

SARS, MERS AND COVID-19; the Story Continues

From the beginning of millennium three, corona viruses (SARS-CoV: 2003, MERS-CoV: 2012 and SARS-CoV-2: 2019) emerged and caused outbreaks producing considerable global health problems. Although these three viruses have similarities especially regarding clinical features, there are key differences between them that limit the relevance of experiences from previous crises (1). SARS-CoV-2 replicates rapidly in respiratory epithelial cells, including nasal cavity, bronchi, bronchioles, and alveoli. Replication in the upper respiratory tract results in transmission between hosts, while replication in the lower respiratory tract results in the development of lung disease. These three viruses are zoonotic ones which spread from animals and have person to person transmission ability (2). AKI probably via direct renal cytotoxicity through DDP4 receptors which are largely represented in tubules and glomeruli is more frequent with MERS than other two ones (3). COVID-19 generally has a less severe clinical picture, and because of higher R0 can spread in the community more easily than MERS and SARS, which has frequently been reported in the nosocomial setting. Allah kaleth et al., in their report about mortality rate, case fatality rate and years of potential life lost of these three viruses, showed that knowing these information is critical to characterize the severity and understand the pandemic potential of COVID-19 in the early stage of the epidemic. They confirmed that despite lower fatality rate and because of higher transmission rate of COVID-19, it causes a large number of infected patients and more deaths. They also showed that given that COVID-19 has a non-fatal effect on a large number of patients, the estimation of disease burden using the mentioned indices can be an appropriate way for future decision making regarding health policy (4).

Physicians should notice that there are some differences between these three viruses regarding viral spread. The first one is the different viral tropism for respiratory tract and the second can be different inoculum dose at the time of infection which defines the severity of disease. Another point is viral load at the time of symptom onset. In COVID-19, viral load progressively decrease during days but in SARS-CoV recorded highest after 10 days from symptom onset which proves much easier transmission of virus during the initial mild symptomatic or even asymptomatic period (5, 6). Another important point is intrinsic virulence which is higher in MERS-CoV

compared to others. This feature results in more severe clinical symptoms and hospital/ICU admission which reduces community transmission of virus. Gastrointestinal involvement and diarrhea are more common in SARA especially in Hong Kong outbreak and GI spread is also shown in MERS-CoV (7). As SARS-CoV can bind to ACE2 receptor and replicate in enteric epithelium, it can be the same for SARAV-CoV-2 transmitting via this route (8). There are still many unanswered questions especially regarding its epidemiological features such as mortality and capacity to spread on a pandemic level. In the absence of a vaccine, we should notice that speed of it spreads will depend on how conscientiously members of the public and hospital workers observe well-established infection prevention and control (IPC) principles—hand hygiene, cough etiquette, social distancing and, in healthcare settings, isolation of affected patients, and appropriate use of personal protective equipment (PPE). Using convalescent healthcare workers for care of confirmed COVID-19 patients can be very important. Establishing national guideline and revising them during the time are some of the most important responsibilities for ministry of health and CDC. During crisis, using telemedicine (9) can decrease the rate of transmission and appropriate using of social media can increase the social information regarding disease, pathogenesis, symptoms, transmission method, diagnosis and treatment. Primary healthcare programs (e.g. maternal and child health, antenatal, and vaccination programs) during the crisis should cope with the ongoing waves of transmissions and ready for interventions that improve patients outcome. As there is not any effective treatment for COVID-19, its early diagnosis and appropriate management of patients especially critically ill patients with COVID-19 are the cornerstone of its management (10, 11). Regarding treatment in the absence of vaccine, it is recommended an the first stage to use pharmaceuticals and their combinations (protease inhibitors, interferon compounds, antiviral antibodies) aiming to suppress diverse targets during virus propagation and during the second disease stage, it seems crucial and reasonable to rely on administration of pathogenetic drugs to restrict life threatening events resulting in marked inflammation, intoxication, hypoxia and infection (12). In future research, some aspects like microbiota/ microbiome, microRNAs and mesenchymal stem cells should be considered as some potential

interventions. The future research should be focused on the biological properties of these viruses using virus isolation, reverse genetics and in vitro and in vivo infection evaluations to help the prevention and control of emerging crisis. The rapid spread of viruses from natural hosts to humans is largely due to human activities, including modern agricultural practices and urbanization. Therefore, the most important method to prevent viral zoonosis is to maintain the barriers between natural reservoirs and human society. Finally, we hope to be able to identify the most suitable approach to combat this crowned dragon as soon as possible and make this world a healthy place to live again.

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Table1, Different characteristics of three types of coronavirus

	SARS-CoV	MERS-CoV	SARS-Cov-2
Phylogenic origin	Clade I, Cluster IIB	Clade II	Clade I, Cluster IIa
Animal reservoir	bat	bat	bat
Receptor	ACE2	Dipeptidyl peptidase	ACE2
Mode of transmission	Droplets, spread from bats, which infected civets. Human to human transmission through close contact	Droplets, touching infected camels or eating their milk or meat. Limited human to human transmission through close contact	Droplets, Touching or eating an infected. Human-to-human transmission occurs through close contact
Mean incub [±] period	5-6 days	2-7 days	7-14 days
At risk	Elderly, med comorbid	Elderly, med comorbid	Elderly, med comorbid
R ₀	1.7-1.9	0.7	1.5-3.5
Treatment	No specific	No specific	No specific
Vaccine	no	no	no
ICU admission	23-34%	53-89%	24%
ARDS *	20%	20-30%	18-30%
AKI *	6.7%	40-50%	3%
Mortality (Hosp ^α patient)	3.6-15.7%	30-4%	10-11%
Overall Fatality	9%	34.4%	2-4%

±: incubation period

α: hospitalized patients

*ARDS: acute respiratory distress syndrome; AKI: acute kidney injury

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