SARS-CoV-2 infection – spread and pathogenicity

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Abstract

In December 2019, a new disease, similar to severe acute respiratory syndrome (SARS), was reported in Wuhan, China. It was quickly indicated that the causative agent of this new coronavirus disease 2019 (COVID-19) is a previously unknown coronavirus, now called SARS coronavirus 2 (SARS-CoV-2). The result of the global outbreak of COVID-19 in the world (currently COVID-19 is present on all continents, except Antarctica) is the pandemic status 2019–2020, as declared by the WHO and Public Health Emergency of International Concern (PHEIC).The virus has a high epidemic potential and is effectively transmitted between humans. The primary route for SARS-CoV-2 infection to spread is air-droplet transmission. In addition, SARS-CoV-2 can be transmitted through direct contact with an infected person, or indirectly via coronavirus-contaminated materials or objects. Current data also indicate the possibility of an alternative route of SARS-CoV-2 infection – through the gastrointestinal tract. The article discusses various SARS-CoV-2 transmission options, with particular attention paid to the role of saliva and gastrointestinal tract in the spreading of the virus. Current data on SARS-CoV-2 pathogenicity and clinical symptoms of COVID-19 are also analyzed. Expanding knowledge about SARS-CoV-2 infection, especially in terms of its spread, will contribute to further actions aimed at preventing transmission of this pathogen.

Key words

SARS-CoV-2, saliva, dentistry, gastrointestinal infection, COVID-19

INTRODUCTION

In December 2019, in the city of Wuhan, Hubei Province, China, there emerged a new disease with an unexplained etiology. Symptoms of the disease were similar to severe acute respiratory syndrome (SARS), but infection with SARS-CoV and MERS-CoV as causative factors were excluded [1]. At the beginning of January 2020, the new virus was identified and described as 2019-novel coronavirus (2019-nCoV) [2]. The World Health Organization (WHO) named the virus severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and caused the rapid spread worldwide of coronavirus disease 2019 – COVID-19 (the name of this disease was introduced and recommended for use by the WHO). SARS-CoV-2 belongs to the Coronaviridae family, Betacoronavirus genus, and is phylogenetically closely related to other SARS-like coronaviruses [3, 4]. The result of the global outbreak of COVID-19 (currently COVID-19 is present on all continents, except Antarctica) is the pandemic status 2019–2020, declared by the WHO and the Public Health Emergency of International Concern (PHEIC) on 11 March 2020 [5].

OBJECTIVE

SARS-CoV-2 has a high epidemic potential. Knowledge about possible transmission routes of SARS-CoV-2 is therefore particularly necessary for more effective reduction of its transmission. The aim of the study is to present current scientific data about possibilities of spreading SARS-CoV-2 between humans, and its pathogenicity.

Spreading of SARS-CoV-2. It is already well known that SARS-CoV-2 is efficiently transmitted between humans. Cases of familial clustering have been reported [6, 7] and it is currently accepted that SARS-CoV-2 is one of the super-spreading pathogens. The main, human-to-human transmission occurs through respiratory droplets when inhaled air contains saliva with secreted droplets from respiratory tract during coughing, sneezing, and some surgical/medical procedures [8, 9]. In addition, SARS-CoV-2 can be infected through direct contact with an infected person or indirectly via coronavirus-contaminated materials or objects [4, 9]. Recently, a very high viral load contained in the saliva from patients with COVID-19 (reaching 1.2 × 10⁶ copies/ml) and the possibility of virus transmission via saliva among patients without coughing or other respiratory symptoms, are indicated [10]. It is also suggested that the presence of SARS-CoV-2 in the oral cavity can result from major- or minor-salivary gland infection, with subsequent release of particles in saliva via the salivary ducts [10]. This suggestion may be supported by earlier studies documenting that salivary gland epithelial cells can be infected by SARS-CoV within a short time after infection in rhesus macaques [11]. The presented data prove that human saliva may play a significant role in the spreading of SARS-CoV-2.
COVID-19 is therefore a new and serious problem in dentistry. Possible virus transmission through direct contact with saliva droplets and aerosols, generated during dental clinical procedures, is a reason for the high prevalence of COVID-19 among dentists [12]. Indirect contact infection is also possible: SARS-CoV-2 has the ability to survive on paper or copper surfaces from 4 – 24 hours, on steel instruments for at least 48 hours, and 72 hours on plastic surfaces [13]. In light of the above, the use of diagnostics of SARS-CoV-2 in patients’ saliva could improve effective strategies for COVID-19 prevention, especially for dentists and healthcare professionals who perform aerosol-generating procedures.

Current data also indicate the possibility of an alternative route of SARS-CoV-2 infection – through the gastrointestinal tract [14]. The presence of live SARS-CoV-2 or viral nucleic acids has been shown in stool samples of patients. The occurrence of gastrointestinal symptoms in some patients with COVID-19 may confirm that digestive tract as a potential source of faecal–oral transmission of SARS-CoV-2 [15, 16].

SARS-CoV-2 pathogenicity. SARS-CoV-2 is characterized by tropism to human cells, showing specific receptors – angiotensin-convertase enzyme 2 (ACE 2) and co-localized in its proximity – cellular transmembrane serine protease 2 (TMPRSS 2) [17, 18]. ACE 2 determines virus attachment to the target cells, while TMPRSS 2 is necessary for viral entry into cells and for viral spreading in the infected host [18]. ACE 2 is expressed commonly on the surface of epithelial and endothelial cells. Its high expression was found on type 2 lung alveolar cells (AT2), oesophagus upper and stratified epithelial cells, cholangiocytes, myocardial cells, kidney proximal tubule cells, and bladder urothelial cells [19, 20]. It has also been shown that high expression of ACE 2 can occur on mucosa of the oral cavity, especially on the oral tongue [19]. In addition, ACE 2 expression was found on lymphocytes within oral mucosa or various organs of the digestive system, and in the lungs [17].

The presented data indicate that although SARS-CoV-2 pathogenicity is manifested principally in respiratory tract, other organs can also be involved. The replication cycle of SARS-CoV-2 takes place in the cytoplasm of target cells, similarly as in most RNA viruses. However, the unique way of replication strategy of coronaviruses (via the transcription of multiple subgenomic RNAs serve as miRNAs for the structural and accessory genes) allows for a very high recombination frequency, which ensures their genetic variability and the emergence of new strains. The process of virus assembly and formation of progeny viral particles is associated with cellular membrane structures of the cell, called ERGIC (endoplasmic reticulum–Golgi intermediate compartment) [9]. Mature progeny viruses are released from cells by exocytosis. SARS-CoV-2 may exert a cytopathic effect (expressed in some SARS patients by the presence of multinucleated syncytial cells) leading to cytolysis, which initiates lung pathology [21]. However, the association of pulmonary damage with severe inflammation in SARS-CoV-2 infection with focal peribronchiolar mononuclear inflammatory infiltrates, is also very possible [22, 23]. The pathogenesis of SARS-CoV-2 infection in humans and lung injury remains unclear. Lung pathology found in COVID-19 is difficult to explain only by the impact of the virus per se.

Recently, it has been indicated that the severity of COVID-19 is accompanied by a significant increase in inflammatory cytokines, mainly IL-6, as well as high levels of D-dimer in peripheral blood [24, 25]. These data suggest that SARS-CoV-2 may induce a cytokine storm, which is the result of an overreaction of the immune system in COVID-19 patients, resulting in severe pneumonia, acute respiratory distress syndrome (ARDS), and multiple organ dysfunction syndrome (MODS).

COVID-19 – clinical symptoms. Although SARS-CoV-2 infection may involve many organs, the most common and the most severe are respiratory symptoms of COVID-19 [23, 26]. The incubation period of the disease is 2 – 14 days, usually 3 – 7 days. The earliest common symptoms of COVID-19 are fever, cough and dyspnea. Recently presented meta-analysis reports their occurrence as follows: 87.3%, 58.1% and 38.3%, respectively [27]. It has also been reported that frequent clinical feature are atypical pneumonia (75.7%) and ground glass opacification (69.9%) in chest X-ray.

Among analyzed patients, the incidence of ARDS was found in 28.8% and MODS in 8.5%. Patients with COVID-19 may also present, although less often, symptoms of damage to various other organs, such as: oral tongue, kidney, liver and small intestine. Most often, the manifestations of the disease include symptoms from the gastrointestinal tract, e.g. diarhoea, nausea/vomiting and abdominal pain. Meta-analysis reported the occurrence of these symptoms as follows: 9%, 6% and 4%, respectively [28]. In addition, approximately 60% of COVID-19 patients had liver impairment, indicating SARS-CoV-2-associated hepatotoxicity, including elevated aminotransferases, hypoproteinaemia, and prothrombin time prolongation [20]. SARS-CoV-2 infection can also be asymptomatic, often especially in children up to 14 years old (28.6%) and adults ≥70 years (27.3%) [29, 30]. Children and the elderly can be estimated as the main population with asymptomatic infection. This phenomenon is difficult to explain only by the reduced effectiveness of the immunological response, especially in the context of the possibility of the rapid spread of SARS-CoV-2 in the host organism.

CONCLUSIONS

SARS-CoV-2 infection represents a global and serious health and life concern for people worldwide; therefore, knowledge about possible transmission routes is essential. The article presents the important role of saliva in the spread of the virus and its potential faecal–oral transmission. In addition, it has been pointed out that COVID-19 pathogenesis is not only determined by SARS-CoV-2 activity per se, but may also be the result of a strong inflammatory response of the host organism. Simultaneously, it is worth notification is that SARS-CoV-2 infection can often be asymptomatic, mainly in children under the age of 14 years, and adults less than 70 years old.

REFERENCES


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