1 \times 1,256$ , followed by two more kernels of size  $3 \times 3,256$  and size  $1 \times 1,1024$ ; this is repeated six times, giving a total of 18 layers. Finally, a  $1 \times 1,512$  kernel was added, followed by two more kernels of  $3 \times 3,512$  and  $1 \times 1,2048$ . This process was done three times, giving a total of nine layers. Then, it do an average pool, fnish it with a completely linked layer made up of 1000 nodes, and add a softmax function to produce one layer [\[45](#page-18-21), [46](#page-18-22)].

#### **Transfer Learning**

Transfer learning is a sophisticated deep learning technique that entails training a CNN model on a problem that is comparable to the one being solved. It is possible to avoid having to create a new model from scratch by using this method for feature representation from a previously trained model. It facilitates the flow of knowledge about the issue from one source to another. A pre-trained model is often trained on a large dataset like ImageNet, and the weights gained from the trained model can be used with the custom neural network to solve any other similar problem [\[47](#page-18-23), [48](#page-18-24)]. Through this method, weights are reused to train a model more quickly and produce results with less generalisation error. These recently developed models can be utilised to make predictions directly on relatively untested problems or to train algorithms for related applications. Since all of the pre-trained models have already been trained on a sizable dataset, the fnal layer contains numerous parameters regarding the original dataset. Therefore, a new classifcation layer must be added in place of the pre-trained models' fnal layer. The degree of similarity between the source data and the target data affects how well the model performs [[49,](#page-18-25) [50\]](#page-18-26).

#### **Full‑Training Approach**

Here, five deep learning models termed VGG16, VGG19, ResNet50, MobileNetV2 and InceptionV3 were used for classification of ILD. All models were trained with RMSprop optimizer with learning rate as 0.00001 each for 100 epochs. All layers in the models underwent full training using the ILD datasets. The image acquisition, preprocessing, and data augmentation stages in this method were the same as those in the transfer learning method. Since every model was built from scratch, the training process took longer than transfer learning. The number of layers, blocks, and input image resolution of each model varied. However, since each model was a sequential model, the order of the layers was always sequential. The ImageDataGenerator from keras.preprocessing was used to import each image and its associated label into the models. To prevent sending negative values to the following layers, the ReLU activation function was implemented for each layer. Two units of dense layers with a softmax classifer were added at the end after making all the convolution blocks. The output values from the softmax layer were 0 and 13, with 0 being a healthy case and 1 to 13 for various categories of ILD. The evaluation of each full-training model followed the same procedures as the transfer learning evaluation. Similar to how transfer learning was evaluated, each full-training model was evaluated.

# **ImageNet**

It is a vast collection of labeled images intended for computer vision research. This dataset contains over 14 million photographs with over 21,000 classifcations. The ImageNet LargeScale Visual Recognition Challenge (ILS-VRC) was created specifcally for image classifcation issues with transfer learning in a deep learning framework.

This challenge serves as a standard in image classifcation challenges based on transfer learning.

#### **Soft‑Voting Ensemble**

Soft-voting ensemble is a technique used in ML and DL to combine the predictions of multiple models. In a softvoting ensemble, each model in the ensemble assigns a probability to each possible class label, rather than making a hard prediction. The fnal prediction is then determined by taking the average or weighted average of the predicted probabilities from all the models. Once the models are trained, they are used to generate predictions on a new, unseen dataset (e.g., a validation or test set). Instead of making binary or categorical predictions, each model assigns probabilities to each possible class label. For example, if there are three classes (A, B, and C), a model might assign probabilities [0.2, 0.6, 0.2] for a given instance, indicating a higher likelihood for class B. To determine the fnal prediction, the predicted probabilities from all the models are combined. One common approach is to take the average of the predicted probabilities for each class across all models [\[51,](#page-18-27) [52\]](#page-18-28). Alternatively, if certain models are deemed more reliable or accurate, their predictions can be weighted more heavily in the ensemble. Weighted averaging involves assigning weights to each model's predicted probabilities based on their performance or confdence. Once the aggregated probabilities are obtained, the fnal prediction can be made by selecting the class label with the highest probability. In some cases, multiple class labels may have similar probabilities, leading to a tie. In such situations, further tie-breaking techniques can be applied, such as selecting the class label with the highest confdence from a specifc model or applying a predetermined rule. The performance of the soft-voting ensemble is assessed using appropriate evaluation metrics, such as accuracy, precision, recall, or F1 score, on a separate evaluation dataset. These metrics help gauge the efectiveness of the ensemble in making accurate predictions compared to individual models or other ensem-ble techniques [\[53,](#page-18-29) [54\]](#page-18-30). Prediction is almost identical to the preceding example, but since this is a binary classifcation problem, utilise only class B:

Classifer 1 correctly predicts class B with a probability *m*.

Classifer 2 correctly predicts class A with a probability *q*.

Classifer 3 correctly predicts class B with a probability *v*.

Therefore, class B will be predicted by the ensemble model with probability  $p = (m + (100-q) + v)/3$ .

#### **Proposed Approach**

The plan of our research is depicted in Fig. [1](#page-6-0) which includes HRCT and MRI dataset acquisition, data extraction, preprocessing, data augmentation, selection of pre-trained models for transfer learning, feature extraction, classifcation using the VGG16, VGG19, ResNet50, InceptionV3, and Mobile-NetV2 models, and ensemble using soft-voting and hardvoting techniques are the steps that make up this process. The images from the converted ILD dataset were in RGB format with a pixel range of [0,255]. During the preprocessing phase, all of those photos were rescaled into the [0,1] range to meet the requirements of the pre-trained model. The classifcation layer of pre-trained models may not be useful for the new classifcation problem in transfer learning. As a result, we replaced it with a completely connected layer at the top layer of each model. Because the pre-trained network was frozen, only the weights of the top four layers and one classifer layer were changed during training. As a classifcation layer, a SoftMax layer was utilised on top of all models in this suggested strategy. In several experiments, Adam and RMSprop optimizers with various learning rates were used to train 100 epochs. For the process of fne-tuning all the models, we took into account a variety of factors, including the number of trainable layers, including the original and additional added layers, epochs, learning rate, and optimizers. Then, utilising statistical data, ML models such as Logistic, BayesNet, SGD, RandomForest, and J48 were employed to detect ILD. The statistical dataset was constructed by extracting and storing eighteen diferent features from HRCT images in a CSV fle.

The soft-voting ensemble was applied on the proposed models, transfer learning model and ML models separately to achieve better accuracy. Later, we again applied second-tier soft-ensemble by utilizing the prediction value from each ensemble training which can ensure the prediction more accurately. The primary advantage of a two-tier soft-voting ensemble is the potential for improved prediction accuracy. By combining predictions from multiple models in a hierarchical manner, the ensemble can leverage the strengths of each model to make more accurate predictions. The frst tier of models can capture diferent aspects or patterns of the data, while the second tier integrates their predictions for a fnal decision. This can help reduce bias and increase overall accuracy. A two-tier soft-voting ensemble can enhance the robustness of the prediction process. Since multiple models are involved, the ensemble can handle noisy or uncertain data more efectively. If one model produces erroneous predictions due to outliers or specifc data biases, the infuence of that model can be reduced or mitigated at the second-tier voting stage. This robustness makes the ensemble more reliable in real-world scenarios. Two-tier soft-voting ensembles encourage model diversity, which can



<span id="page-6-0"></span>**Fig. 1** The architecture diagram of the proposed work

lead to better performance. The frst-tier models are typically diverse in terms of their algorithmic choices, hyperparameter settings, or input data variations. This diversity helps to capture diferent perspectives or representations of the underlying data, enhancing the ensemble's ability to generalize and make accurate predictions on unseen instances. Ensemble methods, including two-tier soft-voting ensembles, tend to reduce overftting. Overftting occurs when a model becomes too specifc to the training data and fails to generalize well to new data. By combining predictions from multiple models, the ensemble can mitigate the risk of overftting by averaging out individual model biases or errors, resulting in improved generalization performance. Two-tier soft-voting ensembles offer flexibility and adaptability in terms of incorporating new models or replacing existing ones. If a more accurate or efficient model becomes available, it can be introduced into the ensemble at either the frst or second tier. This adaptability allows the ensemble to evolve and improve over time as new models or algorithms emerge. Another advantage of two-tier soft-voting ensembles is their potential to provide more interpretable predictions. By using a hierarchical structure, the ensemble can provide insights into the decision-making process. For example, the frst-tier models' predictions can be analyzed to understand which aspects or features of the data contribute more strongly to the fnal decision. This interpretability can be valuable in domains where explainability is crucial, such as healthcare or fnance.

#### **CNN**

The fundamental building block of any model that works with picture data is a Convolutional Neural Network. Visuals were considered when developing convolutions. There is an n-dimensional weights or flter matrix, where n is typically smaller than the image size. The input's flter size patch is used to multiply or dot product this matrix. The flter is applied sequentially to each overlapping section or fltersized patch of the incoming data, working from left to right and then from top to bottom [\[55](#page-18-31)]. CNN architecture is based on convolutional layers. The convolution layers transform the image's data before passing it on to the next layer as input. The transformation is referred to as the convolutional operation. It is necessary to specify the number of flters for each convolution layer. Patterns in objects, textures, edges, forms, curves, and even colors are what these flters look for. It identifes things or patterns with deeper layers that are more intricate. An image kernel, which can be described as a tiny  $3 \times 3$  or  $4 \times 4$  matrix applied to the entire image, is the fundamental component of a flter.

The images' input shape are (299,299,3), because their height and width were previously specifed. And the number 3 stands for the colour channel, which is represented by the fact that the images are RGB. The output size is  $(299-3+1,$ 299−3+1)=(298,298) when a First Conv2d layer Convolution operation is performed on an image of (299,299) with a kernel size of (3,3), strides and dilation are set to 1 by default, and padding is set to 'valid.' Since we defned 32 flters, the output shape is now (None,297,297,32). We obtain  $((297-2/2)+1,(297-2/2)+1)=(148,148)$  assuming that the input image size is (297,297), that the kernel size of the frst Max Pooling layer is (2,2), and that strides are by default (2,2). The Flatten layer creates a one-dimensional vector from all of the pixels along all of the channels without taking batch size into account. The input of  $(7, 7, 64)$  is flattened to  $(7 \times 7 \times 64) = 3136$  values as a result. Piecewise linear function known as the rectifed linear activation function, the short-term ReLU directly outputs the input if the input is positive; if not, it will return zero. The corrected linear activation function makes it possible for models to learn more quickly and perform better by addressing the issue of vanishing gradients.

### <span id="page-7-0"></span>**Results and Analysis**

We assessed the model's performance using various accuracy metrics. The training and validation accuracies were measured to ensure that the model obtained enough knowledge during training and that overfitting was kept to a minimum. After training, the testing accuracy was estimated using 200 test images. We determined the average training and validation accuracies of the last 20 epochs for all of these metrics. Individual model outputs were subjected to soft-voting ensemble technique.

#### **Evaluating Full‑Training Models**

The outputs of full-training of individual models and soft-voting ensembles are presented in Table [1](#page-8-0). In all of the experiments, the epoch accuracies grew over time. The InceptionV3 model, which underwent 30 training epochs, had the greatest test accuracy of 69.37%, while the VGG16, mobileNetV2, VGG19, and resNet50 models provided accuracy of 68.39%, 67.56%, 66.2%, and 67.42%, respectively. But at this point, the soft-voting ensemble accuracy had reached 76.56%, surpassing the test accuracy from InceptionV3. During this time, the InceptionV3 model likewise had the highest training accuracy of 72.35% and validation accuracy of 71.63%. The diference between the training and validation scores of all fve models was just about 2%, ensuring that no overftting occurred during the models' training. After 60 epochs of training, the MobileNetV2 model had the greatest training and test accuracies with 74.86% and 72.72%, respectively, whereas InceptionV3 had the highest validation accuracy with 73.44%. However, the ensemble score of 73.14% was somewhat higher than the test accuracy of the Mobile-NetV2 model. Among all the diferent models, the InceptionV3 had the best training accuracy with 87.59%, validation accuracy with 84.95%, and test accuracy with 76.11%, while VGG16 had the lowest test accuracy with 78.41% after completing training of 100 epochs. All individual models' gaps between training and validation scores were about 3% or less, ensuring that there were no overftting issues. All three of these models were used in an ensemble with soft voting, and the accuracy was 76.56%.

The training graph of fully trained InceptionV3 model is given in Fig. [2](#page-8-1). It includes the accuracy and loss measurements needed to assess the model's performance on both training and validation datasets. The fgures clearly illustrate that no overftting occurred during the model's training. For each epoch, the training and validation accuracies gradually improved. There was considerable volatility in both accuracies at the start of the programme, but

<span id="page-8-0"></span>**Table 1** The results from fulltraining models

Model	<b>Trained layers</b>	Epochs	Training Acc $(\%)$	Validation Acc $(\%)$	Testing Acc $(\%)$
Inception V3	316	30	72.35	71.63	69.37
Inception V3	316	60	74.21	73.44	72.26
Inception V3	316	100	87.59	84.95	78.41
VGG16	23	30	70.48	68.54	66.39
VGG16	23	60	72.34	70.83	68.37
VGG16	23	100	76.26	74.14	73.52
MobileNetV2	158	30	72.63	71.47	67.56
MobileNetV2	158	60	74.86	73.21	72.72
MobileNetV2	158	100	77.06	76.19	75.32
VGG19	26	30	69.36	67.76	66.2
VGG19	26	60	72.03	70.62	68.18
VGG19	26	100	76.38	74.28	73.86
ResNet50	55	30	70.64	67.94	67.42
ResNet50	55	60	73.26	71.52	70.08
ResNet50	55	100	76.56	74.36	74.28
Ensemble		30			70.08
Ensemble		60			73.14
Ensemble		100			76.56

<span id="page-8-1"></span>



this settled down as the training progressed. It is also seen that the loss measurements dropped steadily until the fnal epoch. Based on this research, we can infer that the model produced efficient outcomes with no overfitting.

#### **Evaluating Transfer Learning Models**

Table [2](#page-9-0) lists the results of soft-voting ensembles and individual models for transfer learning. In all transfer learning experiments, the accuracy grew with the passing of each epoch. After 30 training epochs, the InceptionV3 model had the highest test accuracy of 79.28%, while the test accuracies of the VGG16, MobileNetV2, VGG19, and ResNet50 were 76.36%, 78.52%, 76.68%, and 78.26%, respectively. The soft-voting ensemble accuracy, which was higher than the Inception V3 test accuracy at the time, was 79.48%. The InceptionV3 model likewise had the highest validation accuracy with 81.74%, whereas

<span id="page-9-0"></span>**Table 2** The results from transfer learning models



ResNet50 had the highest training accuracy with 82.13%. There was no overftting during model training as evidenced by the less than 2% variations in scores between training and validation for all fve models. The InceptionV3 model once more showed the best test accuracy of 85.31% after 60 training epochs. However, the ensemble score of 85.62% was a little higher than the InceptionV3 model. At that time, the InceptionV3 showed the highest training accuracy with 88.26% and validation accuracy with 87.32%. Among all the individual models, it also achieved the best test accuracy with 92.48%, training accuracy with 94.64%, and validation accuracy with 94.52%, whereas the least test accuracy of 89.22% for 100 epochs resulted from VGG16. The accuracy of the soft-voting ensemble of all fve models was 92.81%, the highest so far. All individual model gaps between training and validation scores were less than 2%, ensuring that no overftting concerns existed. The training graph of transfer learning MobileNetV2 model is given in Fig. [3.](#page-9-1)

<span id="page-9-1"></span>

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<span id="page-10-0"></span>**Table 3** Training results of all developed CNN models using RMSprop optimizer



#### **Evaluating Newly Constructed CNN Models**

We developed three CNN models for ILD prediction and results from those models are demonstrated in Table [3.](#page-10-0) ILDNetV1 achieved highest training accuracy of 96.37%, validation accuracy of 94.88% and test accuracy of 94.08% after completing 100 epochs of training. The accuracy trend was same at 30th and 60th epochs break. The ILDNetV2 and ILDNetV3 also performed well and reached test accuracy of 92.04% and 93.72% after 100 epochs of training. We applied soft-voting ensemble to achieve better accuracy since all models were achieved good results without any over-ftting. The ensemble test accuracies were higher than all three developed CNNs with test accuracy of 84.12% for 30 epochs, 88.17% for 60 epochs and 97.42% for 100 epochs. The training graph of developed CNN ILDNetV1 is given in Fig. [4.](#page-10-1)

In subsequent experiments, alternative optimizers, including Adam, Adagrad, and Adadelta, were employed following the same methodology as the previous employment of the RMSprop optimizer. Employing these optimizers, the newly devised trio of CNN models underwent training for a total of 100 epochs. The outcomes of these training experiments are presented in Table [4.](#page-11-0) Despite the notable performance of the Adam optimizer, achieving an ensemble accuracy of 94.86%, it fell short of matching the level set by the RMSprop optimizer. The utilization of the Adagrad optimizer led to discernible accuracy discrepancies between training and validation scores. The ensemble test accuracy attained using Adagrad reached a modest 92.76%. Although the disparity between training and validation accuracies observed with the Adadelta optimizer was less than with Adagrad, the resultant accuracy metrics remained below expectations. After a hundred epochs of training, the

<span id="page-10-1"></span>



SN Computer Science A SPRINGER NATURE journal <span id="page-11-0"></span>**Table 4** Training results of all developed CNN models using other optimizers



Adadelta optimizer yielded an ensemble accuracy of merely 91.84%. Through these conducted experiments, it is evident that the RMSprop optimizer exhibited superior performance compared to the Adam, Adagrad, and Adadelta optimizers. In our modeling process, we implemented the EarlyStopping method in Python, which consistently terminated training for all models upon reaching 100 epochs. This strategic utilization of the EarlyStopping technique efectively mitigated issues related to overftting. The rationale behind this decision lies in the observation that, following the 100th epoch, there was limited discernible improvement in the validation accuracy over the subsequent six epochs. Consequently, we adopted a standardized training protocol of limiting all models to a maximum of 100 epochs.

# **Evaluating ML Classifers**

Then we used ML classifers such as Logistic, BayesNet, SGD, RandomForest and J48 for detecting ILD using statistical data. The statistical dataset was created by extracting eighteen diferent features from HRCT images and storing it in CSV fle. We considered fve ILD combinations to train the ML classifers. The combinations were created using fve ILD cases such as Consolidation, Fibrosis, Groundglass,

Micronodules and Reticulation with Healthy cases. The J48 showed highest accuracy with all combinations (Table [5](#page-11-1)). Among all the combinations, J48 achieved 93.08% accuracy with C1 combination which was the highest accuracy from ML models. The other accuracy measurements of this model is given in Fig. [5.](#page-12-0)

Our next target was to fnd out which features contribute more in accuracy. Therefore, we considered the highest accuracy providing J48 with each features separately for training. The training results of J48 with each feature are given in Table [6.](#page-12-1) It shows that all features are contributing almost equal level in accuracy even though diagonal:SD (standard deviation) was the highest. The highest accuracy of 85.20% was obtained from Healthy\_Micronodules combination for diagonal:SD feature.

In the next phase, the Gabor flter was applied on the HRCT and trained all ML models in the same manner. Gabor flters are used to analyze textures in images and are widely used in computer vision and image processing tasks. The image were converted into the binary format with pixel values either 0 or 1. The threshold value is set to 127. The J48 showed highest accuracy this time too. It achieved 95.55% accuracy with C1 combination which was the highest accuracy from ML models (Table [7\)](#page-13-0).

<span id="page-11-1"></span>



C1: Healthy\_Consolidation C2: Healthy\_Fibrosis C3: Healthy\_GroundGlass C4: Healthy\_Micronodules C5: Healthy\_Reticulation

<span id="page-12-0"></span>**Fig. 5** Measurement scores from developed CNN ILD-NetV1



=== Confusion Matrix ===

a b <-- classified as

123 42 |  $a =$  consolidation

11 590  $|$  b = Healthy

<span id="page-12-1"></span>



0.846 consolidation 0.957 Healthy

0.933

In the next step, the highest accuracy providing J48 was considered again with each features separately for training. The training results of J48 model with each feature are given in Table [8](#page-13-1). It demonstrates that all features contribute almost equally to accuracy: The highest diagonal:SD was found. The Healthy\_Micronodules combination achieved the highest diagonal accuracy of 86.43% diagonal:SD feature.

# **Evaluating ML Classifers After Applying Segmentation and Masking**

The marker-controlled watershed transformation segmentation technique was applied on the images to achieve better results (Table [9\)](#page-14-0). Marker-controlled watershed transformation is a segmentation technique used in image processing and computer vision. It is based on the concept of <span id="page-13-0"></span>**Table 7** Training results of all used ML classifers after applying Gabor flter

ML classifers Accuracy from  $C1 (%)$ Accuracy from  $C2$   $(\%)$ Accuracy from C3 (%) Accuracy from C4 (%) Accuracy from C5  $(% )$ Logistic 86.70 51.48 54.10 65.40 80.95 BayesNet 83.62 82.66 86.31 89.13 79.19 SGD 89.92 76.10 72.72 87.75 81.33 RandomForest 87.24 82.30 85.17 89.75 84.36 J48 95.55 85.01 88.45 94.78 91.29

<span id="page-13-1"></span>

	Table 8 Training results of J48
for each feature	



watersheds, which are regions in an image that are separated by watershed boundaries. In marker-controlled watershed transformation, markers or seed points are manually or automatically placed in the image to guide the segmentation process. These markers indicate the regions of interest or objects that need to be segmented. The watershed transformation algorithm then treats the image as a topographic surface, where the intensity values represent the elevation. The algorithm foods the image from the markers, and as the water level rises, it forms basins around the markers. The boundaries between these basins are the watershed boundaries, which represent the segmentation result. The markercontrolled watershed transformation is particularly useful when dealing with complex images that have overlapping or touching objects. By providing accurate markers, the algorithm can accurately segment the objects of interest.

The morphological masking approach was utilized to create a mask for the images of the dataset as shown in

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Table [10](#page-14-1). Morphological masking is a technique used in image processing and computer vision to extract or enhance specifc regions or features in an image using morphological operations. Morphological operations are mathematical operations that manipulate the shape and structure of objects in an image. The two most commonly used morphological operations are erosion and dilation. In morphological masking, a binary mask, also known as a structuring element, is defned to represent the desired region or feature to be extracted or enhanced. The structuring element is a small binary image that defnes the shape and size of the neighborhood around each pixel. To apply morphological masking, the structuring element is scanned over the image, pixel by pixel. At each pixel location, the corresponding pixels in the image and the structuring element are compared. If the pixels match according to a specifc condition, the pixel at that location is preserved or modifed based on the desired operation.

#### <span id="page-14-0"></span>**Table 9** Marker-controlled watershed transformation segmentation on HRCT images

# Marker Sample image

Internal marker

External marker



#### Watershed marker

<span id="page-14-1"></span>**Table 10** Morphological masking on HRCT images



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<span id="page-15-0"></span>**Table 11** Training results of all used ML classifers after applying segmentation and masking



The same five ML models were again used to assess the performance of these segmentation and masking techniques. The J48 model showed highest accuracy this time too. It achieved 95.73% accuracy with C1 combination which was the highest accuracy from ML models (Table [11\)](#page-15-0).

In the following step, the ML model with the highest accuracy, the J48 model, was looked at once more, with each feature used separately for training. Table [12](#page-15-1) displays the J48 model's training results for each feature. It exhibits that all highlights contribute similarly to precision: The most extreme diagonal: SD was found. With an accuracy of 86.60%, the Healthy\_Micronodules combination had the best diagonal accuracy: SD include.

# **Assessing Performance of the Proposed Two‑Tier Ensemble Framework Using the Untrained Images**

To check the efficiency of our proposed work for classifying and categorizing ILD diseases, we applied some untrained HRCT images of various ILD category into the trained model. The proposed approach was able to predict ILD category accurately with high similarity score. The predictions were there happened by considering the majority voting ensemble mechanism with predictions from our own three newly developed CNN models, fve pre-trained deep learning models and fve ML models. The predictions and their merging from the various models ensure the sufficiency of prediction results and capacity and efficiency for our system's prediction ability. Some samples of such prediction results are demonstrated in Figs. [6](#page-16-0) and [7.](#page-16-1)

The given results represent the performance of our proposed two-tier ensemble prediction approaches for



<span id="page-15-1"></span>**Table 12** Training results of J48 model for each feature after applying segmentation and masking



**Fig. 6** Model predication for frst sample image

<span id="page-16-0"></span>

**Fig. 7** Model predication for second sample image

<span id="page-16-1"></span>classifying ILD using two diferent untrained HRCT Images. It combines the predictions of multiple base models or classifers to make a fnal prediction. According to Fig. [5](#page-12-0), the ensemble prediction from proposed CNN models achieves a test accuracy of 98.23% and predicts the ILD category as "reticulation". In the transfer learning ensemble approach, the test accuracy is slightly lower than the proposed CNN ensemble, at 96.74%. It also predicts the ILD category as "reticulation". The Ensemble Prediction from Machine Learning achieves a test accuracy of 97.12% and, like the previous two, predicts the ILD category as "reticulation". This is the fnal ensemble prediction that combines the outputs of the previous three ensembles or models. It achieves a test accuracy of 97.36% and predicts the ILD category as "reticulation". All three ensemble approaches (Proposed CNNs, Transfer Learning, and Machine Learning) perform well in classifying ILD categories, with high test accuracies ranging from 96.74 to 98.23%. The fnal 2nd Tier Ensemble maintains a high accuracy of 97.36%. Since it is a softvoting ensemble approach, it predicts the ILD category as "reticulation" with all three votes, because all three frst-tier ensemble approaches also predict the same ILD category.

According to Fig. [6](#page-16-0), the result suggests that the proposed CNN models achieved an ensemble test accuracy of 99.06% when predicting the ILD category, and it classifed the ILD as "consolidation". The transfer learning ensemble approach achieved a test accuracy of 94.28%. The predicted ILD category was "fbrosis". The ensemble prediction approach which used machine learning techniques, achieved a test accuracy of 97.84%. It classifed the ILD as "consolidation". The fnal prediction appears to be a combination of the predictions from the three previous ensemble methods in a second-tier ensemble approach. The resulting predicted ILD category is "consolidation", and the test accuracy is 98.45%. The reason for predicting the category as "consolidation" is that it achieved most votes (from proposed CNN and Machine learning ensemble), only ensemble of transfer learning predicted a diferent category. It seems that the ensemble prediction method using multiple techniques has achieved high test accuracy for ILD category prediction. This approach likely aims to increase prediction accuracy by leveraging the strengths of various models or techniques.

# **Conclusion**

In this proposed work, a new deep learning approach was proposed to detect ILD using HRCT images. In the frst stage, three CNN models were developed and also used fve full trained and fve pre-trained models on the ImageNet dataset such as VGG16, VGG19, ResNet50, MobileNetV2, and InceptionV3 to detect ILD from HRCT images. Our proposed frst model achieved highest individual model test accuracy with 94.08% than all other deep learning models. In each experiment, the RMSprop optimizers were used to train the models for 100 epochs using 25,000 augmented images generated from 2000 original images. In the second stage, a new algorithm was developed to extract various features from HRCT images and applied those features into fve ML algorithms such as Logistic, BayesNet, SGD, RandomForest and J48. The J48 model showed better accuracy among ML models which reached highest accuracy of 93.08%. The diagonal-wise standard deviation feature was contributing more on models' training by analyzing each features separately with J48 model. The ensemble of models helped to achieve better results. Therefore, the majority voting mechanism with each kind of models separately was applied in the third stage. We achieved highest ensemble test accuracy of 97.42% after ensemble of all our three newly built CNN models. In the fnal stage, the two-tier ensemble concept was applied. The outputs of these ensemble models were fed into majority voting ensemble once again which helped to ensure the reliability of the model's prediction on unseen images. Our results suggest that the proposed approach can be used to improve ILD detection accuracy compared to other deep learning methods, which may assist the doctors. When compared with the previous state-of-theart methods, our approach achieved better results.

**Supplementary Information** The online version contains supplementary material available at<https://doi.org/10.1007/s42979-023-02524-3>.

**Funding** No funding received.

**Data Availability** The data that support the fndings of this study are available from St. John's Medical College, Bengaluru, India but restrictions apply to the availability of these data, which were used under licence for the current study, and so are not publicly available. Data are, however, available from the authors upon reasonable request and with permission of St. John's Medical College, Bengaluru, India.

#### **Declarations**

**Conflict of Interest** No confict of interest.

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