

**Demographic and Laboratory Evaluation of the Mucormycosis Received During or Post COVID-19 Treatment**Alpana Singh<sup>1</sup>, Ajay Kumar<sup>2</sup><sup>1</sup>Tutor, Department of Microbiology, Government Medical College, Bettiah, Bihar, India<sup>2</sup>Professor, Department of Microbiology, Government Medical College, Bettiah, Bihar, India

Received: 08-10-2023 Revised: 16-11-2023 / Accepted: 28-12-2023

Corresponding author: Dr. Alpana Singh

Conflict of interest: Nil

**Abstract****Aim:** The aim of the present study was to detect mucormycosis in the clinical species received during or post COVID-19 treatment in our laboratory.**Material & Methods:** A cross-sectional observational study conducted in a tertiary care hospital for five months in association with Department of Microbiology. 50 patients were included in the study.**Results:** Total of n=60 suspected sputum; nasal swab and BAL samples from N=50 of participants were received in our microbiology laboratory during study period. Age, sex and other demography details were collected before sample collection, the average age of the participants was 63.7 ±8.4 years and the majority of participants were male (80%). Although, 44% participants belong to 41-60 year age and 56% of participant belongs to 61-80 year age.**Conclusion:** The present study concluded that the cases of life-threatening MC increase day by day in central India as post complication of covid-19 disease.**Keywords:** COVID-19; Corticosteroid; Diabetes mellitus; Fungus; Mucormycosis

This is an Open Access article that uses a funding model which does not charge readers or their institutions for access and distributed under the terms of the Creative Commons Attribution License (<http://creativecommons.org/licenses/by/4.0>) and the Budapest Open Access Initiative (<http://www.budapestopenaccessinitiative.org/read>), which permit unrestricted use, distribution, and reproduction in any medium, provided original work is properly credited.

**Introduction**

The novel corona-virus 2019 (nCoV-2019) or Severe Acute Respiratory Syndrome Coronavirus-2 (SARS-CoV-2) was an outbreak from Wuhan, China in 2019 and spread rapidly on a global platform forming a global pandemic. [1,2] One may suffer mild to moderate respiratory issues if infected with COVID-19, but in many cases, the virus is affecting different systems of the body at the same time. The covid19 disease causes a range of mild to deadly pneumonia with association of other bacterial, viral and fungal co-infection. Long hospital stay promotes secondary infections in the covid-19 patients and the immune-compromised patients are more prone to develop severe opportunistic infections. [3,4] Apart from attacking the respiratory tract, COVID-19 is also angioinvasive and affects our immunity also. But of late, a number of cases have been reported of mucormycosis in COVID-recovered patients from different states of India. [5]

Although COVID-19-associated pulmonary aspergillosis has been the primary focus in the literature of COVID-19 secondary infections, [6,7] other fungal superinfections, including *Candida* infections, [8] rare mould infections (*fusariosis*), [9,10] and COVID-19-associated mucormycosis.

[11-13] Mucormycosis is a rare but severe fungal infection caused by the Mucorales species of phylum Zygomycota. Naturally, Mucorales occur in soil, their spores spread by air often contaminate foods, water, and clinical specimens. It is aggressive fungal disease and it mainly affects patients with poorly controlled diabetes mellitus and severely immunocompromised patients. [14]

The risk factors predisposing patients to mucormycosis are uncontrolled diabetes, neutropenia, haematological malignancies, organ transplantation, trauma and burn, and use of immunosuppressants such as corticosteroids. [11,15,16] COVID-19-associated mucormycosis can also be mistaken for other angioinvasive fungal infections, particularly with COVID-19-associated pulmonary aspergillosis being the predominant mould disease in COVID-19-associated acute respiratory distress syndrome. The reversed halo sign, predominantly in the peripheral locations of the lung, has been considered to be suggestive of pulmonary mucormycosis in patients with immunodeficiency and useful for the initiation of pre-emptive antifungal therapy; [17] Although cavitory lung lesions might be more specific for mould disease in COVID-19 than the reversed halo

sign, these lesions are frequently observed in both COVID-19-associated pulmonary aspergillosis and pulmonary COVID-19-associated mucormycosis. [18] In the absence of serum antigenic biomarkers and because the availability of PCR testing is low, particularly in low-income and middle-income countries, COVID-19-associated mucormycosis diagnosis is also challenging, with conventional culture and histopathological demonstration of Mucorales being the mainstay of diagnosis, albeit with low sensitivity. In this Covid19 era, the rate of MC cases rapidly growing in the Covid19 patients in India. Mucormycosis is difficult to diagnose which affects outcomes and results in a poor prognosis. Delay in diagnosis increase the mortality rate by about 35- 66%. [19,20] Hence the aim of the study was to detect MC in the clinical species received during or post covid-19 treatment in our laboratory.

### Material & methods

A cross-sectional observational study conducted in a tertiary care hospital for five months months in association with Department of Microbiology, Government Medical College, Bettiah, Bihar, India. 50 patients were included in the study.

### Methodology

The demographic details and clinical diagnosis were recorded from medical records. During the study period, our microbiology laboratory received various specimens such as nasal swabs, ET secretion, sputum, and tissues from our IPD departments such as ICU, and Covid19 ward for the detection of fungal infection in the specimens. Patients admitted in our hospital with a history of fever, cough, body ache and shortness of breath for 4-5 days with have positive report of nasopharyngeal/ oropharyngeal swab for covid19 RT-PCR were included in the present study. Wound swabs were rejected. If the specimens were transported to the laboratory in a sterile container and swabs, the aspirates were immediately performed direct microscopy, KOH smear preparation and gram's stain. Identification was done on the basis of morphology in the microscopy.

### Statistical Analysis

Statistical analysis (Mean, % Value etc.) was done using MS excel 2013.

### Results

**Table 1: Demographic profile of participants**

Gender	N	%
Male	40	80
Female	10	20
<b>Age in years</b>		
21-40	0	0
41-60	22	44
61-80	28	56
<b>Past history of Disease (Immuno-compromised)</b>		
Male	40	88.88
Female	5	11.12
<b>Positive RT-PCR report of nasopharyngeal/ oropharyngeal swab for covid-19</b>		
Male	40	100
Female	10	100
<b>Total stay in hospital (in days)</b>		
10-20 days	32	64
>20 days	18	36
<b>Type of specimens N=60</b>		
Sputum	30	50
Nasal swab	18	30
BAL	12	20

Total of n=60 suspected sputum; nasal swab and BAL samples from N=50 of participants were received in our microbiology laboratory during study period. Age, sex and other demography details were collected before sample collection, the average age of the participants was 63.7 ±8.4 years and the majority of participants were male (80%). Although, 44% participants belong to 41-60 year age and 56% of participant belongs to 61-80 year age.

### Discussion

One may suffer mild to moderate respiratory issues if infected with COVID-19, but in many cases, the virus is affecting different systems of the body at the same time. After months of research, it is now known that apart from attacking the respiratory tract, COVID-19 is also angioinvasive and affects our immunity also. But of late, a number of cases have

been reported of mucormycosis in COVID-recovered patients from different states of India. This post-COVID complication has mainly been seen in diabetics or those treated with steroids. Glucocorticoids are inexpensive, widely available, and have been shown to reduce mortality in hypoxemic patients with COVID-19. [5] The post-COVID-19 state shall be defined as four weeks from the date of being reverse transcription–polymerase chain reaction (RT–PCR) negative (or equivalent laboratory estimation, or clinical criteria constituted by three days without fever and any other symptom. [21]

Total of n=60 suspected sputum; nasal swab and BAL samples from N=50 of participants were received in our microbiology laboratory during study period. Age, sex and other demography details were collected before sample collection, the average age of the participants was 63.7 ±8.4 years and the majority of participants were male (80%). Although, 44% participants belong to 41-60 year age and 56% of participant belongs to 61-80 year age. The genera of Mucorales are one of the best decomposers of organic materials and are often found in decaying organic materials such as rooted fruits and vegetables, plant litter, and animal manure. [22] The *Mucor* sp., *Rhizopus* sp., *Asidia* and *Cunninghamella* are the main causative agent for MC in humans. [23] Spores of the mucorales are highly prevalent in the air. Patients acquire the infection by inhalation, ingestion or traumatic inoculation of the spores from the environment. [24] Other than environmental factors, uncontrolled diabetes mellitus, inappropriate steroid therapy, increased iron accumulation, and the damage caused by the COVID-19 virus may responsible for the MC. [25] Mortality rate of MC is very high, early diagnosis is very essential to reduce the sever morbidity and mortality of patients. [26] The standard approaches for the treatment of MC are usually based on the combination of antifungal therapy and surgical removal of involved tissues. [27]

### Conclusion

Mucormycosis is serious threats in this covid-19 pandemic situation; cases of MC continue to increase in post covid-19 disease patients in India. It is a life-threatening disease that happens due to black mold. The MC occurring in the post Covid-19 patients are a secondary infection and directly linked with the virus, poor glycemia control, widespread use of corticosteroids, and invasive ventilation. Therefore, early screening and diagnosis are much-needed to prevent is a life-threatening event cause by the black mold in post Covid-19 infection.

### References

1. Sharma A, Tiwari S, Deb MK, Marty JL. Severe acute respiratory syndrome coronavirus -2 (SARS-CoV-2): a global pandemic and

treatment strategies. *Int J Antimicrob Agents*. 2020;56(2):106054.

2. Lai CC, Shih TP, Ko WC, Tang HJ, Hsueh PR. Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and coronavirus disease-2019 (COVID-19): The epidemic and the challenges. *Int J Antimicrob Agents*. 2020;55(3):105924.
3. Mahalaxmi I, Jayaramayya K, Venkatesan D, Subramaniam MD, Renu K, Vijayakumar P, et al. Mucormycosis: An opportunistic pathogen during COVID-19. *Environ Res*. 2021;201(6):111643.
4. Chavda VP, Apostolopoulos V. Mucormycosis – An opportunistic infection in the aged immunocompromised individual: A reason for concern in COVID-19. *Maturitas*. 2021
5. Garg D, Mutthu V, Sehgal I, Ramachandran R, Kaur H, Bhalla A, et al. Coronavirus Disease (Covid-19) Associated Mucormycosis (CAM): Case report and systematic review of literature. *Mycopathologia* 2021; 186:289-98.
6. Koehler P, Bassetti M, Chakrabarti A, et al. Defining and managing COVID-19-associated pulmonary aspergillosis: the 2020 ECMM/ISHAM consensus criteria for research and clinical guidance. *Lancet Infect Dis* 2020;21:e149–62.
7. Prattes J, Wauters J, Giacobbe DR, et al. Diagnosis and treatment of COVID-19 associated pulmonary aspergillosis in critically ill patients: results from a European confederation of medical mycology registry. *Intensive Care Med* 2021; 47: 1158–60.
8. Arastehfar A, Carvalho A, Nguyen MH, et al. COVID-19-associated candidiasis (CAC): an underestimated complication in the absence of immunological predispositions? *J Fungi (Basel)* 2020; 6: e211.
9. Poignon C, Blaize M, Vezinet C, Lampros A, Monsel A, Fekkar A. Invasive pulmonary fusariosis in an immunocompetent critically ill patient with severe COVID-19. *Clin Microbiol Infect* 2020; 26: 1582–84.
10. Hoenigl M, Salmanton-García J, Walsh TJ, et al. Global guideline for the diagnosis and management of rare mould infections: an initiative of the European Confederation of Medical Mycology in cooperation with the International Society for Human and Animal Mycology and the American Society for Microbiology. *Lancet Infect Dis* 2021; 21: e24 6–57.
11. Cornely OA, Alastruey-Izquierdo A, Arenz D, et al. Global guideline for the diagnosis and management of mucormycosis: an initiative of the European Confederation of Medical Mycology in cooperation with the Mycoses Study Group Education and Research

- Consortium. *Lancet Infect Dis* 2019; 19: e405–21.
12. Zurl C, Hoenigl M, Schulz E, et al. Autopsy proven pulmonary mucormycosis due to *Rhizopus microsporus* in a critically ill COVID-19 patient with underlying hematological malignancy. *J Fungi (Basel)* 2021; 7:88.
  13. Rudramurthy SM, Hoenigl M, Meis JF, et al. ECMM/ISHAM recommendations for clinical management of COVID-19 associated mucormycosis in low- and middle-income countries. *Mycoses* 2021; 64: 1028–37.
  14. Steinbrink JM, Miceli MH. Mucormycosis. *Infect Dis Clin* 2021; 35:435e52.
  15. Lamoth F, Chung SJ, Damonti L, Alexander BD. Changing epidemiology of invasive mold infections in patients receiving azole prophylaxis. *Clin Infect Dis* 2017; 64:1619–21.
  16. Jenks JD, Reed SL, Seidel D, et al. Rare mould infections caused by *Mucorales*, *Lomentospora prolificans* and *Fusarium*, in San Diego, CA: the role of antifungal combination therapy. *Int J Antimicrob Agents* 2018; 52: 706–12.
  17. Georgiadou SP, Sipsas NV, Marom EM, Kontoyiannis DP. The diagnostic value of halo and reversed halo signs for invasive mold infections in compromised hosts. *Clin Infect Dis* 2011; 52:1144–55.
  18. Koehler P, Bassetti M, Chakrabarti A, et al. Defining and managing COVID-19-associated pulmonary aspergillosis: the 2020 ECMM/ISHAM consensus criteria for research and clinical guidance. *Lancet Infect Dis* 2020; 21: e149–62.
  19. Chavda VP, Apostolopoulos V. Mucormycosis – An opportunistic infection in the aged immunocompromised individual: A reason for concern in COVID-19. *Maturitas*. 2021.
  20. Bhatt K, Agolli A, Patel MH, Garimella R, Devi M, Garcia E, et al. High mortality co-infections of COVID-19 patients: mucormycosis and other fungal infections. *Discoveries (Craiova)*. 2021;9(1):e126
  21. Mucormycosis Advisory from ICMR in COVID19 time, (Screening, Diagnosis & Management of Mucormycosis) pdf, Department of Health Research, Ministry of Health and Family Welfare, Government of India.
  22. Divakar PK. Fungal Taxa Responsible for Mucormycosis/“Black Fungus” among COVID-19 Patients in India. *J Fungi*. 2021;7(8):641.
  23. Ibrahim AS, Spellberg B, Walsh TJ, Kontoyiannis DP. Pathogenesis of mucormycosis. *Clin Infect Dis*. 2012;54(Suppl 1):S16–22.
  24. Richardson MD, Rautemaa-Richardson R. Biotic Environments Supporting the Persistence of Clinically Relevant Mucormycetes. *J Fungi (Basel)*. 2019;6(1):4.
  25. Moorthy A, Gaikwad R, Krishna S, Hegde R, Tripathi KK, Kale PG, et al. SARS-CoV-2, Uncontrolled Diabetes and Corticosteroids- An Unholy Trinity in Invasive Fungal Infections of the Maxillofacial Region? A Retrospective, Multi-centric Analysis. *J Maxillofac Oral Surg*. 2021; 6:1–8.