



BS NGUYỄN TẤN DŨNG

1



Contents

- 1. SARS COV2 pathologies.
- 2. The S protein and its mutations in
 - different genetic variants.
- 3. CT in SARS COV2.
- 4. Presented 2 clinical case.
- 5. 2 SARS COV2 drugs for treatment.





https://jbiomedsci.biomedcentral.com/articles/10.1186/s12929-020-00703-5#Fig2

Major severe COVID-19 pathologies and infection routes

5



https://jbiomedsci.biomedcentral.com/articles/10.1186/s12929-020-00703-5/figures/2



Published online 2021 Apr 10. doi: 10.7150/ijbs.59137



The S protein and its mutations in different genetic variants.

Table 1

Summary of mutations on S proteins in the five SARS-CoV-2 variants

Amino acid position in S protein	Wuhan-Hu-1	D614G	Cluster 5	VOC 202012/01	501Y.V2	P.1	Note
			Residues in	S protein			
18	L	*	*	*	F	F	NTD
20	Т	*	*	*	*	Ν	
26	Р	*	*	*	*	S	
69,70	H, V	*	Delete	Delete	*	*	
570	A	*	*	D	*	*	
614	D	G	G	G	G	G	
655	H	*	*	*	*	Υ	
681	Р	*	*	Н	*	*	
692	Ι	*	V	*	*	*	
701	A	*	*	*	V	*	
716	Т	*	*	I	*	*	
982	S	*	*	A	*	*	Heptad repeat 1
1027	Т	*	*	*	*	Ι	
1118	D	*	*	Н	*	*	
1229	М	*	Ι	*	*	*	Transmembrane doma

Int J Biol Sci. 2021; 17(6): 1476–1485. Published online 2021 Apr 10. doi: 10.7150/ijbs.59137

The summary of notable genetic variants

SARS-CoV-2 Variants of Concern (VOCs)

• Alpha (B.1.1.7 lineage)

 \mathbf{E}) (

- Beta (B.1.351 lineage)
- Gamma(P.1 lineage)
- Delta (B.1.617.2 lineage)

SARS-CoV-2 Variants of Interest (VOIs)

8

Epsilon (B.1.427 and B.1.429);

Zeta (P.2); Eta(B.1.525);

Theta (P.3); lota (B.1.526);

Kappa(B.1.617.1)

and Lambda(C.37).

The CDC has designated the epsilon variants as VOC and zeta, thetaas VOIs.

 \mathbf{O}

Transmissibility or virulence, reduction in neutralization by antibodies obtained through natural infection or vaccination, the ability to evade detection, or a decrease in the effectiveness of therapeutics or vaccination

https://www.ncbi.nlm.nih.gov/books/NBK554776/



Chest CT in SARS COV2

9

Robert M. Kwee Author Affiliations. Published Online:Oct 23 2020https://doi.org/10.1148/rg.2020200159 Thomas C. Kwee ,



Chest CT Protocol

Patients

- A surgical mask and placed in an isolation room. A strong case need distance of 2 m in combination with wearing a face mask.
- Non–CE chest CT, unless detect pulmonary embolism (PE).
- Patients of all ages infected with SARS-CoV-2 and minimize radiation ALARA (as low as reasonably achievable) principle.

Robert M. Kwee Author Affiliations Published Online:Oct 23 2020<u>https://doi.org/10.1148/rg.2020200159</u> Thomas C. Kwee,

Chest CT Protocol

CT personnel

- Should use appropriate PPE, including face masks, eye protection, gown, and gloves. Increasing the air-exchange per hour or using high-efficiency particulate air (HEPA) filtration in CT examination rooms.
- Deep cleaning of the CT room is necessary before imaging the next patient. All material coming into or near contact with a patient with (suspected) COVID-19 should be disinfected.
- After chest CT is performed, the CT examination room downtime may be between 30 minutes to 1 hour to allow for room decontamination and passive air exchange, according to a policy implemented by the University of Washington. As a result, patient throughput will be limited.

Robert M. Kwee Author Affiliations Published Online:Oct 23 2020https://doi.org/10.1148/rg.2020200159 Thomas C. Kwee,



Normal Chest CT

The incidence of normal chest CT findings in symptomatic patients with COVID-19 is estimated at about 10.6% (95% CI: 7.6%, 13.7%)

- + During the first 4–5 days after symptom onset (in 13.9%–33.3% of patients).
- + During the later stage of the infection (in 1.2%–4.0% of patients).
- + In asymptomatic patients with COVID-19 is considerably high (an estimated 46% of patients).

Low viral loads, confinement to the upper respiratory tract and likely host factors that lead to false-negative chest CT findings.

Chest CT Abnormalities with High Incidence (>70%)



ED

Bilateral ground-glass opacities in a peripheral distribution, 59M



13

Bilateral ground-glass opacity 70 M dilated segmental vessels



Bilateral ground-glass opacities in a peripheral distribution, 59M

Including ground-glass opacities, vascular enlargement, bilateral abnormalities, lower lobe involvement, and posterior predilection.

Robert M. Kwee Author Affiliations Published Online:Oct 23 2020https://doi.org/10.1148/rg.2020200159 Thomas C. Kwee,

Chest CT Abnormalities with Intermediate Incidence (10%–70%)



TEDIC

Ground-glass opacity 63 M a subpleural curvilinear opacity



Crazy-paving pattern, 66 M



Halo sign, 55-year-old man

consolidation (51.5%), linear opacity (40.7%), septal thickening and/or reticulation (49.6%), crazy-paving pattern (34.9%), air bronchogram (40.2%), pleural thickening (34.7%), halo sign (34.5%), bronchiectasis (24.2%), nodules (19.8%), bronchial wall thickening (14.3%), and reversed halo sign (11.1%). unilateral (15.0%), multifocal (63.2%), diffuse (26.4%), single and/or focal (10.5%), middle or upper lobe involvement (49.3%– 55.4%), peripheral location (59.0%), and central and peripheral location (36.2%)

14





Chest CT Abnormalities with low incidence (10%).

Several chest CT findings have been reported to be uncommon in RT-PCR test–proven COVID-19 cases, and these include pleural effusion (5.2%), lymphadenopathy (5.1%), tree-in-bud sign (4.1%), central lesion distribution (3.6%), pericardial effusion (2.7%), and cavitating lung lesions (0.7%)

Development of cavitating lung lesions with COVID-19

16



(a, b) Axial nonenhanced CT images (lung window) obtained at hospital admission show ground-glass opacities in both lungs (early progressive stage). (c, d) Axial nonenhanced CT images (lung window) obtained after 10 days show progressive organizing consolidation (peak stage). (e, f) Axial nonenhanced CT images (lung window) obtained 40 days after the baseline CT images (a, b) show cavitating lesions in both lower lobes (arrow) (late stage).



Imaging Classification and CT Features of COVID-19 Pneumonia

17

Table 2: Imaging Classification and CT Features of COVID-19 Pneumonia

Imaging Classification	Rationale	CT Features
Typical appearance	Commonly reported imaging features of greater specific- ity for COVID-19 pneumonia	 Peripheral, bilateral, ground-glass opacities with or without consolida- tion or visible intralobular lines ("crazy-paving" pattern) Multifocal ground-glass opacities of rounded morphology with or with- out consolidation or visible intralobular lines (crazy-paving pattern) Reverse halo sign or other findings of organizing pneumonia (seen later in the disease)
Indeterminate appearance	Nonspecific imaging features of COV- ID-19 pneumonia	Absence of typical features AND the presence of the following features: multifocal, diffuse, perihilar, or unilateral ground-glass opacity with or without consolidation lacking a specific distribution and that are nonrounded or nonperipheral Few small ground-glass opacities, with a nonrounded and nonperiph- eral distribution
Atypical appearance	Uncommonly or not reported features of COVID-19 pneu- monia	Absence of typical or indeterminate features AND the presence of the following features: isolated lobar or segmental consolidation without ground-glass opacities; discrete small nodules (centrilobular, "tree-in-bud" appearance); lung cavitation; smooth interlobular septal thickening with pleural effusion
Negative for pneumonia	No features of pneu- monia	No CT features to suggest pneumonia.

Source.—Adapted and reprinted under a CCBY 4.0 license from reference 51.

Findings classified as indeterminate appearance of COVID-19 pneumonia



Findings classified as indeterminate appearance of COVID-19 pneumonia in a 26year-old woman. Axial nonenhanced chest CT image (lung window) shows an area of groundglass opacity (arrow) in the posterior basal segment of the left lower lobe. No other lung abnormalities were visualized. The RT-PCR test results were positive for SARS-CoV-2.



18

Findings classified as indeterminate appearance of COVID-19 pneumonia according to the RSNA chest CT classification system in a 24-year-old woman. Axial nonenhanced chest CT image (lung window) shows ground-glass opacities (arrow) in the right upper lobe. In addition, there are discrete centrilobular opacities in the upper lobes. The RT-PCR test results were negative for SARS-CoV-2 but positive for influenza type A.



Imaging Classification and CT Features of atypical COVID-19 Pneumonia



Findings classified as atypical appearance of COVID-19 pneumonia in a 94-year-old woman. Axial nonenhanced chest CT image (lung window) shows subtle centrilobular treein-bud opacities (arrows) in the left lower lobe. The RT-PCR test results were negative for SARS-CoV-2 but positive for influenza type A.

19



Mixed chest CT findings

20



Mixed chest CT findings in an 86-year-old man. (a) Axial nonenhanced CT image (lung window) obtained at hospital admission shows sublobar consolidation (arrow) in the posterior segment of the right upper lobe, a finding more consistent with lobar pneumonia than COVID-19. (b) Axial CT image obtained at a more superior level shows the presence of ground-glass opacities (arrows) Altogether, the findings were classified as indeterminate for COVID-19 pneumonia, according to the RSNA chest CT classification system. The RT-PCR test results were positive for SARS-CoV-2.



Chest CT of COVID-19 Complications

In cases of clinical worsening, chest imaging is advised to assess for COVID-19 progression or secondary cardiopulmonary complications such as acute respiratory distress syndrome (ARDS), PE, superimposed pneumonia, or heart failure that can potentially be secondary to COVID-19– induced cardiac injury



Acute Respiratory Distress Syndrome

22



Development of ARDS in a 60-year-old man with COVID-19. (a) Axial nonenhanced CT image (lung window) obtained at hospital admission shows peripherally located ground-glass opacities (arrow), mainly in the right lung. Note the preexisting centrilobular and paraseptal emphysema. (b) Axial contrastenhanced CT image (lung window) obtained after 3 days shows a marked progression of lung abnormalities (arrows).



Pulmonary Embolism



PE in a 73-year-old man with COVID-19. (a) Axial nonenhanced CT image (lung window) at baseline shows peripherally diffuse ground-glass opacities in both lungs. (b) Axial contrast-enhanced CT image (lung window) obtained after 10 days shows increased consolidation in both lungs. Note the bronchial dilatation within involved portions of the lungs. (c, d) Axial contrast-enhanced CT image (mediastinal window) (c) and sagittal reconstruction (d) obtained 10 days after the baseline images show a filling defect (arrow) in a segmental pulmonary artery branch in the right lower lobe, consistent with PE.



Superimposed Pneumonia

/4



Superimposed pneumonia in a 69-year-old man with COVID-19. (a) Axial nonenhanced CT image (lung window) at baseline shows ground glass opacities (arrows) posteriorly located in the left upper lobe and both lower lobes and an area of consolidation (arrowhead) in the right lower lobe. (b) Axial contrast-enhanced CT image (lung window) obtained after 22 days shows increased consolidation in both lower lobes (red arrows) and consolidation with central cavitation in the left upper lobe (yellow arrow). The culture of puslike bronchial fluid was positive for *Staphylococcus aureus*. Note the presence of pneumomediastinum (arrowhead), which is probably due to long-lasting positive-pressure ventilation.

Cardiac Injury





Cardiac injury occurs in 12.5%–19.7% of hospitalized patients with COVID-19 and is an independent risk factor for in-hospital mortality. Pericardial effusion manifests in an estimated 5.2% of patients with COVID-19, with a higher incidence in those with severe or critical illness cardiac injury in COVID-19. Although pericardial effusion is a nonspecific finding, radiologists should suggest the injury when pericardial effusion is depicted on chest CT images.

https://www.ncbi.nlm.nih.gov/pmc/articles/PMC7927542/



26

Abdoninal CT of SARS COV2

Abdominal CT in COVID-19 patients: incidence, indications, and findings

Erica B. Stein, Katherine Bojicic, Bamidele Otemuyiwa, Dharshan Vummidi, Aamer Chughtai, and James H. Ellis Author information Article notes Copyright and License information Disclaimer

Purpose

The purpose of this study was to evaluate the frequency, indications, and findings of abdominal CTs ordered in the initial evaluation of patients who had a positive COVID-19 test performed in our institution.

Methods

Retrospective chart review was performed on all patients who had a positive test for COVID-19 performed at a single quaternary care center from 1/20/2020 through 5/8/2020. In a subset of patients undergoing abdominal CT as part of the initial evaluation, the demographics, suspected COVID-19 status at the time of scan, presenting complaints, and abdominal CT findings were recorded. Cardiothoracic radiologists reviewed and scored the visualized lung bases for the likelihood of COVID-19.

Results

Only 43 (4.1%) of 1057 COVID-19 patients presented with abdominal complaints sufficient to warrant an abdominal CT. Of these 43 patients, the vast majority (39, 91%) were known or suspected to have COVID-19 at the time of the scan. Most (27/43, 63%) scans showed no acute abdominal abnormality, and those that were positive did not share a discernable pattern of abnormalities. Lung base abnormalities were common, and there was moderate interreviewer reliability.

Conclusion

A minority of COVID-19 patients present with abdominal complaints sufficient to warrant a dedicated CT of the abdomen, and most of these studies will be negative or have abdominal findings not associated with COVID-19. Appropriate lung base findings are a more consistent indication of COVID-19 infection than abdominal findings.



Diagnostic Accuracy of Chest CT

A meta-analysis, which included six studies comprising a total of 1431 patients who were mainly symptomatic and at high risk for COVID-19, reported a chest CT pooled sensitivity of 94.6% (95% CI: 91.9%, 96.4%) and a pooled specificity of 46.0% (95% CI: 31.9%, 60.7%) in the detection of COVID-19.

- A negative chest CT examination result certainly does not exclude COVID-19. The proportion of false-positive chest CT examination results is substantial and due to overlapping imaging features with numerous other diseases, including other viral pneumonias
- It is important to realize that CT is not the standard for the diagnosis of COVID-19, but its findings help suggest the diagnosis in the appropriate setting. It is crucial to correlate chest CT findings with epidemiologic history, clinical presentation, and RT-PCR test results.



Additional Chest CT in Patients Who Undergo CT of Other Body Regions

This scenario was not addressed in the Fleischner Society consensus statement. CT is widely used in the emergency department, with the head and abdomen being among the most commonly imaged body regions . In COVID-19 endemic areas, additional chest CT may be performed to help detect COVID-19 in patients who undergo extrathoracic CT.

- The results of several studies in COVID-19 endemic regions have shown that incidental chest CT findings suggestive of COVID-19 pneumonia can be detected in the visualized lung parenchyma in patients who underwent CT of other body regions, such as CT angiography of the head and neck, CT of the cervical or thoracic spine, and CT of the abdomen. These patients should undergo subsequent RT-PCR tests before the diagnosis of COVID-19 can be confirmed



Role of Chest CT in Diagnostic Decision Making

According to the Fleischner Society consensus statement

Asymptomatic Patients and Patients with Mild Respiratory Symptoms

chest imaging is not indicated as a screening test for COVID-19 in asymptomatic patients or in patients with mild respiratory symptoms of COVID-19 (ie, absence of significant pulmonary dysfunction or damage)

Patients with Moderate to Severe Respiratory Symptoms

chest imaging is indicated in patients with moderate to severe respiratory symptoms (ie, presence of significant pulmonary dysfunction or damage) and any pretest probability of COVID-19 infection, when RT-PCR test results are negative, and in any patient for whom an RT-PCR test is not performed or not readily available



SpO2/FiO2 ≤ 315 is suggestive of ARDS.

symptoms.



Clinical case at MEDIC

Case 1

6419197 [H]ọ: * Đặng Thị								
* 196	63 Địa chỉ: * Thôn K4, X. Hương Phú, H. Na							
h Án Chí Định [F2] [T]oa thuốc 🗌 Tái khám 🗌 Toa EN								
	160.0 Nặng (Kg): 59.00 Nhiệt độ (°C): 36.9							
g):	115/51							
n/Phút):	91							
ệnh:	BN VÀ NGƯỜI NHÀ KHAI ĐÃ SỐNG TẠI SG NHIỀU NĂM, 1 THÁNG NAY KHÔNG ĐI KHỎI SG, KHÔNG TIẾP XÚC VỚI NGƯỜI ĐẾN TỪ VÙNG DỊCH, KHÔNG YẾU TỐ							
	PK NHIË	M CHUY	ÊN	8H56 – 10	H30			
m:	4 NGÀY	(: ỚN LẠ	NH, HO ÍT	Khám nhiễ	m			
	VIÊM PHỔI P VÀ T							
:	HO ÍT, ỚN LẠNH, KHÔNG SỐT. TỔNG TRẠNG TỐT, SINH HIỆU ỔN, ĐI LẠI BÌNH THƯỜNG							
				.::	Chuyển			
	ĐK 19/05/2021 10:37 - BS Khám: Khám phối							



Clinic case at MEDIC





RT PCR dương tính BV PNT

Bilateral ground-glass opacities in a peripheral distribution, low lobe and posterior.







Cas	se 2	Clinica	case	at M	EDIC		
* 588 h: * 197	82999 [H]ọ: * 71 Địa chỉ: *	Đặng Thị 70 Ấp 3, X. Phướ	c Kiếng, H. Nhà I	Tên: * Điện thoạ Khóm: 01/03	i: 0708885232	Nữ TH Nghề nghiệp: <mark>nộ</mark> i	IU P i trợ
định: n): Hg): ần/Phút): bệnh:	158.0 Nặng (Kg): 125/78 110 VIÊM GAN SIÊU K CỔ TỬ CUNG P	56.00 Nhiệt độ (℃): VI B MẠN TRỊ BV NH T VÀ XẠ TRỊ 25 TIA	Chẳr bộ: Phâr HỆT ĐỚI BVUB 2020 Chẳr	a đoán sơ n loại:	lhấn F3 ÔN THƯƠNG PHO COVID ÂM TÍNH	ÔI/ CT/ TEST NHA	NH
nám: ng:	ỚN LẠNH KÉO E KHÓ THỜ, KHÔI MẤT MÙI - VI, BI CHI ĐINH CT PH TẦM SOÁT COV KÉO DÀI, KHÔN CÓ TỒN THƯƠN	DÀI, KHÔNG HO, K NG ĐAU NGƯC, KH N KHÁM IỚI CÓ TON THƯC ID BĂNG PCR, ỨN G HO, KHÔNG ĐA IG PHỔI ĐA NỚT \	HÔNG HÔNG I LẠNH U HỌNG, /ÙNG	:hú BS:	<mark>BV NGUYĒN TRI P</mark>	hương	
	NGOẠI VI HAI BI	ÊN/ CT ĐK: 01/07/2021 07:	Chuy 54 - BS Khám	ển khoa: 📘	Ihấn F3		

Ngay 01/07/2021 (Da du ket qua)							
TÊN XÉT NGHIỆM	KÉT QUẢ	KHOẢNG THAM CHIẾU	MÃ QT				
I. MIĚN DỊCH - IMMUNOLOGY							
XN nhanh kháng nguyên SARS CoV-2	Âm tính	Negative					
(Dịch tỵ hầu)							



RT PCR âm tính **BV NTP**



ge:60 years

MEDIC













34



Clinical case at MEDIC

35



Treatment drugs of SARS COV2

FDA Approves First Treatment for COVID-19

mdesivir injects

100 mg/20 ml



For use under Emergency Use Authorization (EUA)

Each mL contains 5 mg of remdesivir in 20 mL solution

For Immediate Release: October 22, 2020 Español

Today, the U.S. Food and Drug Administration <u>approved</u> the antiviral drug Veklury (remdesivir) for use in adult and pediatric patients 12 years

Remdesivir is administered by intravenous infusion over a period of 30–120 minutes once daily for 5 days, with a maximum total treatment duration of 10 days if there is no clinical improvement.¹ Molnupiravir Shows Promise for Treatment of Moderate COVID-19 in Phase 2/3 Trial

36

MOLNUPIRAVIR

Part 1, 302 participants, symptom onset within 7 days, receive either 200 mg, 400 mg or 800 mg of molnupiravir or a placebo. No infectious virus could be recovered at study day 5 in any molnupiravir-treated patients.

Part 2, evaluate an 800 mg dose of molnupiravir administered twice daily.

Estimated phase 3 in September or October of 2021. Five Indian pharma giants to collaborate on clinical trials of Molnupiravir.









- Typical appearance CT features of COVID: ground-glass opacities, vascular enlargement, bilateral abnormalities, lower lobe involvement, and posterior predilection. a subpleural curvilinear opacity, crazy-paving pattern, halo sign.
- Atypical appearance CT features of COVID: pleural effusion (5.2%), lymphadenopathy (5.1%), tree-in-bud sign (4.1%), central lesion distribution (3.6%), pericardial effusion (2.7%), and cavitating lung lesions (0.7%).
- Secondary cardiopulmonary complications on CT of COVID such as acute respiratory distress syndrome (ARDS), PE, superimposed pneumonia, or cardiac injury (Pericardial effusion).
- Although CT scan is not a diagnostic and screening tool, familiarity with different imaging findings and their differential diagnosis can be helpful in rapid and accurate decision-making.
- FDA Approves first treatment for COVID-19 (remdesivir 22-10-2020) and molnupiravir shows promise for treatment of moderate COVID-19 in Phase 2/3 Trial (Estimated phase 3 in September or October of 2021)



Thanks for your attention best regards

MED





References

1. Sarah Alsobaie. Understanding the Molecular Biology of SARS-CoV-2 and the COVID-19 Pandemic: A Review. Accepted for publication 4 May 2021 Published 16 June 2021 Volume 2021:14 Pages 2259—2268. Department of Clinical Laboratory Science, King Saud University, Riyadh, 11451, Saudi Arabia.

2. Marco Cascella; Michael Rajnik; Abdul Aleem; Scott C. Dulebohn; Raffaela Di Napoli. Features, Evaluation, and Treatment of Coronavirus (COVID-19). Last Update: July 17, 2021. Books. NCBI

3. Thomas C. Kwee, Robert M.Kwee. Chest CT in COVID-19: What the Radiologist Needs to Know. Published Online:Oct 23 2020https://doi.org/10.1148/rg.2020200159

4. Loannis P. Trougakos, Kimon Stamatelopoulos, Ourania E.Tsitsilonis, Evmorfia Aivalioti, Dimitrios Paraskevis, Efstathios Kastritis, George N. Pavlakis and Meletios A. Dimopoulos. Insights to SARS-CoV-2 life cycle, pathophysiology, and rationalized treatments that target COVID-19 clinical complications. *Journal of Biomedical Science* volume 28, Article number: 9 (2021) <u>Cite this article</u>