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# **Review Article**

# High Rate Prevalence of Asymptomatic SARS-CoV-2 Infection: Comparative Epidemiology with other Pandemic Flu and Impact on COVID-19 Control Measures

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# ABSTRACT

Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) is a novel coronavirus responsible for the COVID-19 pandemic, an international public health emergency. Conventional routes of transmission of SARS-CoV-2 are respiratory droplets and direct surface-based contact, similar to SARS-CoV, the Middle East respiratory syndrome-related coronavirus (MERS-CoV), and highly pathogenic influenza. Some clinical features of infectivity and viral susceptibility of SARS-CoV-2 are similar to SARS-CoV-1, MERS-CoV, suggesting a difference in the viral tropism despite the phylogenetic homogeneity. The incubation period of SARS-CoV-2 is in line with other coronaviruses, but with a lower case fatality rate, although the presence of comorbidities can make it highly lethal. The major concern with SARS-CoV-2 is its ability to spread silently by the asymptomatic and presymptomatic carriers. About 70-80% positive cases of COVID-19 are coming out to be asymptomatic, presymptomatic, or very mild symptomatic, which is dangerously higher compared to SARS-CoV-1, MERS-CoV, H1N1, and seasonal influenza. Asymptomatic and presymptomatic carriers-based silent spreading of SARS-CoV-2 has become a major concern due to short serial interval, and soaring level of virus shedding from the upper respiratory tract. Asymptomatic transmission is making containment measures difficult to implement, now early detection and isolation of asymptomatic and presymptomatic persons can be an effective strategy to control spread. High transmutability and silent infection rate SARS-CoV-2 will hopefully help in the fast development of community herd immunity, assuming to have 12 to 14 months active spread period compared to 18 to 24 months for previous pandemic flu. Mass screening by rapid antibody tests, especially in congregate living conditions, mandatory use of face masks, social-distancing, and strict execution of sanitization practices even after the relaxation of lockdown, can effectively help to control the COVID-19 infection rate.

# COVID-19 OUTBREAK

The current outbreak of coronavirus disease, first identified in Wuhan, China, has received official name COVID-19 from the World Health Organization (WHO). COVI comes from coronavirus, D stands for disease, and 19 represents 2019, as the virus was first identified in December 2019. The virus causing COVID-19 has been named "severe acute respiratory syndrome coronavirus 2" SARS-CoV-2 by the international committee on taxonomy of viruses (ICTV). The ICTV has recognized this virus as a sister to severe

acute respiratory syndrome causing coronaviruses. This novel coronavirus is a new coronavirus that has not been previously identified. On 30th January 2020, the WHO declared that the SARS-CoV-2 outbreak constituted a public health emergency of international concern. Although coronaviruses endemic in humans normally show mild infection, such as, the common flu, the cross-species transmission has produced some unusually virulent strains that can cause pneumonia and, in serious cases, acute respiratory distress syndrome and death.

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As per the WHO, the coronavirus outbreak follows a fourphase outbreak pattern. Stage I: Imported and sporadic cases causing the first appearance of the disease. In this phase, the disease is first introduced to the population of a country, and positive cases begin to emerge by the people with travel histories to the infected areas. In this phase, a minimal number of persons got the infection, and disease is totally contained. Stage II: Local transmission starts to develop through positive individuals with travel history, or the persons come in direct contact with an infected person. This stage typically confers with transmission virus onto family members, friends, neighbors, and anyone coming in close vicinity of an infected person. In this stage, the viral transmission is monitored by contact tracing, isolating people with symptoms, strict screening measures, social distancing, and lockdown efforts. Stage III: In this stage, community transmission starts when it is difficult to trace the source of the infection spread. Individuals having no travel history to infected "hotspots," or contact with any positive person get infected. In this stage, it becomes impossible to contain the disease, and to stop the chain of transmission; large-scale lockdowns become the only option. Stage IV: Widespread outbreak is the final stage of transmission, making it a pandemic as the number of cases and deaths rapidly multiply. As per Indian Council of Medical Research (ICMR), currently, India is in stage II of coronavirus infection in spite of rapid spread without active contact or travel history. India has surpassed Russia as the third worst-hit country after the USA and Brazil. China has witnessed all stages of transmission and now is in stage IV, where the infection spreads in clusters. USA, Brazil, India, Russia, South Africa, Mexico, UK, Spain, Italy, and Iran are the most affected countries.[1]

# THE SARS-COV-2

Genomic sequencing showed that this pathogenic coronavirus is 96.2% identical to bat coronavirus, and shares a 79.5% sequence identity to SARS-CoV. SARS-CoV-2 virus falls within the subgenus *Sarbecovirus* of the genus *Betacoronavirus*. The virus has zoonotic origins and has close genetic similarity to bat coronaviruses, suggesting it emerged from a bat-borne virus. SARS-CoV-2 is an enveloped, positive-sense, and single-stranded RNA virus. Each SARS-CoV-2 virion is approximately 50 to 200 nm in diameter, having four structural proteins, as S (spike), E (envelope), M (membrane), and N (nucleocapsid) proteins. The N protein holds the RNA genome, and the viral envelope is created by S, E, and M proteins together. [2] The virus uses spike protein to attach and fuse with the membrane of a host cell.

SARS-CoV-2 enters human cells by binding to the receptor angiotensin-converting enzyme 2. For cellular entry, coronavirus uses the ACE2 receptor as a keyhole to unlock the cell. Interferons act as the primary

defenses of the body against viruses, and function as "first responders" in case of foreign pathogen invasion. The immune system releases interferons to alert about the presence of pathogenic coronavirus in the body. The gene that encodes the ACE2 receptor gets stimulated by interferons ramping up expression of ACE2 receptors, which then acts as a carrier for SARS-CoV-2 to infect even more cells, thus, using body's immune response against it. No clinically effective vaccines or specific antiviral drug is currently available for the treatment of COVID-19 infections.<sup>[3]</sup>

# **Infectivity and Viral Susceptibility**

Conventional routes of transmission of SARS-CoV-2 consist of respiratory droplets produced from coughs or sneezes and direct contact, mechanisms that probably occur with SARS-CoV, the MERS-CoV, and highly pathogenic influenza. [4] Indirect contact via contaminated surfaces has become a serious matter of concern. The virus can remain viable on plastic and steel for up to three days, on cardboard for up to one day, and copper for approximately four hours. The virus is inactivated by soap, which destabilizes its lipid bilayer. SARS-CoV-2 can be detected in the gastrointestinal tract, saliva, and urine of an infected person. Viral RNA has also been found in stool samples from infected individuals. [5] Some clinical characteristics of SARS-CoV-2 and SARS-CoV are different despite the phylogenetic homogeneity. The clinical features of COVID-19 mimic fever and cough as the dominant symptoms similar to SARS-CoV. In contrast, gastrointestinal symptoms were uncommon, suggesting a difference in the viral tropism of SARS-CoV-2 compared to SARS-CoV, MERS-CoV, and seasonal influenza. [6]

The median age of the COVID-19 patients was 47 years, with 41.9% of the female patients. Young children and females were less likely to get infected with SARS-CoV-2 than adolescents and males. The lower incidence of positive results in these two groups may have resulted from less exposure to the virus or biologic resistance.<sup>[7]</sup> In other studies, investigators have found that infected children and females were less likely to have a severe disease than adults and males, respectively. [6,8] The most common symptoms of COVID-19 are fever, cough, and lymphocytopenia, and less commonly, diarrhea, nausea, or vomiting. Chest computed tomography showed ground-glass opacity as the most common radiologic finding.<sup>[6]</sup> Initially, it was estimated that the median incubation period SARS-CoV-2 is 2 to 7 days. Analysis based on 158 confirmed cases in China estimated a median incubation period of 5 days with a range of 2 to 14 days, [9] with onset symptoms in 3 to 6 days after assumed exposure.[10] The incubation period of SARS-CoV-2 is 2 to 14 days, which is in line with other human coronaviruses, SARS, and MERS. [11,12] The incubation period of non-SARS human coronavirus is 2 to 5 days. [13] The knowledge of the incubation period is vital for public health activities,



including active monitoring, surveillance, control, and modeling of the disease control system.

Despite the number of deaths associated with COVID-19, it appears to have a lower case fatality rate than either SARS-CoV or MERS-CoV. Approximately 35% of reported patients with MERS-CoV infection have died, [14] whereas, SARS-CoV has a fatality rate of 11%, which can be more than 50% with patients above 65 years. [15] The case fatality rate among people infected with highly pathogenic avian influenza has ranged from 36 to 60%, which is alarmingly high. [16] The death rate with SARS-CoV-2 can vary from 1.4% as reported in China<sup>[6]</sup> to 3.2% reported by WHO, [17] although much higher lethality of 13 to 15% was reported by other studies conducted in China. [10,18] The overall fatality rate of persons with confirmed COVID-19 in the Italian population, based on data up to 17th March, was 7.2%. Whereas between ages 60 to 69, 70 to 79, and above 80, the death rate was 8.6, 35.6, and 52.3%, respectively. The presence of comorbidities, like heart disease, diabetes, and cancer, can increase the risk of mortality by 25%. [8] The fatality rate is defined as the number of deaths in persons tested positive, divided by the number of SARS-CoV-2 cases. India's fatality rate per million population currently is lowest in the world, which is below 2%. The testing strategy has a significant influence on the fatality rate, as countries like the Republic of Korea, Singapore, and Germany have adopted a wide testing strategy for SARS-CoV-2. This has led to the identification of a large number of individuals with mild or limited symptoms, with a much lower case-fatality rate compared to Italy or the USA.

# **Transmission**

People get infected with coronavirus without showing symptoms. This period can be of several days before symptoms kick in. The infected person does not have a cough, fever, and other signs of illness but can transmit the virus to others. Evidence shows that SARS-CoV-2 has this ability to spread silently. Analysis of COVID-19 cases indicates that most of the infected people are generally fine, living a healthy daily life. In many instances, the infected persons do not even know that they have already been infected and can transmit the virus to family members, neighbors, friends, and strangers. This category of infected people is designated as "silent spreaders." Silent spreaders remarkably contribute to the transmission of the COVID-19 disease, basically called the third phase of spread or community transmission.

The first cases of novel coronavirus (2019-nCoV) infected pneumonia (NCIP) occurred in Wuhan, Hubei Province, China, in December 2019 and January 2020. Reports soon started to come out about the transmission of the virus in other geographical regions by infected travelers. Person-to-person transmission of this novel coronavirus in hospital and family settings among close contacts is very high. An extremely soaring rate of COVID-19 outbreak

correlates transmission from the asymptomatic patient and the three-phase outbreak patterns. [18,19] COVID-19 infection in Germany appears to have been transmitted from a mildly symptomatic convalescent patient 1 (German citizen) via an asymptomatic contact with the index patient (Chinese citizen) during the incubation period, in whom the illness was brief and nonspecific. A high sputum viral load observed in the patient 1, lately confirmed prolonged shedding of the virus after recovery. The fact emphasizes that asymptomatic persons are potential sources of COVID-19 infection transmission dynamics of the current outbreak. [20] Recent evidence of SARS-CoV-2 transmission by mildly symptomatic and asymptomatic persons signals that time from exposure to onset of infectiousness (latent period) can be shorter than the incubation period with important implications for transmission dynamics. Based on the 14 days incubation period currently, the length of quarantine or active monitoring of persons potentially exposed to SARS-CoV-2 is done. In some rare cases, longer monitoring periods might be justified as among infected persons, 1.01% can develop symptoms after the end of a 14 day quarantine period. [21]

# Silent Spreaders of Coronavirus

A "silent transmission" occurs when someone who has contracted the virus shows no symptoms but passes the virus to someone else. Silent spreaders are divided into three categories, asymptomatic, presymptomatic, and very mildly symptomatic. Asymptomatic people carry the active virus but never develop any symptoms. Epidemiologists verify that asymptomatic people do not have common symptoms like fever, gastrointestinal problems, cough, or breathing difficulty. It is very tough for medical professionals to identify someone as infected, while showing no symptoms. Asymptomatic carriers can only be confirmed by finding and testing people who were in close contact with any COVID-19 patients. As per the WHO, about 25% of positive tested asymptomatic people, continued to show no signs in follow-up checkups. [22] Since asymptomatic infected person does not know about their infection status, they may not take precautions advised while interacting with people. Presymptomatic people are infected and incubating the virus, but not showing any symptoms yet. Following coronavirus infection, it takes five to six days, sometimes, even two weeks to develop symptoms according to the research conducted between January and February 2020, published in Annals of Internal Medicine.<sup>[21]</sup> The time lag between getting infected with the virus and showing symptoms is called the presymptomatic phase.

Very mildly symptomatic people feel slightly unwell from a COVID-19 infection, but continue to come in close contact with others. Though spreading COVID-19 by symptomatic people having very mild cough or fever cannot be counted as silent transmission, though this happens when people confuse the condition with common

seasonal flu. Very mildly symptomatic people, when they continue to go to public places, may unknowingly spread the disease in the early days of incubation period when they are most infectious. A completely asymptomatic person who never develops any symptoms can transmit the infection. Though silent spreaders have no or mild symptoms, they still infect others who can become very sick with COVID-19. It is truly impossible to determine the impact of asymptomatic cases on spread except mass-scale testing. The data of a small group clinical study on 24 positive tested people in Nanjing, China, shows that those who do not have overt symptoms at the time of testing continued to be asymptomatic one to three weeks after diagnosis had a median age of 14.

Presymptomatic cases of COVID-19 is more common category than asymptomatic. About 75% of people who test positive without showing symptoms turn out to be presymptomatic, displaying coughing, fatigue, fever, and other signs in the later follow-up checks. [22] As per WHO, a presymptomatic person can transmit this virus, most likely in the one to three days period before they start showing symptoms. However, presymptomatic people appear to be most infectious around the time when it starts showing symptoms. [22] In mid-March, 82 residents of a nursing home in King County, Washington, tested positive for the coronavirus. Half of them were free from any symptom when tested for the virus, and most went on to develop symptoms. The coronavirus spread rapidly through the facility within two weeks despite applying isolation policy suggesting transmission from asymptomatic and presymptomatic residents, who were not recognized as having SARS-CoV-2 infection and, therefore, were not isolated. [23] Another study in Singapore, done between February and March, had reported similar evidence of presymptomatic spread among people going to public places. [24]

# **Silent Transmission Fueling Pandemic**

Silent transmission of infectious disease occurs via the asymptomatic carrier who is infected with the pathogen without any symptoms. Asymptomatic carriers play a critical role in the transfer of common infectious diseases, such as, typhoid, influenzas, tuberculosis, measles, cholera, dengue, human immunodeficiency virus (HIV), and now the COVID-19. Asymptomatic carriers are associated with not only the viral disease but also with diseases caused by bacteria and mycobacterium. Controlling asymptomatic disease has become crucial in concern to public health for mitigating the spread of common infectious diseases. The history of the asymptomatic carrier for the first time was connected with Mary Mallon, interestingly nicknamed as Typhoid Mary. She was an asymptomatic carrier of Salmonella typhi, causing typhoid fever in New York City during the late 1800s. She was a cook for several families and soldiers, and nearly 50 cases of typhoid fever were traced back to her. She was guarantined on an island surrounding Manhattan until her death.

WHO estimates that approximately 390 million people get infected with dengue virus worldwide each year, out of which only 96 million develop clinical manifestations, with approximately 75% of asymptomatic patients. Chen et al. reported the presence of dengue virus in people of southern Taiwan, despite the fact that no epidemic has been reported in the past 10 years, demonstrating silent transmission.<sup>[25]</sup> The prevalence of asymptomatic carriage of influenza infections ranges from 60 to 75%, which has a very short generation time and spread very quickly. [26] Asymptomatic measles infections are common among measles-immune persons, but transmission from asymptomatic cases is very rare. As per the European Centre for Disease Prevention and Control (European Union), an estimated 1.7 billion people had latent tuberculosis having a "dormant" form of tuberculosis. Latent tuberculosis infection carriers are asymptomatic and not infectious but can spread disease at a later stage resulting in active tuberculosis. [27] Asymptomatic carriers can transfer cholera usually for a short period, though a few individuals may excrete the Vibrio cholerae for a prolonged period. Asymptomatic HIV infection is the second stage of acquired immunodeficiency syndrome (AIDS), when there are no symptoms, also called clinical latency. During this stage, the virus multiplies in the body, and the immune system slowly weakens. The patients are called longterm nonprogressors, and the duration can be 10 years or more, depending on the HIV multiplication rate and how the immune system responds to the virus. Asymptomatic carriers have a significant role in fueling the epidemic and pandemic of typhoid, influenzas, dengue, HIV, and COVID-19.

# **Role of Silent Spreaders in COVID Pandemic**

The evidence of presymptomatic transmission in Singapore and China, between January to March, supports the likelihood that viral shedding occurs in the absence of symptoms and before symptom onset.<sup>[28,29]</sup> Asymptomatic and presymptomatic SARS-CoV-2 infections in residents of a nursing facility in King County, Washington, suggests that symptom screening initially fails to identify approximately one-half of residents with SARS-CoV-2 infection. Unrecognized asymptomatic and presymptomatic infections contribute to large scale transmission in such settings.<sup>[30]</sup> Even though it was a nightmare for the 3,700 passengers and crew members trapped off the coast of Japan in Diamond Princess cruise ship, it gave researchers a unique chance to study COVID-19. This cruise ship docked at Yokohama Bay, Japan, temporarily became the largest cluster of COVID-19 cases outside China in 1st week of February. Passengers were held under quarantine over two weeks, and researchers found that 46.5% of infected people did not have symptoms at the time of testing. Many eventually did develop symptoms; still, 18% of infected cases remained asymptomatic.[31]



Silent carriers of coronavirus may play a significant role in spreading the disease, according to research conducted in Iceland. Researchers from deCODE genetics and Iceland's Directorate of Health and National University Hospital, Iceland, examined more than 20,000 people, which is about 6% of the country's population in the 3rd and 4th week of March. Targeted testing of more than 9,000 people with COVID-19 symptoms identified 13% positive cases. Out of more than 10,000 people, who voluntarily joined screening on an open invitation without exhibiting symptoms, about 0.8% was tested positive for coronavirus. More than 2,000 people were randomly selected for screening, out of which 0.6% were tested positive, indicating that a significant number of people could have the virus without exhibiting symptoms. About 60% of the participants who tested positive reported having had contact with a known infected person. Notably, 43% of the participants have tested positive but did not have any symptoms, although symptoms developed later in some of them.<sup>[7]</sup> Evidence shows that transmission from asymptomatic infector to close contacts can lead to severe COVID-19 pneumonia. Clinical characteristics of 24 cases with asymptomatic infection from close contacts were screened that showed the transmission potential of asymptomatic COVID-19 virus carriers. The communicable period could be up to three weeks, and some of the communicated patients develop severe illness.<sup>[32]</sup>

Air, surface, environmental, and personal protective equipment got contaminated from infected patients. [33] Environmental contamination with SARS-CoV-2 has been reported, and the possibility of indirect transmission through fomites is a big concern now, as after lockdown also people are getting infected while at home. Vegetables, grocery items, and packed food objects can be contaminated directly by droplets or through contact with an infected person's contaminated hands due to nonrigorous hygiene practices. Researchers have sequenced more than 600 samples of the virus and found more than 291 mutations. [7] The extent of infectivity and lethality of coronavirus changes with its mutation. Human natural defense mechanisms can react differently towards different mutated strains of COVID-19.

# Transmissibility Difference of COVID with other Pandemic Flu

Dr. Anthony Fauci, director of the US National Institute of Allergy and Infectious Diseases, said in a press conference on 28th January, that "in all the history of respiratory-borne viruses of any type, the asymptomatic transmission has never been the driver of outbreaks." On 1st February 2020, the WHO also indicated that "transmission from asymptomatic cases is likely not a major driver of transmission."<sup>[34]</sup> All these assumptions went incorrect soon enough, when reports started to come of rapid escalation of COVID positive cases form Italy and Iran, followed by Germany and the USA. The latest

worldwide data suggests that silent transmission of the COVID-19 virus is playing a dangerous role in electrifying the spread of the pandemic locally, as well as, across the globe. The data reveals that currently, around 60 to 70% of the tested people in India have been found antibody positive despite showing no symptoms. Such individuals look healthy, and as silent carriers would have played a vital role in the spread of the virus, if not contained by the strict lockdown. Collectively all the current research conducted on COVID cases between January to April in Singapore, China, Iceland, Germany, and the USA shows the extent of SARS-CoV-2 spread by the asymptomatic carriers. COVID has started with such ferocity that mostly all the countries effected had to implement partial or strict lockdown policy. Countries like India, applying lockdown sooner before induction of phase II, were able to control the phase III spike effectively, whereas, the countries implementing in later stages are paying a high price. However, asymptomatic carriers of the dreaded virus have been ignored by most of the western countries.

The influenza pandemic of 1918-1919 killed nearly 50 million (5 crores) of people more than World War I, making it one of the deadliest pandemics in human history. The Spanish influenza virus had a profound virulence, with a mortality rate of 2.5%, with the death rate of 20 times higher for 15 to 35-year-olds. People were struck with flu and died rapid deaths, some times within hours. Though no documented research was found on asymptomatic carriers of the Spanish flu, this level of contagious spread would have been possible only with silent carriers. This possibility can be correlated with the fact that officials of San Francisco, one of the hard-hit cities of the US, ordered everyone to wear face masks and banned public gatherings. The restrictions worked, and the numbers of cases and fatalities decreased within weeks. Ensured that the disease spread had slowed down, ban on public gatherings was abolished along with the mandatory wearing of masks, and just after four weeks, the flu rebounded with a prolonged second wave reporting more cases and deaths than the first one.

The asymptomatic infection rate was reported to be less than 0.1% for MERS cases.<sup>[35]</sup> Despite a more than 40% mortality rate, due to the slower spread of MERS, fewer than 200 cases worldwide were reported during the 18 months' period, since the outbreak in 2012. All cases were virtually directly or indirectly related to transmission in the Arabian peninsula, minimizing the advent of super spreaders or a mutation in the MERS virus. Arbitrary estimates of asymptomatic infection were in the range of 30 to 50% for the H1N1 epidemic, [26] which estimated to have caused about 284,000 deaths in 20 months duration. The incidence of asymptomatic cases was found to be 7.5 to 13% out of all SARS-CoV-1 positive cases. [36] SARS-CoV-1 outbreak in the 2003-2004 had infected approximately 8,100 persons in limited geographic areas and killed close to 10% of those infected in 8 months. Whereas, within 8 months, SARS-CoV-2 has infected more than 25 million people and continues to spread rapidly around the world while more than one third (approximately 2.5 billion) of the world's population was under lockdown for 2 to 3 months.

High level of SARS-CoV-2 shedding from the upper respiratory tract of presymptomatic patients was observed.[37] Influenza asymptomatic persons, generally, have lower quantitative viral loads in upper respiratory tract secretions, and a shorter duration of viral shedding. [38] Live coronavirus clearly sheds at high concentrations from the nasal cavity even before symptom development. Because of this, traditional symptom-based screening or only contact tracing-based testing does not have much utility in the COVID containment measure. Ultimately, the rapid spread of COVID-19 across the globe is clear evidence of SARS-CoV-2 transmission from asymptomatic persons.[39] The eventual need to relax current lockdown measures urges for broadened SARS-CoV-2 testing to include asymptomatic persons in prioritized settings of congregate living situations, like in slams, nursing facilities, prisons, jails, mental health facilities, homeless shelters, and hospitalized inpatients. The general public should now be encouraged to use face masks always after the relaxation of lockdown, also when in crowded outdoor or indoor spaces, along with the following common sanitization practices. [40] This unprecedented pandemic calls for unprecedented measures to the level of government, as well as, the general public to defeat it ultimately.

# CONCLUSION

Scientists are suggesting authorities, managing COVID-19 lockdowns, to learn from history and avoid slowing containment efforts too soon. Self-isolation, quarantining, and other social-distancing measures have helped to control the COVID-19 infection rate. All the scientific findings on COVID-19 suggest that to control the pandemic, it is not enough to limit contact with persons having symptoms, as persons without symptoms might also transmit infection. The current rate of asymptomatic and presymptomatic COVID-19 positive persons has increased the challenges of containment measures to control the transmission. Asymptomatic transmission definitely makes containment measures difficult to implement. The strategy must now focus on predicated early detection and isolation of asymptomatic persons along with extensive control measures, like isolation, quarantine, school closures, travel, and mass gatherings restrictions. Early detection of asymptomatic and presymptomatic persons are only possible with quick result oriented mass population testing measures based on reverse transcription polymerase chain reaction (RT-PCR) or more specifically, detection of IgM/IgG antibodies against SARS-CoV-2 by rapid lateral flow immunoassay. The magnitude of this impact will be

purely dependent upon the extent and duration of transmissibility, while a patient is asymptomatic or presymptomatic, which, to date, has been extended from 14 to 28 days by authorities of many countries, as many asymptomatic and mild presymptomatic persons are now tested to be SARS-CoV-2 positive up to 28 days.

The researchers have found that the time interval between infection and subsequent transmission, called serial interval for SARS-CoV-2, is approximately 4 to 8 days. The serial interval for SARS-CoV-1 was 7 days. A shorter serial interval makes an epidemic spread rapidly and harder to contain. Naturally, pandemics eventually ends, when the pathogen runs out of individuals to infect in its course of progression. This period helps the population to develop "herd immunity," as, during this disease susceptible period, enough of the population became immune. All the previous respiratory borne pandemics like Spanish flu, MERS, H1N1, and SARS-CoV-1 were active on an average of 18 to 24 months. Depending on the viral strain, lethality susceptible individuals die between these active disease spread periods. SARS-CoV-2 is spreading at a much faster rate than MERS, H1N1, and SARS-CoV-1, as it is showing a very high rate of asymptomatic positive cases, in addition to the shorter serial interval. Despite many similarities between SARS-CoV-1 and SARS-CoV-2, including genetic relatedness, transmission characters, and incubation period, trajectories of the two epidemics have veered in dramatically speculated to be due to drastically high rate of asymptomatic carriers of SARS-CoV-2. The current pandemic of SARS-CoV-2 has a much higher silent infection rate, and thus, presumably, herd immunity will also arrive swiftly and assumed to have 12 to 14 months active spread period. Rapid antibody tests can be used to determine the percentage of the population that has contracted the disease, and that is, therefore, presumed to be immune.

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