



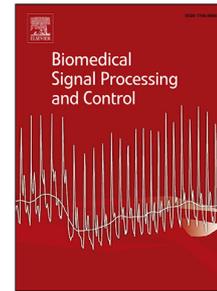
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Automated diagnosis of COVID-19 using radiological modalities and Artificial Intelligence functionalities: A retrospective study based on chest HRCT database

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Automated Diagnosis of COVID-19 Using Radiological Modalities and Artificial Intelligence Functionalities: A Retrospective study based on chest HRCT database

Abstract

(Background and Objective): The spread of coronavirus has been challenging for the healthcare system's proper management and diagnosis during the rapid spread and control of the infection. Real-time reverse transcription-polymerase chain reaction (RT-PCR), though considered the standard testing measure, has low sensitivity and is time-consuming, which restricts the fast screening of individuals. Therefore, computer tomography (CT) is used to complement the traditional approaches and provide fast and effective screening over other diagnostic methods. This work aims to appraise the importance of chest CT findings of COVID-19 and post-COVID in the diagnosis and prognosis of infected patients and to explore the ways and means to integrate CT findings for the development of advanced Artificial Intelligence (AI) tool-based predictive diagnostic techniques.

(Methods): The retrospective study includes a 188 patient database with COVID-19 infection confirmed by RT-PCR testing, including post-COVID patients. Patients underwent chest high-resolution computer tomography (HRCT), where the images were evaluated for common COVID-19 findings and involvement of the lung and its lobes based on the coverage region. The radiological modalities analyzed in this study may help the researchers in generating a predictive model based on AI tools for further classification with a high degree of reliability.

(Results): Mild to moderate ground glass opacities (GGO) with or without consolidation, crazy paving patterns, and halo signs were common COVID-19 related findings. A CT score is assigned to every patient based on the severity of lung lobe involvement.

(Conclusion): Typical multifocal, bilateral, and peripheral distributions of GGO are the main characteristics related to COVID-19 pneumonia. Chest HRCT can be considered a standard method for timely and efficient assessment of disease progression and management severity. With its fusion with AI tools, chest HRCT can be used as a one-stop platform for radiological investigation and automated diagnosis system.

Keywords: COVID-19, Ground glass opacities, Consolidation, Crazy paving, Halo Sign, Machine Learning, Deep learning.

1. Introduction

With its emergence in late 2019, the persisting pandemic, namely the novel coronavirus (COVID-19), has drastically affected people across the globe. The outbreak of this deadly disease was caused by the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2). The rapid transmission of this disease from human to human is phylogenetically close to SARS-like coronavirus with a separate monophyletic group [1]. The SARS-CoV-2 virus is a member of the human coronavirus (HCoV) family that affects the lower respiratory tract. Due to the high number of confirmed COVID-19 cases and no treatment for the infection, safety precautions are implemented worldwide, such as social isolation, the use of a mask to prevent the virus from entering the respiratory system, quarantine and other containment measures, which are able to lower morbidity and mortality among highly susceptible individuals [2]. Still, studies reveal that, unlike the SARS-CoV and MERS-CoV viruses, the mortality rate is lower in the case of COVID-19 [3]. SARS-CoV-2 is largely spread through droplet inhalation, indirectly through contact with infected fomites, and through airborne inhalation of bioaerosols that are suspended in the air. The symptoms of COVID-19 are inconsistent, but com-

mon symptoms include fever, difficulty in breathing, cough, fatigue, loss of smell, taste, and headache [4]. These symptoms may occur between one to fourteen days if exposed to the virus. An infected person may experience a few of these symptoms, and it may last for a minimum of seven days. If the severity of symptoms continues for more than seven days, then there are chances that the patient may include symptoms like dyspnea, hypoxia, and the involvement of 50% of the lung. Considering the impact factors and statistics, studies disclose that COVID-19 is a severe case of pneumonia that affects the lungs. Such symptomatic patients are easy to identify and take corrective measures for. But the situation worsens when the patient is asymptomatic COVID-19 positive. Even though they often go unnoticed, these people may be contagious and help SARS-CoV-2 spread to healthy people. Despite the fact that such patients are responsible for fewer secondary infections than symptomatic cases, asymptomatic COVID-19 subjects can still spread SARS-CoV-2 to other people. It is noteworthy that asymptomatic COVID-19 cases have viral loads that are equal to or higher than symptomatic cases [5]. Asymptomatology and high viremia can thus coexist and represent a significant risk factor for the spread of COVID-19 infection [6]. Because they don't exhibit symptoms that would make one wonder if they

46 have COVID-19. Several variants of SARS-CoV-2 have been¹⁰³
 47 brought into focus by the World Health Organization (WHO),¹⁰⁴
 48 designated as the Alpha, Beta, Gamma, Delta, and Omicron¹⁰⁵
 49 variants [7]. With their potential for rising transmission and¹⁰⁶
 50 virulence, these variants have contributed to the persistence of¹⁰⁷
 51 this pandemic. The SARS-CoV-2 Delta strain was discovered¹⁰⁸
 52 in India in late 2020 and has since been discovered in almost 60¹⁰⁹
 53 other countries [5][6]. A higher rate of transmission is possible¹¹⁰
 54 in Delta compared to other SARS-CoV-2 variations. Testing¹¹¹
 55 and the infected patients' record study revealed that the Delta¹¹²
 56 variation may result in more serious lung deformities, leading to¹¹³
 57 death in extreme cases. While mass vaccination campaigns are¹¹⁴
 58 currently just getting started, several medications have shown¹¹⁵
 59 in vitro activity against SARS-CoV-2 or potential clinical bene-¹¹⁶
 60 fits. Early identification and appropriate treatment of immuno-¹¹⁷
 61 logic complications can reduce morbidity and mortality in pa-¹¹⁸
 62 tients with COVID-19 infection. The COVID-19 vaccination¹¹⁹
 63 was first administered in India on January 16, 2021. India had¹²⁰
 64 provided more than 2.04 billion doses of currently recognized¹²¹
 65 vaccinations as of August 29, 2022, including the first, second,¹²²
 66 and booster doses. In India, 86.87% of the eligible popula-¹²³
 67 tion (12+) is fully vaccinated, and 94.48% of the eligible pop-¹²⁴
 68 ulation (12+) has received at least one shot [[https://vaccinate-¹²⁵](https://vaccinate-india.in/dashboard)
 69 [india.in/dashboard](https://vaccinate-india.in/dashboard)].¹²⁶

70 It is also true that contaminated vaccine recipients may con-¹²⁷
 71 veniently show only moderate symptoms, stay asymptomatic,¹²⁸
 72 or otherwise go unreported. When significant viral loads are¹²⁹
 73 proven, vaccine recipients may spread the infection even more¹³⁰
 74 subtly. This is due to the possibility that participants in the¹³¹
 75 COVID-19 mass vaccination campaign would neglect their¹³²
 76 need to maintain social distance. Therefore, it's crucial to con-¹³³
 77 sistently follow hygiene practices including hand washing, face¹³⁴
 78 mask use, and other public health safety precautions, such as¹³⁵
 79 social isolation. Additionally, it is important to promote the¹³⁶
 80 proper use of personal protection equipment, such as surgical¹³⁷
 81 masks and filtering face pieces, in high-risk settings, includ-¹³⁸
 82 ing crowded areas, public transportation, and indoor establish-¹³⁹
 83 ments like schools or hospitals. Additionally, it is important¹⁴⁰
 84 to improve occupational health safety practices, such as health¹⁴¹
 85 surveillance, screening, testing mainly self-testing with antigen¹⁴²
 86 tests, and contact tracing activities [4]-[10].¹⁴³

87 The reverse transcription-polymerase chain reaction (RT-¹⁴⁴
 88 PCR) is considered the primary diagnostic test for COVID-19,¹⁴⁵
 89 which can analyze thousands of samples in a single day and¹⁴⁶
 90 has a testing sensitivity of 95%. Thus, the RT-PCR technique¹⁴⁷
 91 is regarded as the gold standard for both qualitative and quan-¹⁴⁸
 92 titative viral nucleic acid detection. The lesser sensitivity in¹⁴⁹
 93 the RT-PCR method demonstrates various analytical issues, in-¹⁵⁰
 94 cluding human mistakes, testing outside the diagnostic window,¹⁵¹
 95 active viral recombination, and insufficiently validated assays,¹⁵²
 96 which compromise the diagnostic accuracy [2][11][12]. The¹⁵³
 97 viral loads in throat swabs are most substantial when the virus¹⁵⁴
 98 first manifests, and the virus may start to shed two to three days¹⁵⁵
 99 prior to the beginning of symptoms, making presymptomatic¹⁵⁶
 100 or asymptomatic transmission easier. Therefore, instead of¹⁵⁷
 101 considering it as a diagnostic standard, we may consider RT-¹⁵⁸
 102 PCR as the primary detection measure. As lung involvement¹⁵⁹

is a part of coronavirus infection, the physicians suggest chest
 computer tomography (CT) as a mandate for the proper diag-
 nosis and prognosis of COVID-19 in its early detection due to
 its high sensitivity (60%-98%), of a viral lung [11]-[13]. Al-
 though chest CT imaging is not considered a standard screening
 test protocol for COVID-19, it is beneficial for people with mild
 symptoms or asymptomatic ones or those with a negative RT-
 PCR experiencing mild symptoms or chest anomalies, or un-
 explained lung pathology. Nevertheless, chest CT also, proves
 to be helpful for recovered cases of COVID-19 (post-COVID).
 Other imaging techniques like X-ray and ultrasound also help
 to evaluate the disease progression, but diagnosis with chest CT
 is preferred due to its three-dimensional pulmonary view and
 versatility [14].

Decision making with the detection of chest CT is typically
 based on several parameters such as whether the patient is
 RT-PCR positive, whether the patient has post-COVID symp-
 toms or any other disease with overlapping distortion in the
 chest/lung. Moreover, diagnosis with the help of chest CT
 sounds advantageous in both alternative diagnosis, prognosis,
 and figuring out complications in COVID-19, reducing the rate
 of severity and mortality in cases [15]. While monitoring these
 parameters in manual and traditional way, in many cases it is
 time consuming, tedious and repetitive. Machine Learning
 (ML) and Deep Learning (DL) tools may provide a helping
 hand to the physicians so as to increase the screening rate [16]-
 [19]. The automated ML and DL algorithms can be used in
 biomedical platforms and can be used as AI-based tools to de-
 sign predictive models. These models, deployed on a computer
 system, learn to detect, classify, or diagnose the CT images.
 The AI-aided predictive diagnostic tool may be configured to
 avoid the tedious and repetitive task of manually evaluating
 each CT image of COVID patients. Whereas AI enables the
 computer to understand from a large set of databases, to iden-
 tify or classify diseased cases, a proper methodology to config-
 ure the system is required to design and deploy it. In a previ-
 ous work [20] on methods for COVID-19 detection, the authors
 have reviewed the current ongoing research with the reported
 databases. In that work, the authors compared various algo-
 rithms and schemes for classification, based on the reported
 findings. As a summary of that paper, in a nutshell, engineers
 need to use the signal processing on these CT scan images with
 ML and DL tools to help the physicians screen at a faster rate
 [21]. In this direction, the first criteria is to understand the CT
 scan images and their findings so that they can be incorporated
 into the algorithms for better identification and classification
 of diseased cases. Studies have brought into focus several
 initial chest CT findings in COVID-19 positive cases, which
 include peripheral ground-glass opacity (GGO), consolidation,
 GGO with consolidation, and crazy paving [11][23]-[25]. The
 later stage of the disease shows less common findings such as
 bronchiectasis, septal and pleural thickening, and subpleural
 involvement. These findings are expected in COVID-infected
 lungs and other viral infections or diseases. A patient with RT-
 PCR positive and the above-mentioned findings is considered
 for COVID-19 infected cases.

Therefore, this study aims to understand the CT image find-

Table 1: CO-RADS level of suspicion for COVID infection [22]

CORADS	Class	CT Findings
CO-RADS 1	Nil	Healthy or Non-infectious anomalies
CO-RADS-2	Minimal	Infectious non-COVID anomalies
CO-RADS-3	intermediary	May be infectious with COVID-19, status unclear
CO-RADS-4	Moderate	Infectious with COVID-19 suspicion
CO-RADS-5	Severe	COVID-19 typical
CO-RADS-6	RT-PCR +	

ings so that they can be incorporated into algorithms of ML and DL with a focus on (a) the role of chest HRCT in the diagnosis and prognosis of COVID-19, (b) significant radiological findings, and (c) its severity based on CT score. Further, in this search work, we characterized HRCT findings in 188 patients presenting a retrospective study on the collected database from Marwari Hospitals, Guwahati, Assam, India. The findings of the database have been compiled in association with a team of radiologists from the Dr. Bhubaneswar Borooah Cancer Institute (BBCI). This study aims to appraise the usefulness of the spectrum of HRCT chest findings on COVID-19 cases and estimate the infection's severity, to rule out the findings for further processing, and explore the possibility of incorporating different AI tools for the design of a predictive diagnostic framework for fast confirmatory screening of COVID-infected patients.

2. Chest CT and Significance of Radiological Modalities

In this section, we highlight the importance of chest CT and the significance of radiological diagnostic aids in combating COVID-19 infections.

2.1. Importance of Chest CT in COVID-19 and its protocol

Chest CT plays a significant role in the automatic diagnosis and prognosis of lung disease detection. Chest CT is advised on the third day of the symptomatic patients of COVID-19. As per record, 56% of cases imaged during the initial two days with the above-mentioned COVID symptoms may show normal lung findings [27]. Moreover, chest CT is a well-chosen diagnosis technique for cases with negative RT-PCR reports but having mild to severe COVID-19 symptoms. Radiological findings in chest CT are beneficial for analyzing the seriousness of the confirmed cases. At the early stage of the disease, about 15-50% of cases have shown normal lung [22]. Considering the chest CT limitations and cost, it is not recommended as a regular screening test for COVID-19. Therefore, cases with false-negative RT-PCR and normal CT scan reports may result in isolation. Again, the chest CT findings of COVID-19 are incomprehensible to other viral infections like influenza, adenovirus, pertussis, swine flu, rhinovirus, etc. Hence, it may mislead the proper diagnosis of infected cases. Another restriction on performing CT is that it should be done after procuring every suspected individual and is time-consuming. Besides, chest CT is useful in suspected COVID cases with negative RT-PCR and normal chest X-ray (CXR) even though the person has

high suspicious index and suffers from mild-to-severe respiratory symptoms. The COVID-19 Reporting and Data System (CO-RADS) developed by the Dutch Radiological Society has become a reliable and convenient assessment scheme for radiologists to classify further COVID-19 based on the CT findings [28]. The CO-RADS shows a five-point assessment scale representing the suspicious COVID cases with pulmonary chest CT involvement, as shown in Table 1. Also, chest CT plays a crucial role in post-COVID diagnosis and prognosis periods where a patient recovering from COVID infection persists with impaired lung dysfunction. In such cases, chest CT puts forward a helping hand in differentiating the post-COVID infection (lung fibrosis) sequence from other lung diseases.

People of any age can get infected with this virus. So, keeping this in consideration, chest CT is always suggested to be done using a low radiation dose [29]. Using a low radiation dose helps reduce the radiation burden as the infected patient may need to undergo a sequence of CT follow-ups. With the continuity of COVID from the past two years worldwide, all medical personalities must know the usefulness of CT in proper management and detection of COVID suspected, infected, and recovered cases to contribute to this disease's diagnosis and prognosis care [30].

2.2. Significant radiological findings

Lung: The lung is a spongy, air-filled, pyramid-shaped organ connected to the trachea by the right and bronchi on the left [31]. Thin tissue layers called pleura cover the lung cavity. The right lung is shorter and broader than the left lung, which occupies a smaller area than the right lung. Both the lungs are separated into lobes by fissures. The right lung consists of superior, middle, and inferior lobes, while the left lung involves only superior and inferior lobes. Three regions can be mentioned to study the lung findings: the central region, placed on the interior boundary of the lung; the peripheral region, which includes the area between the central and the outline of the whole lung; and the basal region, the inferior lobes [32]. Figure 1 shows these anatomies.

Lung findings: Several studies have reported a wide variety of chest CT findings in the detection of COVID-19 [24][27][30]. However, the main CT findings include ground-glass opacity (GGO), consolidation, GGO with consolidation, and crazy paving. The GGO may be bilateral and multilobar with a peripheral, central, and basal distribution, mainly in the lower lobes and less visible in the upper and middle lobes [Figure

Table 2: CT Score severity [26]

Percentage (%) involvement of the five lobes	CT scale	Severity
0% lung involvement	0	None
Less than 5% lung involvement	1	Minimal
Above 5% upto 25% lung involvement	2	Mild
Above 25% upto 50% lung involvement	3	Moderate
Above 50% upto 75% lung involvement	4	Severe
Above 75% lung involvement.	5	Extensive

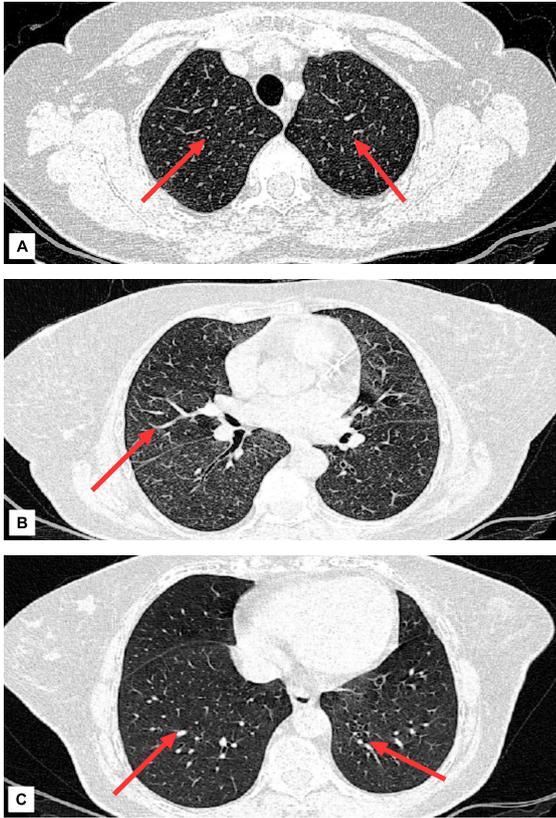


Figure 1: Axial HRCT images of lungs showing normal lobar anatomy of bilateral lungs. (A) shows bilateral upper lobe (B) shows the right middle lobe (C) shows bilateral lower lobe.

261 Other findings include a halo sign lesion which is rounded in
 262 structure. A halo sign is a rounded consolidation surrounded by
 263 ground-glass [Figure 2(e)] [37]. Moreover, vascular dilatation
 264 (widening of the vessels), fibrosis, traction bronchiectasis, sub-
 265 pleural bands, and architectural distortion are some less com-
 266 mon findings mainly seen during the later progression of the
 267 infection or the post-COVID phase.

268 2.3. Severity based on CT score:

269 COVID-19 infected patients have a variable infection rate
 270 where the severity of the infection ranges from mild with less
 271 than 10% of lung parenchyma involvement to severe infection
 272 comprising of white lung on CT [Figure 2(f)]. The severity
 273 of the disease correlates with the involvement of the lung in
 274 HRCT, which can be estimated visually. The prior concern re-
 275 garding the use of chest CT scan imaging was to appraise the
 276 spectrum of imaging findings and to recognize the different typ-
 277 ical, atypical, and indeterminate CT patterns for COVID-19 as
 278 mentioned in the previous subsection.

279 The pulmonary involvement of COVID-19 related anom-
 280 alies from thin-section CT images can be standardized and com-
 281 municated using the 25-point severity score or CT severity
 282 score for COVID-19. Without any additional tools, radiologists
 283 can use this scoring technique, which is fairly reproducible.
 284 Based on an approximation of the pulmonary affected areas,
 285 the COVID-19 lung alterations and involvement are scored us-
 286 ing the CT severity score index [38]-[40]. As per the inform-
 287 ation, it is evident that the combination of individual lobe scores,
 288 from negative to maximum lung involvement, 0 to 25, gives the
 289 total CT score (over 75% involvement of five lobes).

In our work, we have considered the 25-point severity score [26], where the severity of the disease is visually assessed by experienced radiologists. The involvement of all the five lobes is categorized on a five-point scale as given in Table 2. As per this scoring system for each of the 5 lobes (Right lung: Upper, Middle, Lower lobe; and Left lung: Upper, Lower lobe) is awarded a CT score from 0 – 5 depending on how much of the lobe is affected. Score 0 for 0% involvement of the lobe; score 1, corresponding < 5% involvement; score 2, for 5–25% involvement; score 3 for 26–50% involvement; score 4 to 51–75% involvement, and finally score 5 for > 75% involvements of the lobe. The total CT score is calculated as the sum of the individual scores of the five lobes, corresponding from 0 (no involvement) to 25 (maximum involvement) across the lung lobes.

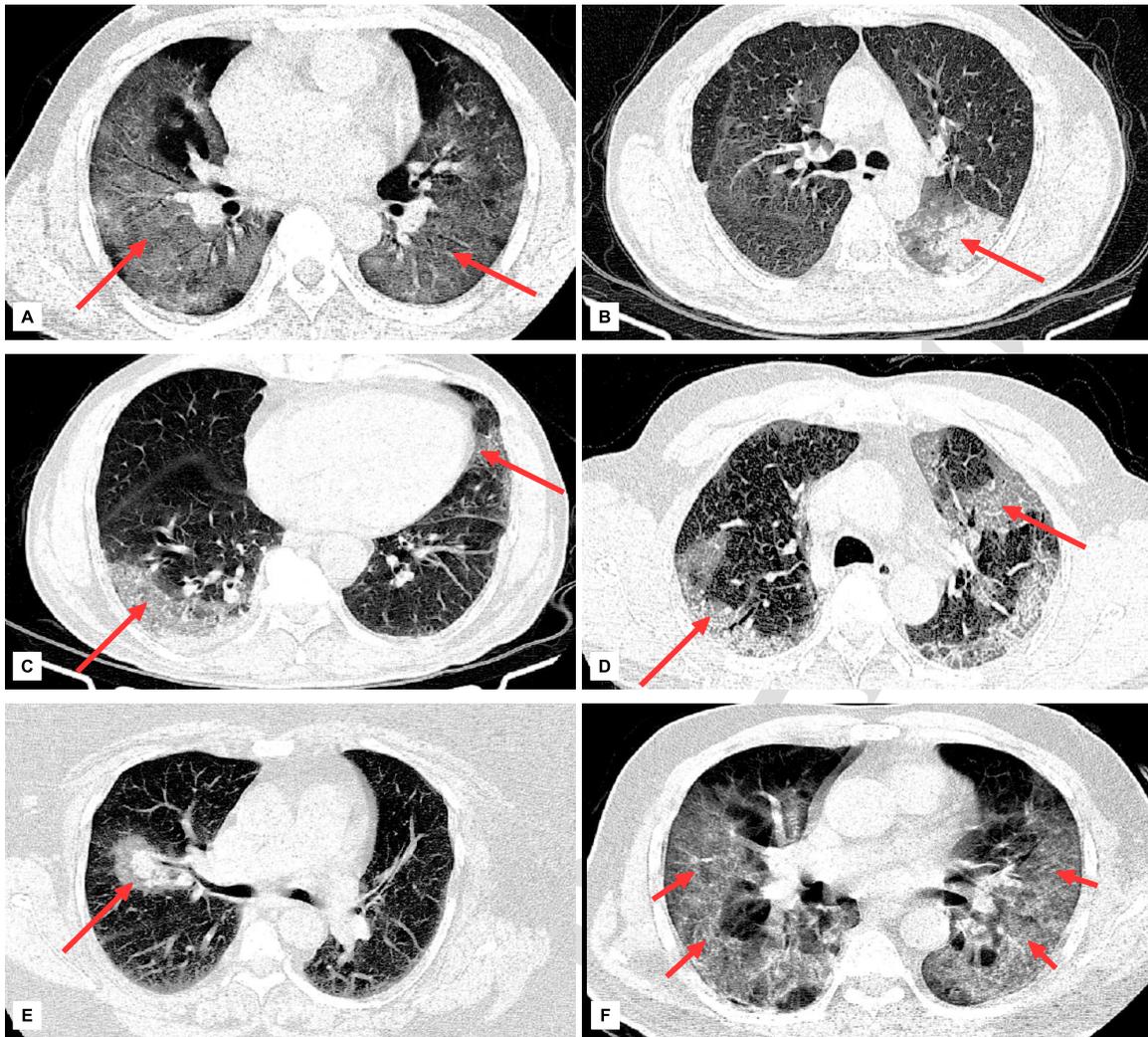


Figure 2: Radiological findings: (A) Axial HRCT of a COVID positive 40 years male showing GGO in bilateral lungs (arrows). (B) Axial HRCT of a COVID positive 45 years male shows an area of consolidation in the left lower lobe (arrow). (C) Axial HRCT of a COVID positive 70 years male showing GGO with consolidation in the right lower lobe (arrow) and a scattered area of GGO in the inferior lingular segment of the left lung (arrow). (D) Axial HRCT of a COVID positive 65 years male shows multifocal areas of GGO with interstitial thickening suggestive of crazy paving in the upper lobes of bilateral lungs (arrows). (E) Axial HRCT of a COVID positive 65 years female showing a nodular opacity with perilesional GGO suggestive of halo sign in right upper lobe (arrow). (F) "white lung" appearance of a COVID positive 62 years male with respiratory distress showing severe GGO(arrows) involving 75% of the bilateral lungs.

3. Materials and methods

The section details the data acquisition procedure, the criteria of the included and excluded data, imaging technique and interpretation, and the de-identification process.

3.1. Data acquisition procedure

The database used for this study was obtained from Marwari Hospitals, Guwahati, Assam, India recorded from May 2021 to February 2022. The patient's identity was kept confidential adhering to the ethical guidelines. The present study is purely retrospective and was carried out on 188 patient databases. The acquired dataset consists of imaging findings inclusive of:

- GGO with minimal to moderate hazy opacity.
- Consolidation pattern.

- GGO with consolidation.
- Crazy Paving patterns.
- GGO with crazy paving.
- Halo sign.

3.2. Criteria of the included and excluded data

All the cases used for this study were diagnosed with COVID-19, detected by the RT-PCR method tested in the hospital, and had undergone HRCT with the proper consent of doctors and laboratory technicians. The subjects used in this study had a few common clinical symptoms: loss of smell and appetite; a severe cold and cough; sore throat; anxiety; and breathing difficulties. Few patients have a history of COVID as per hospital records. Among different age groups of 188 patients,

Table 3: Age distribution of COVID and post-COVID cases

PARTICULARS	COVID				POST COVID			
	MALE		FEMALE		MALE		FEMALE	
	COUNT	% COUNT	COUNT	% COUNT	COUNT	% COUNT	COUNT	% COUNT
AGE UPTO 30 YR	8	05.5	11	07.5	2	04.8	5	11.9
AGE 30 YR TO 60 YR	40	27.4	34	23.3	7	16.7	8	19.0
AGE 60 YR TO 90 YR	35	24.0	17	11.6	8	19.0	12	28.6
AGE ABOVE 90 YR	1	00.7	0	00.0	0	00.0	0	00.0
TOTAL NO. OF CASES	84	57.5	62	42.5	17	40.5	25	59.5

Age distribution chart

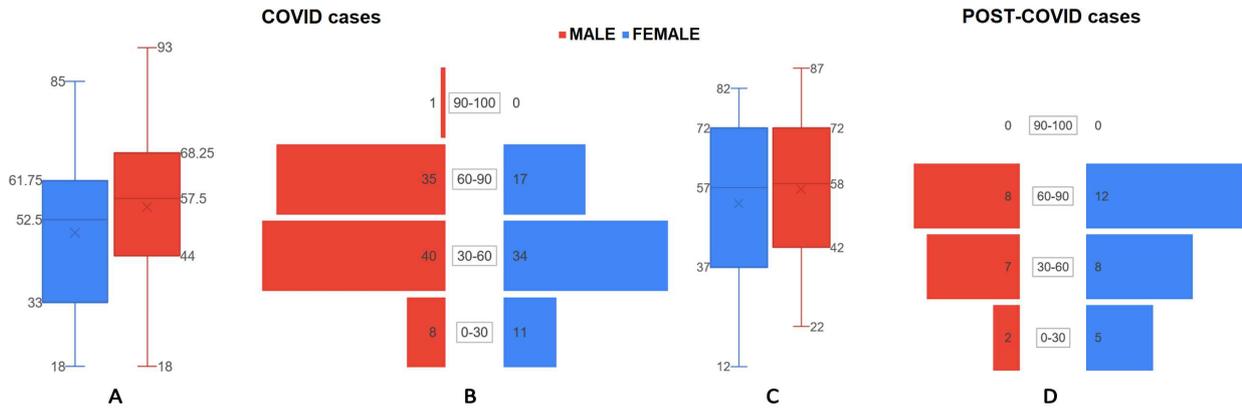


Figure 3: Chart showing the inception of COVID-19 with respect to age distribution and gender: (A) Box plot for male and female COVID-infected cases; (B) Butterfly plot showing patient distribution by age category; (C) Corresponding box plot for post-COVID cases; (D) Corresponding butterfly plot for post-COVID cases.

inclusive of both male and female sex, 144 patients are RT-PCR positive, with a true negative of two patients. The remaining patients are post-COVID cases. This study excludes pregnant women and patients on ventilator support.

3.3. Imaging technique and interpretation

All HRCTs have been carried out using SOMATOM go. Standard CT scan protocols have been applied with a topogram of 512 cm with 120 kV and 35 mA ratings. The lung images have been obtained in an axial window and reconstructed into 1.5mm thin slices. Finally, all the HRCT images acquired are saved in the Digital Imaging and Communications in Medicine (DICOM) format, which further helps in the re-evaluation process. Two radiologists with 33 and 10 years of experience reviewed the findings using the DICOM software. All the images have been viewed in both lung and mediastinal windows. We have evaluated our study using a few parameters, considering the HRCT findings as mentioned above. Laterality of lung involvement, lobar involvement and region of coverage of lungs, and CT Score Severity Grading are the parameters used to diagnose the extent of infection taking place in the lung found in the COVID samples.

3.4. De-identification process

Keeping in view the privacy of the patients, we have de-identified all the CT studies using the DICOM software. Every name, patient ID, and centre-related information has been

removed for further statistical characteristics calculation of the dataset.

4. Data statistics and interpretation results

CT scan images are extensively used as diagnostic aids for the treatment of COVID-19 infected patients. From the collected images, we have obtained the annotation and data classification based on the statistics for the mentioned retrospective study. This section highlights the statistics of the data, truly based on retrospective study to incorporate its applicability for the development of ML and DL algorithms, which are to be used as AI tools to create predictive models based on pure database patterns acquired from CT scanners. At the end of this section, a discussion based on the statistical analysis and a few of the limitations of the available data is brought to the fore.

The allocation based on age group and gender for both COVID and post-COVID is shown in Table 3 and Figure 3. Among the COVID positive cases, 57.5% are male, and 42.5% are female patients. The distribution of patients is shown in the box plot of Figure 3(A) separately for males and females. As per age groups, referring to the plot shown in Figure 3(B) the highest number of sufferers were 30–60 years old, with 27.4% male and 23.3% female. The next group is 60–90 years old, with 24% male and 11% female. The minor sufferer group belongs to the age group of 30 years (05.5% male and 07.5% female). We also have one patient (male) belonging to the age group of 90. The CT scan of this case is shown in Figure 4.

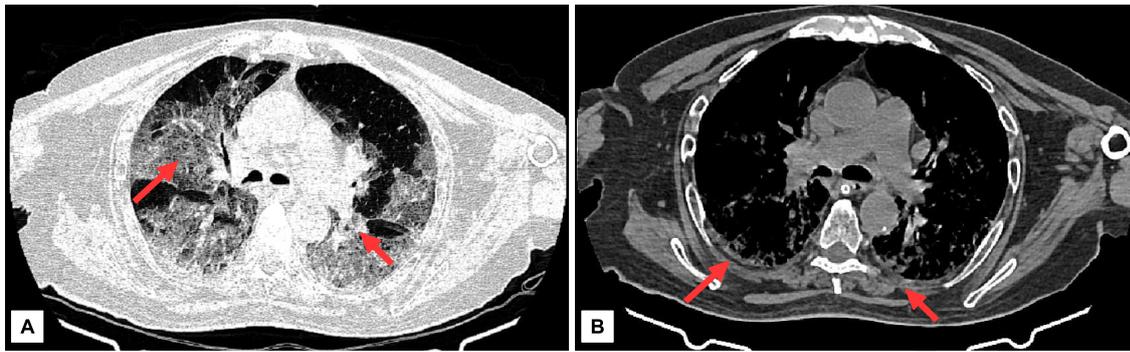


Figure 4: Axial HRCT of a 93-year-old COVID positive man shows (A) extensive GGO in the bilateral lung with small areas of consolidation (arrow). (B) The mediastinal window shows minimal bilateral pleural effusion in the same patient (arrow). The HRCT was obtained 12 days after the infection with a low oxygen level and high fever symptoms.

Table 4: Laterality of lung involvement of COVID and post-COVID cases

PARTICULARS	POSITIVE		NEGATIVE	
	COUNT	% COUNT	COUNT	% COUNT
Bilateral lung involvement of COVID and post-COVID	82	43.6	106	56.4
Laterality of RIGHT lung involvement of COVID and post-COVID	92	48.9	96	51.1
Laterality of LEFT lung involvement of COVID and post-COVID	95	50.5	93	49.5

Table 5: Lobar involvement of lungs of COVID and post-COVID

Distribution of lobes of lung of COVID and post-COVID		NO	UPPER	UPPER	LOWER	UPPER	MIDDLE	LOWER
			MIDDLE	LOWER	MIDDLE	MIDDLE	LOWER	
RIGHT LUNG	COUNT	96.0	54.0	04.0	06.0	05.0	07.0	16.0
	% COUNT	51.1	28.7	02.1	03.2	02.7	03.7	08.5
LEFT LUNG	COUNT	93.0	00.0	65.0	00.0	05.0	00.0	25.0
	% COUNT	49.5	00.0	34.6	00.0	02.7	00.0	13.3

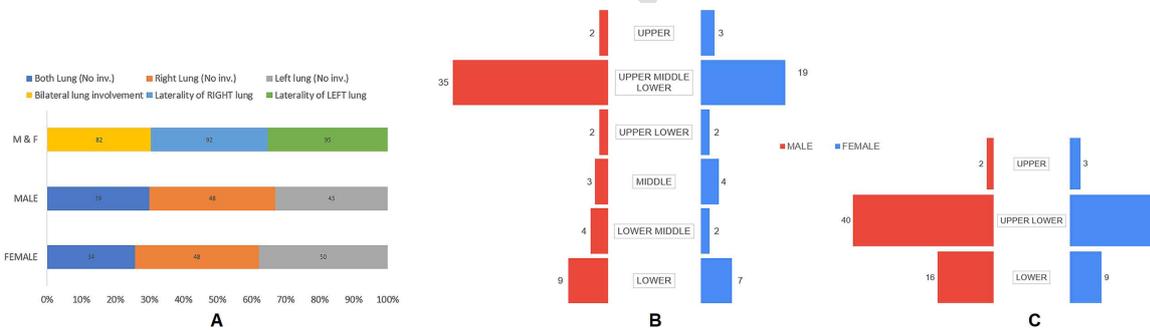


Figure 5: Chart showing HRCT findings for lung involvement of COVID and post-COVID cases with respect to gender: (A) A bar diagram depicting the patient count for lung involvement in both the right and left lungs; (B) a butterfly plot depicting lobar involvement of the right lung in both male and female patients; and (c) a corresponding butterfly plot depicting lobar involvement of the left lung.

Table 6: Region of coverage of COVID and post-COVID cases

REGION OF COVERAGE	POSITIVE		NEGATIVE	
	COUNT	% COUNT	COUNT	% COUNT
CENTRAL	30	16.0	158	84.0
PERIPHERAL	92	48.9	96	51.1
BASAL	92	48.9	96	51.1

383 Similarly, in 42 post-COVID cases, 40.5% belonged to male 385 3(C) shows the corresponding distribution. As seen from the
 384 patients and 59.5% to female patients. The box plot in Figure 386 butterfly plot Figure 3(D), the highest post-COVID cases be-

Table 7: HRCT findings

HRCT findings in COVID and post-COVID case	MALE		FEMALE	
	COUNT	% COUNT	COUNT	% COUNT
Minimal Ground Glass Opacities	07.0	06.9	01.0	01.1
Ground Glass Opacities	40.0	39.6	34.0	39.1
Consolidation	25.0	24.8	16.0	18.4
Ground Glass Opacities with consolidation	15.0	14.9	06.0	06.9
Crazy Paving	06.0	05.9	01.0	01.1
Ground Glass Opacities with crazy paving	05.0	05.0	00.0	00.0
Halo sign	02.0	02.0	03.0	03.4
Nil parameter findings cases	40.0	39.6	44.0	50.6
One parameter findings cases	30.0	29.7	28.0	32.2
Two parameter findings cases	25.0	24.8	12.0	13.8
Three parameter findings cases	04.0	04.0	03.0	03.4
Four parameter findings cases	02.0	02.0	00.0	00.0
Five parameter findings cases	00.0	00.0	00.0	00.0
Six parameter findings cases	00.0	00.0	00.0	00.0
Seven parameter findings cases	00.0	00.0	00.0	00.0

long to the age group of 60-90 years, with 19% male and 28%⁴²⁴ female, followed by 30-60 years, 16.7% male and 19% female,⁴²⁵ and the rest belonging to the age group of 30 years, with 4.8%⁴²⁶ male and 11.9% female.⁴²⁷

Hence, from the available database, we may conclude that⁴²⁸ the highest number of COVID-infected patients belong to the⁴²⁹ age group of 30–60 years, with a high prevalence among male⁴³⁰ patients. The post-COVID cases show a high rate in the age⁴³¹ group of 60–90 years old, with more female cases.⁴³²

HRCT of the COVID symptomatic patients has been per-⁴³³formed between the 4th to 14th days of infection for the 146⁴³⁴ COVID cases. The 42 post-COVID cases underwent the scan⁴³⁵ due to breathing difficulties, low oxygen saturation (SpO_2),⁴³⁶ palpitation, weakness with body ache after testing RT-PCR neg-⁴³⁷ative post-infection. Out of 188 cases, it was found that 43.6%⁴³⁸ of them had bilateral lung involvement, while 48.9% had in-⁴³⁹volvement in the right lung and 50.5% had involvement in the⁴⁴⁰ left lung; the details are given in Table 4 and Figure 5.⁴⁴¹

As observed from Table 5, and Figure 5 (c) both upper and⁴⁴² lower lobes (65%, with 50% to 25% male female ratio) of the⁴⁴³ left lung were the most commonly involved, followed by the⁴⁴⁴ right upper, middle, and lower lobes (54%, with 35% to 19%⁴⁴⁵ male female ratio), referring to Figure 5 (b). Next in the count⁴⁴⁶ is the left lower lobe (25%, 16 : 9) and the right lower lobe⁴⁴⁷ (16%, 9 : 7). Considering the regions as shown in Table 6,⁴⁴⁸ 48.9% of the cases had findings in both peripheral and basal⁴⁴⁹ regions of both the right and left lung, followed by 16% in the⁴⁵⁰ central region.⁴⁵¹

In this study, it is noticeable that 44 female cases (50.6%)⁴⁴² had a normal HRCT scan with the absence of all the parameters⁴⁴³ taken into consideration, followed by 40 male cases (39.6%).⁴⁴⁴ The most common finding is the presence of multifocal, bilat-⁴⁴⁵eral, peripheral GGO, which is significantly found in 40 male⁴⁴⁶ patients (39.6%) and 34 female patients (39.1%), followed by⁴⁴⁷ 7 (06.9%) male and 1 (01.1%) female with minimal opacities.⁴⁴⁸ The HRCT performed on the cases mentioned, with minimal⁴⁴⁹ to severe GGO was done approximately between the 4th to 12th⁴⁵⁰

days from being infected with the virus. The next common find-
ing that was seen in 25 male (24.8%) and 16 female (18.4%)
patients is the presence of consolidation. GGO with consoli-
dation was present in 15 male (14.9%) and 6 female (06.9%)
patients. A crazy-paving pattern was found in 6 male (05.9%)
patients and 1 female (01.1%). GGO with the presence of a
crazy-paving pattern was found only in 5 male patients (05.0%).
A halo sign has been seen in 2 (02.0%) male and 3 (03.4%) fe-
male patients. None of the cases had all the seven parameter
findings. Maximum cases had only one parameter finding (ei-
ther the presence of GGO or consolidation or GGO with con-
solidation or GGO with crazy paving patterns), followed by a
minimum of four to three-parameter findings [Table 7]

The CT score is done visually based on the severity of the
disease [Table 8 and Figure 6]. A total of 85 patients with

Table 8: CT Score Severity Grading

CT score	Patient Count
0	85
1 - 5	40
6 - 10	37
11 - 15	19
16 - 20	6
21 - 24	1

COVID and post-COVID infection have no lung involvement.
The highest CT severity is assigned to 1 patient with a CT value
ranging between 20-25 with bilateral lung involvement, involv-
ing all the lobes [shown in Figure 2 (a)]. 40 patients had CT
scores between 1 and 5, where the cases showed bilateral lung
involvement in a few cases and involvement of either one lung
in a few cases with the presence of minimal to moderate periph-
eral GGO, GGO with consolidation, and halo sign [refer Figure
7 for an example]. 37 patients showed CT severity ranging
between 6-10, showing patches of consolidation, consolidation
with the presence of GGO, and a few crazy paving patterns in
less than 10 patients, for example as shown in Figure 8. 19 pa-

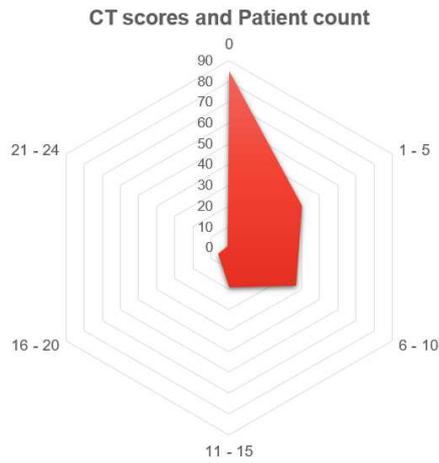


Figure 6: Scatter plot shows the CT Score Severity Grading with respect to CT score division and corresponding patient count.

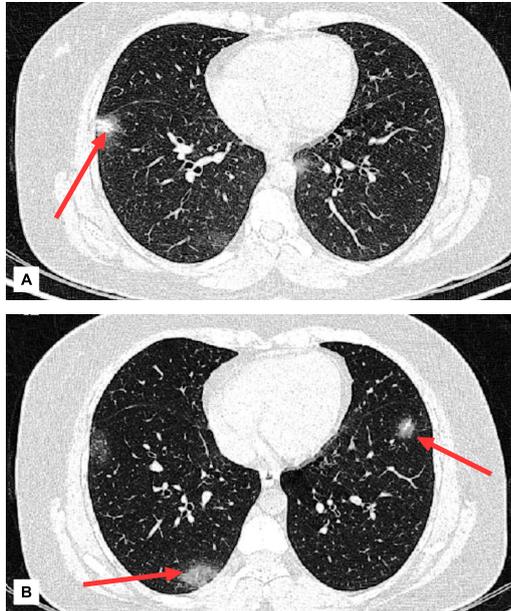


Figure 7: COVID positive 41 years female, scan done on 7th day from infection: (A) axial HRCT showing the occurrence of a nodular opacity with surrounding GGO (arrow), and (B) scattered areas of GGO's in both lower lobes (arrows).

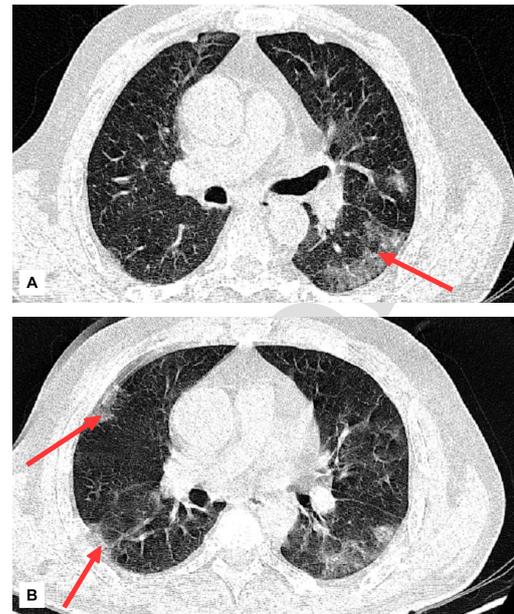


Figure 8: COVID positive 65 years male, scan done on 9th day from infection: (A) axial HRCT showing patchy GGO with crazy-paving pattern in left lower lobe (arrow), (B) GGO with consolidation in the right middle and lower lobes (arrows).



Figure 9: COVID positive 52 years female on the 9th day of infection: axial HRCT shows the occurrence of GGO in bilateral, peripheral areas in the upper lobe and patchy areas of consolidation in left upper and lower lobes (arrows).

451 tients had CT severity between 11-15, involving both lungs with
452 moderate GGO and consolidation, as shown in Figure 9 as an
453 example, followed by 6 patients whose CT severity ranged be-
454 tween 16-20 showing bilateral peripheral GGO with interstitial
455 thickening, GGO's with consolidation, and bilateral pleural ef-
456 fusion [refer Figure 10 for an example]. Several chest HRCTs
457 reported in our study with symptoms of chest pain, pounding
458 heartbeat, shortness of breath and fatigue were found to have
459 opacities parallel to the pleura [as an example refer Figure 11],
460 fibrosis along with bronchiectasis as shown in Figure 12, in-
461 terstitial thickening and subpleural band [Figure 13 for refer-
462 ance]. The evidence of the prior information on HRCT con-

463 firms the presence of the virus in earlier months, along with
464 some pre-existing diseases. However, a few cases with post-
465 COVID symptoms did not show any findings on HRCT done 2
466 months prior to the infection. These few cases may have other
467 pre-existing diseases due to which they might experience the
468 above-stated post-COVID symptoms.

5. Discussions

As per the statistics of North-East India collected during the pandemic stage, thousands of new COVID-19 cases were registered on a daily basis. The gold standard for COVID identification, the RT-PCR test, is efficient, but it shows true negatives and therefore fails to be 100% accurate. Moreover, the case becomes more challenging for asymptomatic patients. As a secondary measure, CT-scans are advised, but generally patients

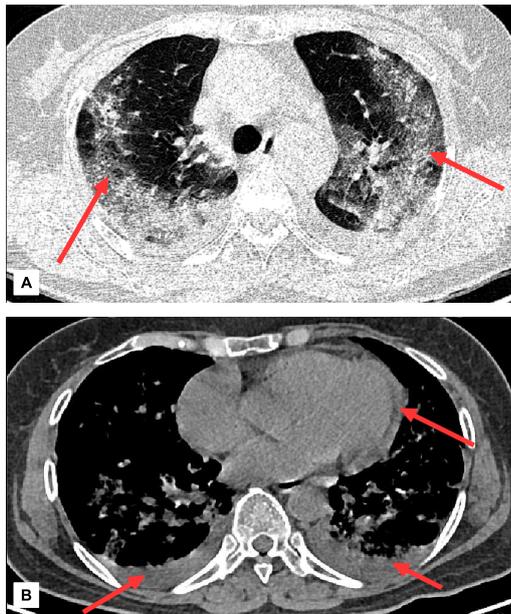


Figure 10: COVID positive 54 years female on the 13th day of infection: (A) axial HRCT shows the occurrence of GGO and consolidation in bilateral upper lobes in the peripheral region (arrow). (B) The mediastinal window shows moderate pleural effusion bilaterally and mild pericardial effusion (arrows).

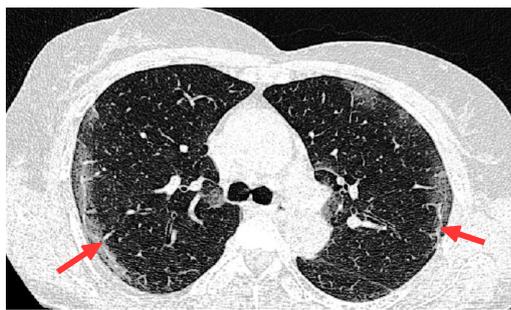


Figure 11: 56 years old female with a history of COVID shows thin linear opacities parallel to the pleura in bilateral upper lobes, representing a subpleural band and adjacent GGO (arrows). She tested positive for COVID two months ago and had persistent shortness of breath. The HRCT was done when she tested negative two months post-COVID.



Figure 12: 58 years old man was discharged from the hospital after testing negative with the help of supportive therapy. He had symptoms of chest pain, fever, and breathing difficulties. HRCT done after one week shows fibrosis and bronchiectasis in the right lower lobe (arrow) and a small fibrotic opacity in the left upper lobe (arrow).

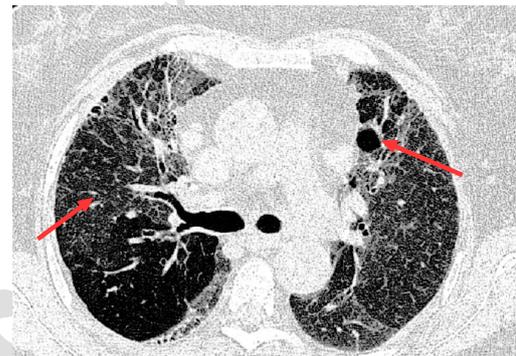


Figure 13: 58 years female with symptoms of low oxygen level, breathing difficulties and cough showed evidence of fibrosis, interstitial thickening, subpleural cyst and bronchiectasis in bilateral upper lobes and interstitial thickening right lower lobe (arrow). She was admitted to the hospital due to a rapid drop in oxygen levels after two weeks of testing RT-PCR negative.

show less interest unless they are suffering from symptoms related to lung disorders. Also, the lung anomalies start at a later stage of COVID inception and when not treated in time, lead to total lung dysfunction and finally death. During the second wave of COVID, the post-COVID anomalies have increased the number of death counts.

As already mentioned, we have collected CT-scans of a total of 188 COVID positive patients. However, as far as the accuracy of the corresponding RT-PCR report is concerned, we have found two HRCT cases that show that they belong to COVID positive patients but were actually evaluated as RT-PCR negative. The CT scan of these two cases is shown in Figure 14 which clearly signifies the presence of COVID. The database statistics showed more infections in the age group of 30–60, with a higher rate of male patients. This demography may vary

depending on location and populations, but there is a probability match of the statistics with those recorded in other parts of India [7][41]. Next, as per the HRCT findings on lung involvement, the pattern is similar in both genders with more effect on the upper-middle-lower lobes in the right and upper-lower in the left lung, respectively. As per the study findings, out of a total of 188 cases, 40 (39.6%) male and 44 (50.6%) female patients showed normal HRCT with zero findings with respect to GGO and consolidation. Thus, CT-scans can't be used as a standalone measure for screening and RT-PCR will continue to be the gold standard for mass identification. But the point to be noted is that for the RT-PCR positive cases or post-COVID, CT-scans are mandatory. These studies were also reported in previous literature [11][24][41][42], but the main objective of this work is to understand the HRCT findings useful to apply image processing tools and for automated AI algorithms to classify and grade COVID-19 severity from the samples.

Considering the versatility of the disease, the detection process based on radiological findings has many shortcomings despite its wide applications in diagnostic centers. Correspondingly, researchers related to the medical and computer fields use AI tools such as ML and DL models for analyzing these ra-

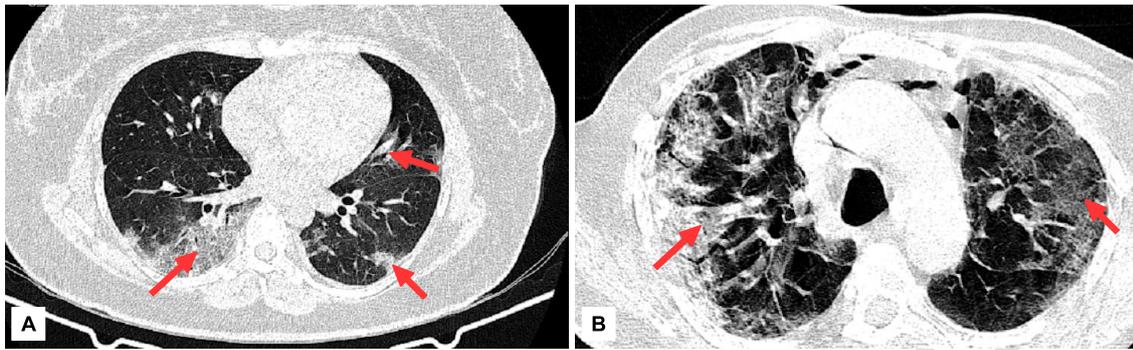


Figure 14: True negative in RT-PCR and corresponding CT scans; (A) Axial HRCT of a 36-year-old female COVID negative patient showing crazy paving and consolidation in the right peripheral and basal area (arrow), and patchy areas of consolidation are seen in the left lower lobe lingular segment (arrow). This patient had symptoms of high fever, headache, and sore throat, which resulted in RT-PCR being negative during testing. The patient underwent HRCT due to non-recovery from the symptoms, which resulted in the findings mentioned above suggestive of COVID positive. (B) axial HRCT of a 62-year-old man with symptoms of COVID infection resulting in breathing difficulties. The patient tested RT-PCR negative, but HRCT showed areas of consolidation in the right lung and GGO's in the left lung in peripheral locations (arrows).

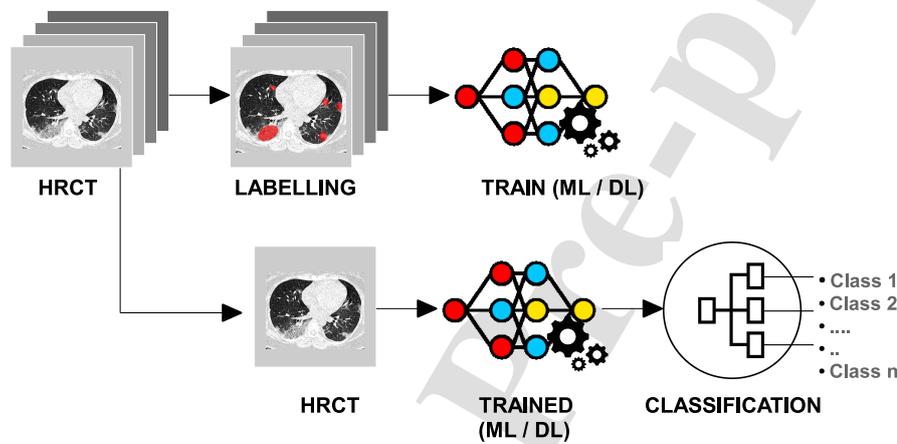


Figure 15: Block diagram of ML-DL model for automated classification of COVID-19

514 biological findings. There were many systematic reviews which 535
 515 showed that ML and DL algorithms can be used for various dis- 536
 516 ease detection and diagnosis processes [21],[43]-[45]. These 537
 517 algorithms have proven themselves to be efficient in the detec- 538
 518 tion and classification of various diseases from different types 539
 519 of databases [46][47]. Using different DL algorithms, stud- 540
 520 ies have embarked on the identification and differential diag- 541
 521 nosis of COVID-19. The DL-based architecture in the field 542
 522 of COVID-19 based on radiological findings helps to reduce 543
 523 false-positive and negative errors in the analysis of the disease, 544
 524 providing a fast and safe diagnosis system for patients. Figure 545
 525 15 provides an overview of a predictive model for the detec- 546
 526 tion and diagnosis of COVID-19 using radiological modalities. 547
 527 Various studies reviewed that DL based detection for COVID 548
 528 diagnosis is classified into two categories: pre-trained models 549
 529 with deep transfer learning (DTL) and customized DL [20]. 550
 530 Pre-trained models use trained data in areas that have simi- 551
 531 larity with the content of the application. Pre-trained models 552
 532 with DTL provide the facility to accelerate convergence with 553
 533 network generalization other than the general training models, 554
 534 which requires more computation time. On the other hand, cus- 555

tom based DL techniques have specific applications and deal
 with feasible architecture development, which gives accurate
 performance due to their consistency. The custom networks
 use specific DL algorithms or hybridization of DL techniques
 with other fields of application like AI, such as data mining,
 ML, etc. [48]. Various popular architectures such as AlexNet,
 GoogleNet, MobileNetv2, Inception, InceptionV3, Xception,
 VGG-16, VGG-19, ResNet, ResNet18, ResNet50, ResNet101,
 Inception ResNetV2, DenseNet201, XceptionNet, CSHNet,
 FGCNet are used in both of these models [49]-[60]. Responses
 reported on these architectures by state-of-the art methods are
 reproduced in Figure 16 where pre-trained model comparison
 is based on its sensitivity of detection, where-as the customized
 methods are categorized as per its accuracy of detection [49]-
 [70].

As per the literature, although efficient algorithms have been
 proposed in this area, there are certain limitations to these stud-
 ies. As reported in our previous publication of COVID-19 re-
 view [71] there are two fold shortcomings, which includes, (i)
 improper database and (ii) limited database. Due to the noisy,
 incomplete, and unmarked data, issues like redundancy, spar-

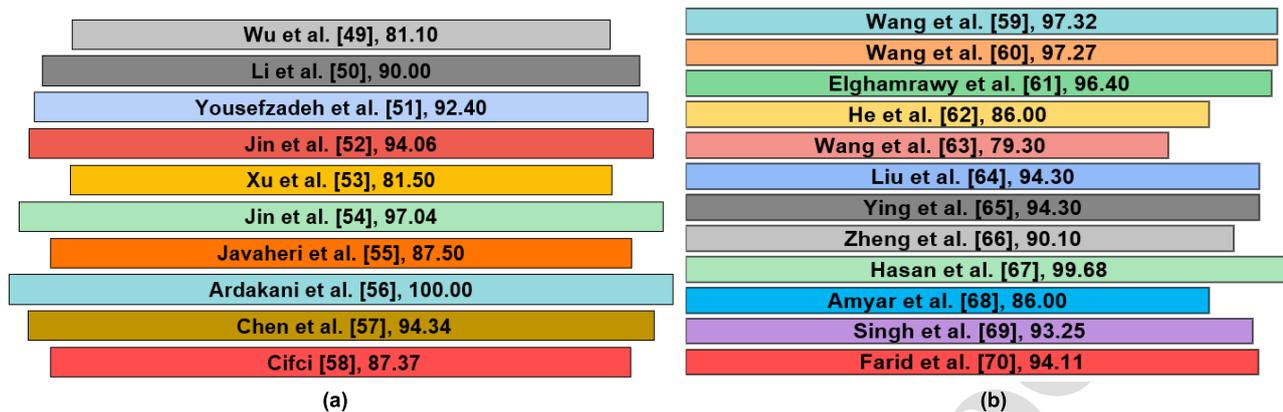


Figure 16: Comparison of state-of-the-art DL algorithms for detection of COVID-19 using HRCT images based on, (a) Sensitivity of Pre-trained models, and (b) Accuracy of Customized models.

sity, and non-availability of values are encountered during the training of classification modules. Again, the unavailability of accessible implies the application of limited local databases, which are very minimal compared to the actual patient data. This leads to underfit or overfit problems while applying DL. Apart from this, with the unavailability of standard databases, it becomes hard to calculate the proper efficiency. Hence, considering these factors and the modality under consideration, we would like to continue our work in the direction of automation of CT-scan grading using ML and DL tools, to increase the sensitivity and specificity of detection.

Limitations: There are a few limitations encountered during this work. Since the study is retrospective, and all the cases diagnosed are COVID-infected, it cannot be used to evaluate the exact sensitivity and specificity of COVID-19 HRCTs. RT-PCR and HRCT cannot be compared for COVID detection. For instance, all the post-COVID subjects considered in this study are not RT-PCR positive. These patients underwent HRCT due to post-COVID symptoms as mentioned in hospital records and clinical results. Again, the slices and lobes reviewed during the study depended on regions with distinctive COVID findings; some subjects had pre-existing lung disorders, resulting in subjective findings, which put hindrances during the study. A few low-quality HRCT scans showed mild motion artefacts that may be due to shortness of breath (dyspnea) experienced by the patients. Apart from these, there are certain other associated infections/diseases which worsen the COVID-19 patient's condition. Some of them are bacterial or fungal co-infections, with mucormycosis being the most common. Mucormycosis is a common side effect of corticosteroids that are prescribed for severe COVID-19. At mucormycosis the patient's immune system weakens and blood sugar level increases, and it becomes life threatening for the diabetic patients [72]. As per the International Diabetes Federation, India being the hub of diabetes, the appearance of mucormycosis is common for most of the databases under consideration. But this information was not available in the current CT scan reports of Marwary Hospitals and therefore not considered in this work.

To summarize, we may say that, with considerable evidence,

chest HRCT proves itself to be an essential module for diagnosis, prognosis, and follow-up for individuals with COVID infection. All the patients confirmed with COVID-19 have the same imaging patterns, which may be helpful for the radiologist for early detection and estimating the severity of the disease. We have performed this study as a familiarization step to HRCT findings of COVID and post-COVID so that it may help the research community to explore possibilities for analysis or prognosis of this challenging disease of the current epoch.

6. Conclusion

The HRCT findings of the chest play an essential role for radiologists in adequately diagnosing the COVID-19 infection. Mass screening by chest HRCT is a must, along with RT-PCR tests, to rule out better disease management. The presence of common imaging patterns of bilateral, peripheral, multifocal GGO, consolidation, and GGO with consolidation, in the initial phase of the infected cases, may help in the early detection of the disease. Asymmetrical and bilateral distribution of opacities of lungs is considered a benchmark for radiologists to detect COVID-19 pneumonia.

The current retrospective study analyses the clinical characteristics and outcomes of 188 patients with COVID-19 and post-COVID infections. It identifies six HRCT findings based on age, gender, laterality of lung involvement, lobe involvement of lungs, region of coverage and lastly, CT severity score for proper assessment of the severity of the disease progression. The strength of this study is to assist researchers in exploring the radiological imaging patterns or findings of COVID-19 to develop knowledge-based systems using AI tools for detecting and diagnosing COVID-19. At the same time, it will be helpful for radiologists and medical institutes as an additional tool for the early detection of the disease. In the near future, we will analyze these findings for implementing AI tools to propose a novel automated algorithm for detecting and grading COVID-19.

Although HRCT outperforms the RT-PCR tests as a mass screening tool, for the time being, RT-PCR is regarded as the

primary gold standard test as being cost-effective and able to detect subclinical cases due to the rising number of infections in the community level transmission.

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Highlights:

- The role of chest CT in diagnostic decision making.
- The typical initial CT findings in COVID-19.
- Chest CT imaging importance in diagnostic work-up and management in patients with COVID-19.
- Retrospective study of collected 188 cases of HRCT images.
- Application of deep learning in the field of COVID-19 radiological imaging

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**Automated Diagnosis of COVID-19 Using Radiological Modalities and Artificial Intelligence Functionalities:
A Retrospective study based on chest HRCT database**

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Declaration of interests

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

The authors declare the following financial interests/personal relationships which may be considered as potential competing interests:

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WITHOUT ANY PREJUDICE

TO WHOM IT MAY CONCERN

This is to certify that Ms. Upasana Bhattacharjya, research scholar from Gauhati University, Department of Electronics and Communication Engineering has taken data from the hospital which contains HRCT Images and corresponding RT-PCR test information regarding her ongoing research on biomedical images related to automatic diagnosis of COVID-19.

Please note, we have no objection in using these data for research purpose only by adhering to the ethical guidelines and also these data do not contain any biological sample.

Yours faithfully,
For, Marwari Hospitals,
(A unit of Shree Marwari Databya Aushadhalaya)

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