Co-infections In People With COVID-19

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Digital Congress of COVID-19 Management and Treatment







- In previous influenza pandemics, bacterial co-infections have been a major cause of mortality.
- Other coronaviruses, such as **MERS-CoV** and **SARS-CoV-1**, show differing levels of bacterial/fungal co-infection.
- whereas there is evidence that *co-infections do not occur in patients infected with MERS-CoV* and occur rarely in patients infected with SARS-CoV-1.

Co-infections in people with COVID-19: a systematic review and meta-analysis; 0163-4453/© 2020 The British Infection Association.



A multicenter retrospective cohort study of

critically ill patients

with MERS-CoV demonstrated that 18% and 5% had

bacterial and viral co-infections, respectively.

Arabi YM, Al-Omari A, Mandourah Y, Al-Hameed F, Sindi AA, Alraddadi B, et al. Critically ill patients with the Middle East Respiratory Syndrome: a multicenter retrospective cohort study. Crit Care Med 2017;45:1683e95.



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Review Article

Co-infections among patients with COVID-19: The need for combination therapy with non-anti-SARS-CoV-2 agents?

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Summary of studies that reported Co-infection in COVID-19

Study	country	No. of patients	Virus	Bacteria	Fungus
Chen	China	99	0	0.1%	Candida (0.4%)
Arentz	USA	21 (critically ill)	3 (14.3)	1 (4.8)	0
Wang	China	104	6 (5.8%): corona influenza A rhinovirus	0	0
Wuhan	China	201	1 (0.6): influenza A	0	0
Kim	USA	116	24 (20.7): influenza/rhinovirus /Enterovirus	0	0

COVID-19

Clinical Infectious Diseases





Article Navigation

ACCEPTED MANUSCRIPT

Bacterial and fungal co-infection in individuals with coronavirus: A rapid review to support COVID-19 antimicrobial prescribing @



Methods:

- MEDLINE, EMBASE, and Web of Science were searched using broad
 - based search criteria relating to coronavirus and bacterial co-infection.
- Articles presenting clinical data for patients with coronavirus infection (defined as SARS-1, MERS, SARS-COV-2, and other coronavirus) and bacterial/fungal co-infection reported.
- Secondary analysis of studies reporting antimicrobial prescribing in



Results:

- 9/18 (50%) studies reported on COVID-19, 5/18 (28%) SARS-1, 1/18 (6%)
 MERS, and 3/18 (17%) other coronavirus.
- For COVID-19, 62/806 (8%) patients were reported as experiencing bacterial/fungal co-infection during hospital admission.
- On secondary analysis, 1450/2010 (72%) of patients reported received
 - antimicrobial therapy.





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Co-infections in people with COVID-19: a systematic review and meta-analysis

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Methods:

- They systematically searched Embase, Medline, Cochrane
- Library, LILACS and CINAHL for eligible studies published from 1

January 2020 to 17 April 2020.

• They included patients of all ages, in all settings. The main

outcome was the proportion of patients with a bacterial, fungal

or viral co-infection.



Results:

- Thirty studies including 3834 patients were included.
- Overall, 7% of hospitalised COVID-19 patients had a bacterial co-infection (95% CI 3-12%, n = 2183, I 2 = 92 ·2%).
- A higher proportion of ICU patients had bacterial co-

infections than patients in mixed ward/ICU settings (14%,

95% CI 5-26, $I 2 = 74 \cdot 7\%$ versus 4%, 95% CI 1-9, $I 2 = 91 \cdot 7\%$).



The commonest bacteria were Mycoplasma

pneumonia, Pseudomonas aeruginosa

and Haemophilus influenzae.





The pooled proportion with a viral co-infection

12 = 62 · 3%), with *Respiratory Syncytial Virus* and

influenza A the commonest.



Conclusions:

• A low proportion of COVID-19 patients have a bacterial co-

infection; less than in previous influenza pandemics.

- These findings do not support the routine use of antibiotics
 - in the management of confirmed COVID-19 infection.



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Invasive aspergillosis in patients admitted to the intensive care unit with severe influenza: a retrospective cohort study



Alexander F A D Schauwvlieghe^{*}, Bart J A Rijnders^{*}, Nele Philips, Rosanne Verwijs, Lore Vanderbeke, Carla Van Tienen, Katrien Lagrou, Paul E Verweij, Frank L Van de Veerdonk, Diederik Gommers, Peter Spronk, Dennis CJJ Bergmans, Astrid Hoedemaekers, Eleni-Rosalina Andrinopoulou, Charlotte H S B van den Berg, Nicole P Juffermans, Casper J Hodiamont, Alieke G Vonk, Pieter Depuydt, Jerina Boelens, Joost Wauters, on behalf of the Dutch-Belgian Mycosis study group



Methods

- They did a retrospective multicentre cohort study. Data were collected
 - from adult patients with severe influenza admitted to <u>seven ICUs</u> across <u>Belgium</u> and The <u>Netherlands</u> during seven influenza seasons.
- Patients were older than 18 years, were admitted to the ICU for more than 24 h with <u>acute respiratory failure</u>, had pulmonary infiltrates on imaging, and a confirmed influenza infection based on a positive airway



PCR test.

Findings

- Data were collected from patients admitted to the ICU between Jan 1, 2009, and June 30, 2016.
- Invasive pulmonary aspergillosis was diagnosed in <u>83 (19%) of 432 patients</u> admitted with influenza (influenza cohort), a median of 3 days after admission to the ICU.
- For patients with influenza who were immunocompromised, incidence of invasive pulmonary aspergillosis was as high as 32% (38 of 117 patients)
- The 90-day mortality was 51% in patients in the influenza cohort with invasive pulmonary aspergillosis and 28% in the influenza cohort without invasive pulmonary aspergillosis (p<0.0001).

COVID-19

Like severe influenza

pneumonia, <u>COVID-19</u> is associated with acute

respiratory distress syndrome (<u>ARDS</u>), which

might be considered

a risk for fungal colonization and infection

of the respiratory tract.



Possible explanations for development of fungal

co-infections include *immuneparalysis*

caused by COVID-19 infection-induced ARDS,

diffuse alveolar damage with severe

inflammatory exudation, and lymphopenia





Mycoses Diagnosis, Therapy and Prophylaxis of Fungal D

REVIEW ARTICLE 🛛 🔂 Free Access

Proven Aspergillus flavus pulmonary aspergillosis in a COVID-19 patient; A case report and review of the literature

Mohammadreza Salehi, Nasim Khajavirad, Arash Seifi, Faezeh Salahshour, Behnaz Jahanbin, Hossein Kazemizadeh, Sayed Jamal Hashemi, Seyed Ali Dehghan Manshadi, Mohammad Kord, Paul E. Verweij, Sadegh Khodavaisy ⊠

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Accepted Articles

Accepted, unedited articles published online and citable. The final edited and typeset version of record will appear in the future.





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Literature review

• The English literature was reviewed for published CAPA cases using search

terms "corona", "COVID-19", "aspergillosis", "CAPA" and "fungal".

• COVID-19 associated pulmonary aspergillosis (CAPA)

• A total of 175 CAPA cases were found.

• Although variable case definitions were used, only 7 (4%) cases were

classified as proven CAPA.



A major challenge remains diagnosing CAPA as the

performance of diagnostic Aspergillus biomarkers remains

suboptimal.

Serum galactomannan (GM)

detection is commonly negative (74%) even in patients with

proven CAPA.



COVID-19- associated invasive pulmonary aspergillosis. Annals of Intensive Care. 2020;10(1):1-4.

Bronchoscopy with bronchoalveolar lavage (BAL)

remains the preferred

diagnostic procedure to diagnose CAPA, and

GM was detected in 79% CAPA patients

who underwent bronchoscopy.



There is conflicting evidence that supports both colonization and invasive infection in Aspergillus positive COVID-19 patients. On the one hand COVID-19 patients with evidence for Aspergillus have survived without antifungal therapy, which suggests that a positive culture or GM represents colonization.

High prevalence of putative invasive pulmonary aspergillosis in critically ill COVID-19 patients. medRxiv. 2020; 8: e48-e49.



There was a trend towards lower mortality in CAPA

patients receiving antifungal therapy, compared

with untreated cases, which suggests that the

mortality, at least in part, may be attributable to IA.

A national strategy to diagnose COVID-19 associated invasive fungal disease in the ICU. 2020;ciaa1298.





Jycoses Diagnosis, Therapy and Prophylaxis of Fungal Diseases



ORIGINAL ARTICLE 🔂 Free Access

Oropharyngeal candidiasis in hospitalised COVID-19 patients from Iran: Species identification and antifungal susceptibility pattern

Mohammadreza Salehi, Kazem Ahmadikia, Shahram Mahmoudi, Saeed Kalantari, Saeidreza Jamalimoghadamsiahkali, Alireza Izadi, Mohammad Kord <mark>... See all authors</mark> v



Mycoses. 2020;63:771–778.

Patients and Methods:

- We here aim to investigate the prevalence, causative agents and antifungal susceptibility pattern of OPC in COVID-19 patients.
- Hospitalised COVID-19 patients with OPC were studied. Relevant clinical data were mined. Strain identification was performed by 21-plex PCR.
- Antifungal susceptibility testing to fluconazole, itraconazole, voriconazole, amphotericin B, caspofungin, micafungin and anidulafungin was performed according to the CLSI broth dilution method.



Results:

- In 53 COVID-19 patients with OPC, cardiovascular diseases (52.83%) and diabetes (37.7%) were the principal underlying conditions.
- The most common risk factor was lymphopaenia (71%).
- C .albicans (70.7%) was the most common, followed by C glabrata (10.7%), C dubliniensis (9.2%), C parapsilosis sensu stricto (4.6%), C tropicalis (3%)
 - and Pichia kudriavzevii (=C krusei, 1.5%). Majority of the Candida isolates

were **susceptible to all three classes** of antifungal drugs.





FIGURE 1 Hospitalised time between diagnosis of COVID-19 and clinical presentations of OPC



Underlying conditions

Cardiovascular diseases	28	52.8
Diabetes	20	37.7
Chronic kidney diseases	11	20.7
Haematological malignancies	5	9.4
Risk factors		
Recipient broad-spectrum	49	92
antibiotics		
antibiotics Corticosteroid therapy	25	47
antibiotics Corticosteroid therapy Admission to ICU	25 26	47 49
antibiotics Corticosteroid therapy Admission to ICU Mechanical ventilation	25 26 16	47 49 30
antibiotics Corticosteroid therapy Admission to ICU Mechanical ventilation Respiratory support	25 26 16	47 49 30
antibiotics Corticosteroid therapy Admission to ICU Mechanical ventilation Respiratory support Non-invasive	25 26 16 49	47 49 30 92.4



Mucormycosis in Patients with COVID-19: A prospective multicenter study

- This was <u>a prospective multicenter study</u> on patients with biopsy proven mucormycosis during 2020 pandemy.
- Inclusion criteria :
 - 1. Mucormycosis should be confirmed on histopathologic examination

2. A verified case was a positive result of real-time reverse-transcriptase– polymerase-chain-reaction (rRT-PCR)SARS COV-2

3. The interval between two infections should not be more than three months.



Results:

- finally **eighteen patients** with laboratory confirmed COVID-19 and Mucormycosis were included and evaluated.
- Mean age of patients was 50 and 60% were male.
- clinical manifestations of mucormycosis were averagely presented within <u>8</u> days (1-40) after COVID-19 infection.
- All patients had underlying diseases including diabetes mellitus and hypertension that were the most prevailing most common comorbodities documented in 85%, and 45%, respectively.
- <u>63% patients had been received intravenous corticosteroids</u> (dexamethasone or methylprednisolone) for their COVID-19 management and one patient had history of <u>mechanical ventilation</u>.





Thank you for your attention