

Topic 08: COVID-19 Pathogenesis

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Characterization of the Nasopharyngeal Microbiota of COVID-19 Patients According to Disease Severity

H. Carrillo-Ng^{1,*}, J. Del Valle-Mendoza^{1,2}, J. Martins-Luna^{1,2}, W. Silva-Caso^{1,2}, I. Peña-Tuesta^{1,2}, L.J. Del Valle³, G. Pérez-Lazo⁴, S. Miranda², C. Tinco-Valdez^{1,2}, M.A. Aguilar-Luis^{1,2}

¹ Universidad Peruana de Ciencias Aplicadas (UPC), School of Medicine, Research and Innovation Center of the Faculty of Health Sciences, Lima, Peru

² Instituto de Investigación Nutricional, Molecular Biology Laboratory, Lima, Peru

³ Universitat Politècnica de Catalunya (UPC), Barcelona Research Center for Multiscale Science and Engineering, Departament d'Enginyeria Química, EEBE, Barcelona, Spain

⁴ Hospital Nacional Guillermo Almenara Irigoyen, EsSalud, Unidad de Infectología, Lima, Peru

Purpose: Increasing evidence indicates a possible relationship between the pathogenesis of COVID-19 and the nasopharyngeal microbiota. This study aimed to compare the nasopharyngeal microbiota of COVID-19 patients with different disease severity.

Methods & Materials: A comparative study was performed on patients classified into four groups according to their disease severity. A total of 26 patients were recruited for each of the following study groups. Group 1 (G1): patients with a confirmatory diagnosis of SARS-CoV-2 and hospitalized in the ICU. Group 2 (G2): patients with a confirmatory diagnosis of SARS-CoV-2 and hospitalized in regular hospitalization wards. Group 3 (G3): patients with a confirmatory diagnosis of SARS-CoV-2 who were not hospitalized and had mild or no symptoms. Group 4 (G4): healthy patients with a negative result for SARS-CoV-2. The diagnosis of SARS-CoV-2 was performed by reverse-transcriptase polymerase chain reaction (RT-PCR). The nasopharyngeal microbiota was characterized using polymerase chain reaction (PCR) targeting 13 representative bacteria genus.

Results: The detection of some bacteria genus was significantly more frequent in the hospitalized groups compared to healthy subjects. This is the case of *Lactobacillus* with 96.15% in G1, 96.15% in G2, 23.08% in G3 and 15.38% in G4. Similarly, *Prevotella* presented 96.15% in G1, 80.77 in G2, 0.00% in G3 and 19.23% in G4. *Veillonella*, *Bacteroidetes* and *Firmicutes* presented a similar prevalence. On the other hand, the detection of *Eubacterium* was more frequent among asymptomatic and healthy subjects, with 3.85% in G1, 19.23% in G2, 50.00% in G3 and 30.77% in G4. The relative abundance of the bacteria was evaluated and *Lactobacillus* and *Veillonella* were predominant in both of the hospitalized groups (G1 and G2). On the other hand, *Actinobacteria* and *Eubacterium* were predominant in the asymptomatic and healthy groups (G3 and G4)

Conclusion: In conclusion a unique nasopharyngeal microbiota profile was found in COVID-19 patients with different disease severity. *Lactobacillus*, *Prevotella*, *Veillonella*, *Bacteroidetes* and *Firmicutes* were the predominant bacteria genus in critical and hospitalized patients. While, *Eubacterium* and *Actinobacteria* were predominant in the groups of asymptomatic and healthy subjects. Further longitudinal studies are required to determine the prognostic role of the nasopharyngeal microbiota.

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Risk Factors Associated with the Mucormycosis Epidemic During the COVID-19 Pandemic

K. Bhanuprasad*, G.M. Varghese, A.M. S

Christian Medical College, Vellore, India

Purpose: The second wave of the COVID-19 pandemic in India is associated with an unprecedented surge of patients with mucormycosis. This study assessed the risk factors driving the epidemic of COVID-19 associated mucormycosis (CAM) and addressing these risk factors could reduce morbidity and mortality among vulnerable populations.

Methods & Materials: This is a case-control analysis of risk factors in 164 adults from a prospective cohort database, the POISE Mucor Study, approved by the institutional review board and ethics committee with proven invasive mucormycosis of paranasal sinuses admitted in a tertiary care hospital in South India from 01.07.2020 to 10.06.2021 included after informed consent. Cases were patients with COVID-19 confirmed by RT-PCR for SARS-CoV-2 on a nasopharyngeal sample within three months of mucormycosis. Patients with ROCM (Rhino-orbital-cerebral mucormycosis) negative RT-PCR for SARS-CoV-2 were the controls.

Results: The mean age of the 164 patients (132 cases and 32 controls) was 51 years, and 78% were men (table 1). Most patients with CAM had mild COVID-19 (76.7%); 16.3% and 7% had moderate and severe/critical disease, respectively. Almost all patients presented with acute ROCM within 3 months of COVID-19 occurrence.

Diabetes mellitus (present in 97%) remained a strong predisposing factor in both groups.

Uncontrolled diabetes mellitus (HbA1c of >7.0%), 40% of which was newly detected, was associated with CAM (OR: 4.6; p=0.026). Diabetic ketoacidosis and mean HbA1c values were not.

Steroid use was common, even in mild disease, and was strongly associated with CAM (OR:38.3; p<0.001). Oxygen use was uncommon among cases (14.4%). Serum ferritin was significantly higher among patients with CAM (p=0.041); whereas C-reactive protein was not. Acute presentation of ROCM was commoner in CAM. Involvement of brain and orbit were similar between groups. Multivariate analysis revealed that steroid use was independently associated with CAM (OR 28.4; p=0.001).

Conclusion: The current Indian mucormycosis epidemic (mostly acute ROCM) was precipitated by a unique confluence of risk factors – diabetes mellitus, widespread use of steroids, and the COVID-19 infection itself.