

Role of HRCT for Rapid Triage of Patients with COVID-19 Pneumonia

Hina Hafeez Abbasi¹, Misbah Durrani², Umme Kalsoom³, Anum Zahoor⁴, Fizza Batool⁵, Shawana Sharif⁶

^{1,3,5} Senior Registrar, Department of Radiology, Benazir Bhutto Hospital, Rawalpindi.

⁴ Assistant Professor, Department of Radiology, Benazir Bhutto Hospital, Rawalpindi.

² Associate Professor, Department of Radiology, Rawalpindi Institute of Cardiology, Rawalpindi.

⁶ Senior Registrar, Department of Dermatology, Benazir Bhutto Hospital, Rawalpindi.

Author's Contribution

¹ Conception of study

^{1,2} Experimentation/Study conduction

^{1,2,3} Analysis/Interpretation/Discussion

^{1,2,3} Manuscript Writing

^{2,4,5,6} Critical Review

^{4,5,6} Facilitation and Material analysis

Corresponding Author

Dr. Hina Hafeez Abbasi

Senior Registrar,

Department of Radiology,

Benazir Bhutto Hospital,

Rawalpindi.

Email: hinahafeez32@gmail.com

Article Processing

Received: 05/03/2021

Accepted: 17/06/2021

Cite this Article: Abbasi, H.H., Durrani, M., Kalsoom, U., Zahoor, A., Batool, F., Sharif, S. Role of HRCT for Rapid Triage of Patients with COVID-19 Pneumonia. *Journal of Rawalpindi Medical College*. 31 Aug. 2021; 25 COVID-19 Supplement-1, 110-116. DOI: <https://doi.org/10.37939/jrmc.v25i1.1657>

Conflict of Interest: Nil
Funding Source: Nil

Access Online:



Abstract

Objective: To assess the diagnostic performances of HRCT for COVID 19 pneumonia for efficient triage of patients, in comparison with RT-PCR reverse transcription-polymerase chain reaction test.

Materials and Methods: It is a retrospective comparative study conducted in Benazir Bhutto hospital affiliated with Rawalpindi medical university from March 25th to April 25th, 2021. HRCT of 500 patients was selected from a central computer server and their RT-PCR results were also obtained from the HMS system of the hospital. HRCT was reported as "Definitely COVID positive", "Possible COVID positive" or "COVID negative" by experienced radiologists. Sensitivity, Specificity, positive predictive value (PPV), and negative predictive value (NPV) were calculated using the final RT-PCR test as a standard of reference.

Results: RT-PCR test of 207 patients were positive, whereas 293 were reported negative. HRCT was reported as "Definitely COVID positive" in 222 cases (44.4 %), "Possible COVID positive" in 24 cases (4.8%), and "COVID negative" in 254 cases (50.8%). Comparing only Definitely COVID positive category with RT-PCR results sensitivity, specificity, PPV and NPV were 90.3%, 88%, 84.2% and 92.8%.

Conclusion: CT chest is the most reliable, sensitive, and rapid tool for triaging of patients as COVID positive or negative in busy emergency departments as compared to RT-PCR which is time-consuming and has limitations such as faulty sampling technique, limited kits, and variable sensitivity.

Keywords: Coronavirus, High resolution Computed Tomography, RT-PCR, Sensitivity, and Specificity, Positive predictive value, negative predictive value.

Introduction

COVID-19 pandemic has changed our lives irrevocably. It has put a huge strain on our already weak health care system.¹ The health care workers and medical resources are stretched to their maximum capacity.² Although in the current pandemic every patient with respiratory symptoms is taken as suspected COVID it also becomes imperative to effectively triage such patients into COVID positive or negative so that effective measures could be taken to timely manage the potential COVID complications. RT-PCR test has variable sensitivity, its time consuming with limitations of shortage of kits as well as faulty sampling technique. In this context, quick triage of patients as COVID positive or COVID negative is essential to support not only critically ill patients, patients requiring emergency surgery but also for the protection and safety of health care workers.^{3,4}

HRCT has an important role in the diagnosis, detection of complications, and prognostication of coronavirus disease 2019 (COVID-19).⁵ The fact of limited availability of FDA-approved RT-PCR testing kits, the delay in test results, and different values of sensitivity of PCR tests across the globe have further emphasized the growing interest in the role of early detection of COVID-19 by HRCT. Although reverse transcription-polymerase chain reaction (RT-PCR) is the standard laboratory test to confirm the diagnosis of COVID-19,⁶ HRCT may represent a quick and valid tool in the initial assessment of the suspected COVID patient population.

Due to resource constraints, all the patients suspected of COVID-19 were not elected for HRCT in our study. The patients who developed worsening of respiratory status or developed complications were referred to our department for HRCT. At the same time, virological testing of these patients was also being performed.

This study aimed to analyze the efficiency of the HRCT for quick triage of patients with COVID-19 as COVID Positive or COVID Negative compared to reverse transcription-polymerase chain reaction (RT-PCR) as a method of reference.

Materials and Methods

This retrospective study was conducted in Benazir Bhutto Hospital (Tertiary care hospital) from March 25th to April 25th, 2021. All CT examinations requested by departments of (medicine, surgery, gynecology,

and ICU) during this period were extracted from our Radiology Electronic Requesting system. Patients under 18 years old, pregnant females, patients already known to have interstitial lung disease and pulmonary edema were excluded. A total of 1300 CT were requested, including 730 chest CT among which 500 were indicated for COVID-19 suspicion and were at risk for disease progression. All these 500 patients (sex-ratio M/F=282/218) also underwent RT-PCR test.

CT Protocol and Analysis:

HRCT was performed on patients with volume reconstruction at 0.625 mm to 1.5 mm slice thickness (gapless). CT images were acquired during a single inspiratory breath-hold

Proper precautions were taken during patient transfer and scanning to reduce the likelihood of the spread of disease with proper PPE, adequate donning and doffing technique, and post scanning decontamination measures.

CT scans were analyzed by senior radiologists from the radiology department.

Common imaging features of COVID disease were ground-glass opacities, crazy paving, consolidations, (preferably bilateral and peripheral) ⁽⁷⁾, focal vessel enlargement, sub-pleural bands, and architectural distortion. Possible complications include pneumothorax, pneumo-mediastinum, ARDS, and pulmonary embolism. ⁽⁸⁾

On the final HRCT report, patients were TRIAGED according to guidelines proposed by the Radiological Society of North America as "Definitely COVID Positive" when typical signs were found including peripheral bilateral ground-glass opacities with or without consolidation or visible intralobular lines (crazy-paving), multiple ground-glass opacities with or without consolidation or visible intra-lobular lines, reverse halo sign or other signs of organizing pneumonia.

Absence of typical findings or presence of features like multifocal, peri-hilar, central, unilateral ground-glass opacities with or without consolidation lacking the specific distribution and are non-rounded or non-peripheral, isolated lobar or segmental consolidation without ground-glass opacities, discrete small nodules (tree-in-bud appearance), lung cavitation or associated with large pleural effusion or lymphadenopathy⁽⁹⁾ were categorized as "Possible COVID"

"COVID Negative" when the HRCT was normal or there were no CT features to suggest pneumonia or HRCT demonstrating another disease⁽¹⁰⁾.

PCR Testing:

This test was performed after RNA extraction by a real-time RT-PCR detection system with internal and external positive controls using the SARS COV-2 protocols. Samples were collected from nasopharyngeal and oropharyngeal swabs. The average time taken to acquire results is 48 hours in our setup.

Results

On the RT-PCR test, 207 patients were tested positive to COVID-19 whereas 293 were reported negative.

Chest CT was triaged as “COVID Positive” in 222 cases (44.4%), “Possible COVID Positive” in 24 cases (4.8%), and “COVID Negative” in 254 cases (50.8%). Our study included 500 patients with Males (56.4%) and Females (23.8%). Common presenting complaints were Shortness of breath (84%), fever (73%), and cough (25%). The majority of patients had co-morbidities like tuberculosis (31%), hypertension (20%), Ischemic heart disease (14%), and diabetes mellitus (16%). The most common complication was Acute Respiratory Distress Syndrome (22.6%). See Table 1 for the detail of these characteristics

Table 1: Base-line characteristics of patients

| <i>Characteristics</i> | <i>COVID-19 Negative</i> | <i>COVID-19 Positive</i> |
|----------------------------|------------------------------|------------------------------|
| Age (years) | | |
| Mean \pm SD | 37 \pm 18.1 | 45 \pm 15.5 |
| Gender | | |
| Male | 163(32.6%) | 119(23.8%) |
| Female | 130(26%) | 88(17.6%) |
| Smoking | | |
| No | 180(36%) | 190(38%) |
| Yes | 74(14.8%) | 56(11%) |
| Symptoms | | |
| Fever | | |
| No | 66(13.2%) | 67(13.4%) |
| Yes | 169(33.8%) | 198(39.6%) |
| Cough | | |
| No | 200(4%) | 174(34.8%) |
| Yes | 41(8.2%) | 85(17%) |
| Shortness of breath | | |
| No | 30(6%) | 47(9.4%) |
| Yes | 137(27.4%) | 286(57.2%) |
| Sore throat | | |
| No | 200(40%) | 234(46.8%) |
| Yes | 31(6.2%) | 35(7%) |
| Loss of smell | | |
| No | 236(47.2%) | 250(50%) |
| Yes | 4 (0.8%) | 10(2%) |
| Diarrhea | | |
| No | 200(40%) | 162(32%) |
| Yes | 96(19%) | 42(8.4%) |
| Travel history | | |
| No | 163(32.6%) | 280(56%) |
| Yes | 39(7.8%) | 18(3.6%) |
| Co-morbidities | | |
| Diabetes | | |
| No | 202(40.4%) | 211(42.2%) |
| Yes | 33(6.6%) | 54(10.8%) |
| Hypertension | | |
| No | 171(34%) | 226(45.2%) |
| Yes | 70(14%) | 33(6.6%) |
| CKD | | |
| No | 292(58%) | 176(35%) |
| Yes | 25(05%) | 07(1.4%) |
| Tuberculosis | | |
| No | 230(46%) | 115(23%) |
| Yes | 135(27%) | 20(04%) |
| Asthma | | |
| No | 290(58%) | 199(19.8%) |
| Yes | 08 (1.6%) | 03(0.6%) |

| | | | |
|---|-----|------------|------------|
| Cardiac problems | No | 168(33%) | 263(52.6%) |
| | Yes | 65(13%) | 04(0.8%) |
| Lung involvement | | | |
| Unilateral (either right/ left lung involved) | No | 22(4.4%) | 17(3.4%) |
| | Yes | 7(1.4%) | 6(1%) |
| Bilateral | No | 141(28.2%) | 74(14.8%) |
| | Yes | 123(24.6%) | 110(22%) |
| Disease distribution | | | |
| Peripheral | No | 68(13.6) | 08(1.6%) |
| | Yes | 49(9.8%) | 70(14%) |
| Central | No | 58(11.6%) | 41(8.2%) |
| | Yes | 17(3.4%) | 7(1.4%) |
| Diffuse | No | 38(7.6%) | 41(8.2%) |
| | Yes | 63(12.6%) | 40(08%) |
| Involvement of Lobes | | | |
| Upper | No | 22(44%) | 20(04%) |
| | Yes | 17(3.4%) | 39(7.8%) |
| Middle | No | 23(4.6%) | 36(7.2%) |
| | Yes | 54(10.8%) | 27(5.4%) |
| Lower | No | 102(20.4%) | 51(10.2%) |
| | Yes | 75(15%) | 34(6.8%) |
| Lymphadenopathy | No | 256(51.2%) | 220(44%) |
| | Yes | 11(2.2%) | 13(2.6%) |
| Pleural effusion | No | 310(62%) | 169(33%) |
| | Yes | 13(2.6%) | 08(1.6%) |
| Lung fibrosis | No | 279(55.8%) | 200(40%) |
| | Yes | 18(3.6%) | 3(0.6%) |
| Attenuation | | | |
| Ground glass appearance | No | 222(70%) | 06(06%) |
| | Yes | 20(30%) | 252(94%) |
| Consolidation | No | 95(19%) | 150(30%) |
| | Yes | 89(17%) | 166(33%) |
| Complications | | | |
| ARDS | No | 190(38%) | 200(40%) |
| | Yes | 40(8%) | 70(14%) |
| Pneumothorax | No | 212(43%) | 215(43%) |
| | Yes | 68(13%) | 5(1%) |
| Pneumo-mediastinum | No | 240(48%) | 25150% |
| | Yes | 4(0.8%) | 5(1%) |

(Figures are presented as whole numbers with percentages in brackets)

We divided the results into two categories and calculated Sensitivity, Specificity, PPV, NPV, and Diagnostic accuracy for each category and then compared the results.

In Category 1, Group I was "Definitely positive" on HRCT (total= 222), and Group II included "Possible Positive" on HRCT and "CT Negative" cases (total= 278). (Table 2)

Table 2: (Category 1)

| | RT-PCR (+) | RT-PCR (-) | Total |
|---------------------------|-----------------------------|-----------------------------|------------|
| CT (+) Group I | A=187 (TRUE POSITIVE) | B=35 (FALSE POSITIVE) | 222 |
| CT(-) Group II | C=20 (FALSE NEGATIVE) | D=258 (TRUE NEGATIVE) | 278 |
| Total | 207 | 293 | |

In Category 2, Group I included both “Definitely Positive on CT” and “Possible Positive on CT” (Total=246), whereas Group II was CT Negative (Total= 254). (Table 3)

Table 3: (Category 2)

| | RT-PCR (+) | RT-PCR (-) | Total |
|-----------------|-------------------|-------------------|------------|
| CT (+) | A=193 | B=53 | 246 |
| Group I | TRUE POSITIVE | FALSE POSITIVE | |
| CT (-) | C=14 | D=240 | 254 |
| Group II | FALSE NEGATIVE | TRUE NEGATIVE | |
| Total | 207 | 293 | |

Comparison between Category 1 & 2 shows Sensitivity, Specificity, PPV, NPV, and Diagnostic Accuracy values. (Table 4)

Table 4:

| | Category 1 | Category 2 |
|----------------------------------|------------|------------|
| Sensitivity | 90.3% | 93.2% |
| Specificity | 88% | 81.9% |
| Positive predictive value | 84.2% | 78.4% |
| Negative predictive value | 92.8% | 94.4% |
| Accuracy | 89% | 86.6% |

Adding the “Possible Positive” group to “Definitely Positive” cases has impaired the Diagnostic accuracy, PPV, and Specificity of HRCT. In this study our purpose was to quickly segregate the patients into COVID Positive and Negative categories, so the inclusion of atypical cases was not suitable as it has decreased PPV of HRCT.

Discussion

With the rapid resurgence in the cases of COVID-19 in the current situation^{11,12} physicians in outpatient departments and emergencies have to face overwhelming patient load and need to decide within the short time frame which patient needs immediate on the spot hospitalization and critical care to minimize potentially serious outcomes. Triage of COVID patients is also important for isolation of COVID + patients in order to control the spread of disease¹³ Although RT-PCR is the investigation of choice¹⁴ for COVID-19, however high number of false-negative cases¹⁵, the limited number of testing kits¹⁶

and time is taken to get the results have convinced the clinicians to consider an alternate test. In our study due to resource constraints, CT chest was not performed for all the patients with clinical symptoms suspicious of COVID-19 but in this retrospective study, we only included the patients referred to our department either due to worsening of respiratory symptoms or those who developed complications.

We further categorized data into two groups. The first one with ‘Definitely COVID +’ on CT was compared with RT-PCR results with a sensitivity of 90.3% and specificity of 88%. On contrary, in previously published studies it was found to be 60% to 98% and 25% to 53% respectively. In this study, specificity turned to be higher as compared to the previous study where it was found to be 25-80%.¹⁵ It can be because a patient who presented to us did not have CT features overlapping with the typical features of COVID-19 thus reducing the number of false-positive cases. In this group, PPV and NPV are 84.2% and 94.8%. NPV can be high due to the reason that patients referred to us were having worsening respiratory symptoms reducing the possibility of false-negative cases on CT. The diagnostic accuracy of the CT chest is 89% making it a reliable tool for COVID -19 testing.¹⁷

In the second group where we have considered “Definitely COVID positive” and “Possible COVID” both as positive cases, the sensitivity and specificity turned out to be 93.2% and 81.9%. PPV has decreased from 84% to 78% as compared to other group. This shows that while triaging the patients into POSITIVE and NEGATIVE cases, the patients with atypical features labelled as ‘POSSIBLE COVID’ should not be included in COVID positive group. The National Health Commission of the People’s Republic of China even stated that diagnosis of COVID-19 could solely be based on chest CT findings⁽⁸⁾. Our findings add value for delivering timely treatment and optimizing the use of medical resources in the pandemic of COVID-19.

To categorize the patients into “Definitely COVID positive”, “Possible COVID” and “COVID negative” we followed the imaging guidelines as proposed by The Radiological Society of North America (RSNA) accredited by the Society of Thoracic Radiology and the American College of Radiology (ACR). The cardinal CT chest findings of COVID 19 pneumonia are Ground glass opacities or consolidations in peripheral distribution, predilection to bi-basal posterior regions.¹⁸

Consolidation or ground-glass opacities could have different morphological appearances, it may present as

patchy, nodular, homogenous, ill-defined, amorphous, or as rounded infiltrates. Other associated features supporting the “definitely COVID-19” pneumonia were thickened interlobular septa superimposed on GGO labelled as crazy paving pattern, reverse halo sign, bronchovascular thickening, and air bronchogram sign. (Figure 1)

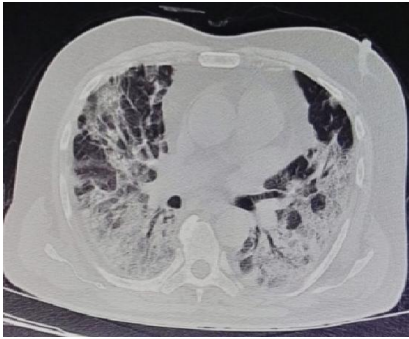


Figure 1: A



Figure 1: B

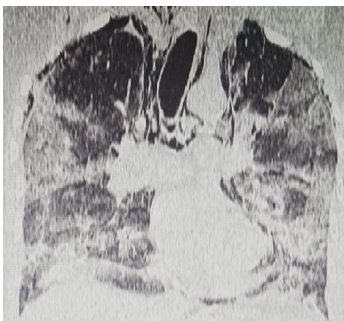


Figure 1: C

Figure 1 shows Axial unenhanced chest CT images (lung window) in a 49-year-old man (A) and a 57-year-old man (B), each with positive RT-PCR test results for SARS-CoV-2, show bilateral areas of confluent mixed ground-glass opacities, inhomogeneous organizing consolidations with air bronchogram in the peripheral distribution along with bronchovascular bundles and subpleural distribution. Few intervening areas of crazy-paving, traction bronchiectasis as well as peripheral subpleural curvilinear parenchymal bands

are also appreciated. Coronal Reformatted image (C) (lung window) is also consistent with classical findings of definitely covid + pneumonia.

While multifocal, diffuse, perihilar, central, or unilateral Ground Glass Opacification (GGO) or consolidation lacking a specific or peripheral distribution are categorized as “Possible COVID” or atypical COVID pneumonia with opacities also being non-rounded in configuration. (Figure 2)

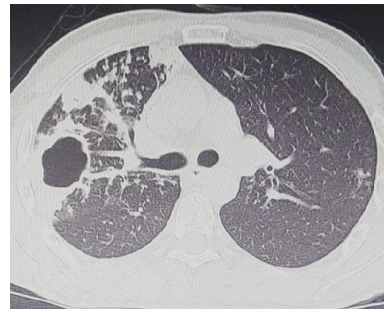


Figure 2: A

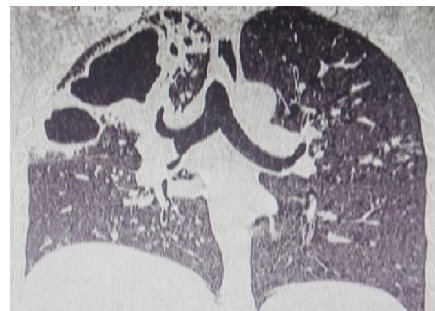


Figure 2: B

Axial and coronal nonenhanced chest CT image (lung window) of a young male patient who presented with fever, Shortness of breath, and mimicking clinical features of covid shows subtle centrilobular tree-in-bud opacities in both upper lobes more pronounced on the right side. At least two large evolving cavities with adjacent interlobular septal thickening are also appreciated. Provisional diagnosis of tuberculosis was made and the patient was triaged and sent back to medical discipline for further clinical and laboratory evaluation. The RT-PCR test results were also negative for SARS-CoV-2.

Large pleural effusion, major lymph node size increase, or bronchiolitis pattern are also included in this domain. Scans were deemed COVID negative when the HRCT was normal or demonstrated another pathology. Ground glass Opacification was defined as partial hazy Opacification of lung parenchyma with no obliteration or bronchovascular markings while consolidation was defined as homogenous dense

opacification with resultant obscuration of underlying lung parenchyma.

Limitations

There are few limitations to our study. Firstly the limited sample size as we have selected patients who were referred to our department and not all the patients visiting the hospital for COVID which can result in higher false-negative results ⁽¹⁹⁾ Another limitation was that we had no follow-up HRCT available owing to heavy workload. Follow-up scans could pave a way for further elaborated studies to look for interval change in the disease process as well as disease complex patterns and varied imaging presentations.

In our country due to limited resources, CT chest is not the first investigation done for COVID-19 diagnosis, however, during this retrospective study we found out that high sensitivity and high NPV of HRCT as compared to PCR should be taken into account and in the future, the clinicians might prefer HRCT as the preferred diagnostic test for COVID-19.

Conclusion

CT chest is the most reliable, sensitive, and rapid tool for triaging of patients as COVID positive or negative in busy emergency departments as compared to RT-PCR which is time-consuming and has limitations such as faulty sampling technique, limited kits and variable sensitivity.²⁰

References

1. A.R. Sahin, A. Erdogan, P.M. Agaoglu, Y. Dineri, A.Y. Cakirci, M.E. Senel, A.M. Tasdog 2019 novel corona virus (COVID-19) outbreak: a review of the current literature *EJMO*, 4 (1) (2020), pp. 1-7
2. Khullar D, Bond AM, Schpero WL. COVID-19 and the Financial Health of US Hospitals. *JAMA*. 2020 Jun 2;323(21):2127-2128
3. Fang Y, Zhang H, Xie J, Lin M, Ying L, Pang P, et al. Sensitivity of Chest CT for COVID-19: Comparison to RT-PCR. *Radiology*. 2020;200432
4. Zu ZY, Jiang MD, Xu PP, Chen W, Ni QQ, Lu GM, et al. Coronavirus Disease 2019 (COVID-19): A Perspective from China. *Radiology*. 2020;200490.
5. Rubin GD, Haramati LB, Kanne JP, Schluger NW, Yim JJ, Anderson DJ, et al. The Role of Chest Imaging in Patient Management during the COVID-19 Pandemic: A Multinational Consensus Statement from the Fleischner Society. *Radiology*. 2020;201365.
6. Mareiniss DP (2020) The impending storm: COVID-19, pandemics and our overwhelmed emergency departments. *American Journal of Emergency Medicine* 38(6), 1293-1294. <https://doi.org/10.1016/j.ajem.2020.03.033>
7. Zhou Z, Guo D, Li C, et al (2020) Coronavirus disease 2019: initial chest CT findings. *European Radiology*, 1-9, 2020. <https://doi.org/10.1007/s00330-020-06816-7>
8. Li M, Lei P, Zeng B, Li Z, Yu P, Fan B, et al. Coronavirus Disease (COVID-19): Spectrum of CT Findings and Temporal Progression of the Disease. *Academic Radiology* 27(5), 603-608, 2020.
9. Chest CT in COVID-19: What the Radiologist needs to know. *RadioGraphics* 2020; 40:1848-1865 <https://doi.org/10.1148/rg.2020200159>
10. Kanne JP, Little BP, Chung JH, Elicker BM, Ketani LH. Essentials for Radiologists on COVID-19: An Update-Radiology Scientific Expert Panel. *Radiology*. 2020;200527.
11. Mareiniss DP (2020) The impending storm: COVID-19, pandemics and our overwhelmed emergency departments. *American journal of emergency medicine*. 38(6) 1293-1294, 2020 <https://doi.org/10.1016/j.ajem.2020.03.033>
12. Mitchell R, Banks C, authoring working party (2020) Emergency departments and the COVID-19 pandemic: making the most of limited resources. *Emergency Medicine Journal* 37(5), 258-259, 2020. <https://doi.org/10.1136/emmermed-2020-209660>
13. Ng K, Poon BH, KiatPuar TH, Shan Quah JL, Loh WJ, Wong YJ, et al. COVID-19 and the Risk to Health Care Workers: A Case Report. *Annals of Internal Medicine* 172(11), 766-767, 2020.
14. CDC 2019-Novel Coronavirus (2019-nCoV) Real-Time RT-PCR Diagnostic Panel; Division of Viral Diseases, U.S. Centers for Disease Control and Prevention: Atlanta, GA, 2020.
15. Xie X, Zhong Z, Zhao W, Zheng C, Wang F, Liu J. Chest CT for Typical 2019-nCoV Pneumonia: Relationship to Negative RT-PCR Testing. *Radiology*. 2020;200343. DOI: 10.1148/radiol.2020200343. pmid:32049601
16. Chen C, Gao G, Xu Y, et al (2020) SARS-CoV-2-positive sputum and feces after conversion of pharyngeal samples in patients with COVID-19. *Annals of Internal Medicine* 172(12), 832-834, 2020. <https://doi.org/10.7326/M20-0991>
17. Anita Kovács 1, Péter Palásti 1, Dániel Veréb 1, Bence Bozsik 1, András Palkó 1, Zsigmond Tamás Kincses 2) The sensitivity and specificity of chest CT in the diagnosis of COVID-19 *Eur Radiol* 2021 May;31(5):2819-2824.
18. Y, Xia L. Coronavirus Disease 2019 (COVID-19): Role of Chest CT in Diagnosis and Management. *AJR Am J Roentgenol*. 2020;1-7.
19. Peiris JS, Chu CM, Cheng VC, Chan KS, Hung IFN, Poon LLM, et al. Clinical progression and viral load in a community outbreak of corona virus-associated SARS pneumonia: a prospective study. *Lancet*. 2003;361:1767-72 [10.1016/S0140-6736\(03\)13412-5](https://doi.org/10.1016/S0140-6736(03)13412-5)
20. Lin C, Xiang J, Yan M, Li H, Huang S, et al. (2020) Comparison of throat swabs and sputum specimens for viral nucleic acid detection in 52 cases of novel corona virus (SARS-Cov-2) infected pneumonia (COVID-19). *medRxiv*: 2020.2002.2021.20026187