

# Patterns of thromboembolic pulmonary vascular disease in COVID-19

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## Abstract

SARS-CoV-2 (COVID-19) is associated with increased thrombosis. Here, we demonstrate patterns of pulmonary vascular disease in COVID-19 including classical acute pulmonary embolism and subsegmental perfusion defects in the absence of acute pulmonary embolism suggestive of microvascular thrombosis.

## Keywords

coronavirus, CT-LSIM, iodine subtraction mapping, pulmonary embolism, microvascular thrombosis

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## Introduction

Severe acute respiratory syndrome (SARS) caused by the coronavirus SARS-CoV-2 (COVID-19) has a high mortality due primarily to respiratory failure. Recent studies have highlighted increased thrombosis in COVID-19.<sup>1</sup> Extensive microvascular thrombosis has been noted at post-mortem and high rates of pulmonary embolism (PE) diagnosed using computed tomography pulmonary angiography (CTPA),<sup>2</sup> alongside lung parenchymal changes.<sup>3</sup> There has been increasing interest in the patterns of pulmonary vascular involvement due to COVID-19 and concern that perfusion abnormalities may represent in-situ thrombosis that may not be appreciated on standard CTPA. The British Thoracic Imaging Society Guidelines recommend unenhanced pulmonary angiography and CTPA. This protocol facilitates lung subtraction iodine mapping (CT-LSIM) for lung perfusion, a clinically sensitive tool in PE. We report the first CT-LSIM images in COVID-19.

## Materials and methods

At our institution, 10 patients (mean age (SD) 70 (16), 40% female) with COVID-19, confirmed on reverse transcription polymerase chain reaction (RT-PCR), underwent CTPA and CT-LSIM for suspected acute PE based on clinical

assessment and elevated d-dimer levels (Table 1). Analysis of CT images was approved by our institution review board.

## Results

Three patients had confirmed PE on CTPA and CT-LSIM (one case is shown in Fig. 1 (1a, 1b, 1c)). Another patient had perfusion defects on CT-LSIM without visible PE where CT-LSIM showed subsegmental perfusion defects without visible PE (Fig. 1 (2a, 2b, 2c)). Six patients did not have perfusion defects on CT-LSIM.

## Discussion

Distinct from classical thromboembolic PE, a high proportion of in situ pulmonary arterial thrombosis exists in COVID-19, and the pathophysiology is not fully understood.<sup>4</sup> Here, we demonstrate patterns of pulmonary vascular disease in COVID-19 including (i) classical acute PE with central clot associated with lung infarction and (ii)

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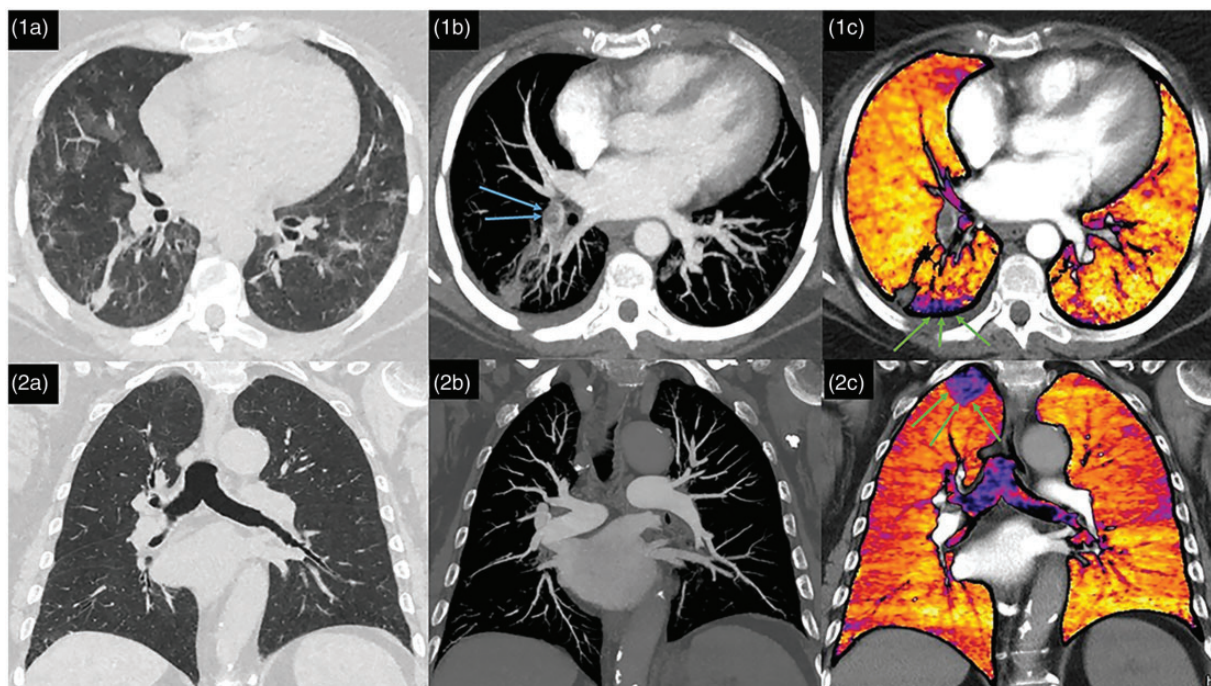


**Table 1.** Patients' characteristics.

Characteristic	Total (n = 10)	No pulmonary vascular changes CTPA/CT-LSIM (n = 6)	Pulmonary vascular changes CTPA/CT-LSIM (n = 4)
Age, years	70 (16)	73 (17)	68 (18)
Female, n	4	3	1
Race or ethnic group, n			
White	8	4	4
Black	1	1	0
Other ethnicity	1	1	0
Comorbidities, n			
Obesity	2	1	1
Smoker	3	2	1
Immunosuppression	2	1	1
Malignancy	1	0	1
Chronic obstructive pulmonary disease	1	1	0
Asthma	2	1	1
Ischaemic heart disease	1	0	1
Hypertension	5	2	3
Diabetes mellitus	2	1	1
Chronic kidney disease	1	0	1
Symptoms and signs around time for CTPA, n			
Tachycardia	4	2	2
Chest pain	2	1	1
Hypoxia	8	4	4
Intubation	2	1	1
Length of hospital stay, days	23 (13)	17 (16)	27 (15)
Critical care admission, n	5	2	3
Peak d-dimer 30 days prior to CTPA, ng/ml <sup>a</sup>	1637 (1075–18,902)	1160 (870–15,240)	8130 (2339–23,317)
Peak c-reactive protein 7 days prior to CTPA, mg/L	192 (127)	204 (84)	181 (170)
Peak Ferritin 30 days prior to CTPA, µg/L <sup>a</sup>	1036 (347–1928)	1089 (218–1360)	982 (342–1801)

Data are presented as mean (SD) or numbers. CTPA: computed tomography pulmonary angiography; CT-LSIM: computed tomography lung subtraction iodine mapping.

<sup>a</sup>Median (interquartile range).



**Figure 1.** Patterns of parenchymal, vascular, and perfusion abnormalities in COVID-19. Patient 1: (a) Peripheral wedge-shaped abnormality in the right lower lobe associated with a segmental filling defect (blue arrows) on CTPA (b) corresponding to a perfusion defect (green arrows) on CT-LSIM (c). Patient 2: (a) Absence of lung parenchymal involvement and normal pulmonary vasculature on CTPA (b) with a perfusion defect in the right upper lobe on CT-LSIM (c, green arrows).

subsegmental perfusion defects in the absence of acute PE which is perhaps suggestive of microvascular thrombosis.

CT-LSIM is potentially widely available for the assessment of lung perfusion in COVID-19. Further studies to understand the pathophysiology of pulmonary thrombotic disease in COVID-19 are required.

#### Author contributions

YS: analysed the data and wrote the manuscript; SR and AJS analysed the CT scans and collected the data; VP; collected the clinical data; DGK and JMW reviewed the images and final manuscript.

#### Conflict of interest

The author(s) declare that there is no conflict of interest.

#### Ethical approval

Imaging analysis was approved by our institutional review board.

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#### Guarantor

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