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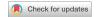
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Airborne transmission of SARS-CoV-2 in indoor environments: A comprehensive review

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The COVID-19 pandemic has caused millions of deaths and great economic loss globally. There has been substantial evidence supporting the airborne transmission of SARS-CoV-2. Airborne route has been considered as a major transmission pathway, which can spread the disease over a longer distance and time. The viral loads in the respiratory tract of a virus carrier are typically below 10° RNA copies/mL and are related to the emission rate of pathogens. Most particles expelled during respiratory activities are smaller than 1–2 µm in diameter. Viral aerosols can remain infectious for hours under typical indoor conditions. Sunlight contributes greatly to the viability inactivation of SARS-CoV-2. The outbreaks in different scenarios are reviewed based on existing data. Most outbreaks were related to long-term care facilities, K-12 schools, restaurants, retail facilities, and offices. The Wells-Riley model for estimating the risk of airborne transmission is introduced, along with model parameters such as the quanta generation rate, virus-containing particle size distribution, and inactivation rate. The effectiveness of various IAQ control strategies for mitigating the airborne transmission risk is analyzed, including PPE, ventilation strategies, partitions, air cleaning, and disinfection technologies, and occupancy control strategies. Both benefits and costs should be considered in designing the control strategies.

Background

The novel coronavirus disease 2019 (COVID-19) pandemic caused by the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) has lasted for over one and half years. It has caused millions of deaths and great economic loss globally (WHO 2021a). Currently, a mass vaccination effort against COVID-19 is getting underway worldwide. But the world is still facing a severe situation of pandemic as new COVID-19 cases have risen to over 600,000 per day over the last few weeks. Many countries are currently facing another serious wave of infections (WHO 2021a). Some emerging SARS-CoV-2 variants, e.g. the delta variant, have increased transmissibility, which further brings the world more unknown challenges (U.S. CDC 2021n). At the current

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stage, minimizing the virus transmission is still essential in reducing the risk of COVID-19 infection.

Studies have revealed that most COVID-19 infections occurred in indoor environments while outdoor outbreaks only account for less than 10% of the infection cases (Bulfone et al. 2021). In addition to fomite route through contacts with infected people or contaminated surfaces and droplet-borne route transmitted by medium (5-100 µm) or large droplets (>100 μm) (Wei and Li 2016), more and more studies have revealed the possibility of airborne transmission through aerosols (<5 μm) for SARS-CoV-2 (D. Lewis 2020a; Y. Li et al. 2020; Luo et al. 2020; Miller et al. 2020; Morawska and Milton 2020; Park et al. 2020; Setti et al. 2020a; Y. Shen et al. 2020; The Lancet Respiratory Medicine 2020a; N. Wilson, Corbett, and Tovey 2020a). Virus-laden aerosols can be expelled by breathing, talking, and coughing by the infectors, and remain suspended in the air over a longer time and distance, and in some circumstances, may travel beyond the most recommended social distance (i.e. 2 m) (Y. Li et al. 2020; Luo et al. 2020; Miller et al. 2020; Y. Shen et al. 2020). Therefore, airborne transmission may play an important role in spreading SARS-CoV-2, particularly in crowded and inadequately ventilated indoor spaces, where virus-laden particles generated by the infectors cannot be removed or diluted efficiently (U.S. CDC 2020d; WHO 2020f, 2020b). Some studies even

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indicated the airborne route as the predominant transmission of COVID-19 (Nature 2021; R. Zhang et al. 2020). It is also believed that airborne transmission is likely the key to explain the superspreading events that vastly occurred in indoor environments (Swinkels 2020). Public health or environment agencies, including WHO, U.S. CDC, China NHC, Europe CDC, Canada PHAC, U.S. EPA, and ASHRAE, have raised the concern on the airborne route of SARS-CoV-2 transmission (ASHRAE 2020b; China NHC 2020; Europe CDC 2020; PHAC 2020; U.S. CDC 2020d; WHO 2020b).

The outbreaks of COVID-19 have been vastly reported in some indoor scenarios like long-term care facilities, manufacturing facilities, correctional facilities, schools and colleges, healthcare facilities and hospitals, retail facilities, restaurants, and offices (CDPHE 2020). Airborne transmission has been suspected as the potential or even predominant route for SARS-CoV-2 transmission in these indoor scenarios. Considering the various indoor settings such as occupant density, behavior and building layout and ventilation settings between different scenarios, the role of airborne transmission and relevant interventions in various scenarios can vary greatly. In order to reduce the airborne transmission risk, many control strategies have been introduced, including personal protective equipment (PPE), improved ventilation and air distribution, standalone air cleaning and disinfection technologies, and occupancy control and restriction (Bazant and Bush 2021; A. K. Melikov, Ai, and Markov 2020; Morawska et al. 2020; J. Shen et al. 2021; J. Zhang 2020). Both benefits and costs of control strategies are crucial for designing the control strategies in a specific scenario.

This study reviewed the previous research on the airborne transmission of SARS-CoV-2 in indoor environments from the perspective of indoor air quality (IAQ) and its control. The existing evidence supporting the airborne transmission of SARS-CoV-2 was reviewed. Then the mechanism of airborne transmission, including the viral load in the respiratory tract, expelled virus-laden particle concentration and size distribution, viral natural inactivation under different environmental conditions, and the transmissibility of some SARS-CoV-2 variants. were introduced. The outbreaks in different scenarios were also reviewed based on the existing data, and the role of airborne transmission in these outbreaks was analyzed. The commonly used models for estimating the risk of airborne transmission were introduced along with the estimation of model parameters. In the end, the IAQ control strategies for mitigating the airborne transmission risk of SARS-CoV-2 are discussed.

Evidence and mechanism of SARS-CoV-2 airborne transmission

Evidence of airborne transmission

There was a controversy about the routes of COVID-19 transmission. During the initial takeoff of the pandemic, close contact with the infectors, contaminated surfaces, and expelled droplets were believed to be the dominant routes. Due to the lack of sufficient evidence, the airborne transmission of COVID-19 was not suggested by major public health agencies in their guidelines

(WHO 2020c, 2020e), although some people raised their concerns on the possible transmission through aerosols (Favre 2020; Offord 2020) based on the experience on other respiratory diseases (Xiao et al. 2018; Yu et al. 2004). With further research on the pandemic, more and more scientific results suggested that airborne transmission played an important role in SARS-CoV-2 infection. Many studies sampled and analyzed the air in different environments where infectors were occupied, including hospital settings, healthcare facilities, quarantined hotels, and other public places and transportations, to figure out whether the virus can be carried by aerosols and travel on airflow. Contrasting results were reported in these studies. Some research did not observe viral RNA in their air samples (Cheng et al. 2020b; Döhla et al. 2020; Dunker et al. 2021; Faridi et al. 2020; U. J. Kim et al. 2020; Lane et al. 2020; Y. H. Li et al. 2020; Masoumbeigi et al. 2020; Nakamura et al. 2020), while most studies detected positive for SARS-CoV-2 in the air (Binder et al. 2020; Chia et al. 2020b; Ding et al. 2021; B. Feng et al. 2021; Z. D. Guo et al. 2020; Hadei et al. 2021; Jiang et al. 2020; Jin et al. 2021; Kenarkoohi et al. 2020; Lednicky et al. 2020; Y. Liu et al. 2020; Ma et al. 2020; Moore et al. 2021; Moreno et al. 2021; Mouchtouri et al. 2020; Razzini et al. 2020; Santarpia et al. 2020; Setti et al. 2020b; Stern et al. 2021; Tan et al. 2020; Zhou et al. 2021). Viral RNA was vastly detected in the air of hospital settings, e.g. patient rooms, intensive care units (ICUs), bathrooms, and corridors, probably due to the high density of infectors in hospitals and the aerosol-generating procedures for the treatment (e.g. intubation), which may generate considerable virus-laden aerosols. Viruses were even detected in the ventilation system in hospitals, indicating the possible viral transmission through ventilation ducts (Horve et al. 2020; Nissen et al. 2020; Ong et al. 2020). Most studies suggested that outdoor air was safe and not infectious for the public excluding crowded sites (Chirizzi et al. 2021; Hu et al. 2020), though viral RNA was detected in some outdoor air samples collected in Italy (Setti et al. 2020b). But the concentrations of virus-laden particles were not evaluated in their study. Therefore, it is unlikely a role of airborne transmission of COVID-19 in outdoor environments, excluding crowded areas.

However, even though viral RNA was detected in the air samples, experts still questioned the airborne viability of SARS-CoV-2 in the carrier aerosols (WHO 2020c). Because most of these studies sought to determine the viral presence, while samples were not evaluated for viral infectivity, which is a key criterion for evaluating airborne transmission (S. Tang et al. 2020). Some studies detected viral RNA in aerosols but none of the positive samples showed evidence of viable virus (Binder et al. 2020; Maestre et al. 2021; Moore et al. 2021; Moreno et al. 2021). However, Santarpia et al. (2020) detected viable viruses in some aerosol samples using the cell culture method in hospitals in Nebraska. Lednicky et al. (2020, 2021) detected viable SARS-CoV-2 isolated from air samples collected in a hospital in Florida and a car driven by a patient. Therefore, the expelled aerosols have the potential to serve as a source of transmission of the virus.

Viral infectivity on aerosols was also vastly supported by laboratory experiments. The virus-laden aerosols generated in experimental conditions can typically remain infectious for hours. van Doremalen et al. (2020) observed that the

Table 1. Major studies on the airborne transmission of SARS-CoV-2.

Peference	Cettings	Findings
	Scinigo	Campin 1
Onsite virus measurements in the air or aerosols	the air or aerosols	
Binder et al. (2020)	Hospital rooms	Very few aerosol samples detected positive (3/195) ^b ; None of the aerosol samples showed evidence of viable vinus:
Cheng et al. (2020h)	Hospital rooms	Air samples detected negative (0/8)
Cheng et al. (2020a)	Hosnital rooms	Air samples detected negative (0/6 natients)
Chia et al. (2020a)	Hospital rooms	Air samples detected positive (2/3 rooms) ² (1.84 × 10 ³ $-$ 3.38 × 10 ³ RNA
de Man et al. (2020)	Nursing home with infectors	Copies in Solution of a mirsing home
Ding et al. (2021)	Hospital rooms	Air samples detected positive (1/46)
Döhla et al. (2020)	Quarantined households (flats or houses) with at	Air samples detected negative (0/15)
Dunker et al (2021)	least one injector Hosnital rooms	Air sammles detected negative
Faridi et al. (2021)	Hosnital rooms	Air samples detected negative (0/10)
B. Feng et al. (2021)	Hospital rooms	Air samples detected positive (1/12) (1112 copies/m ³ in <1 μ m aerosols,
		745 copies/m ³ in >4 μ m aerosols)
Z. D. Guo et al. (2020)	Hospital rooms	Air samples detected positive (14/40)
Hadei et al. (2021)	Public places and transportation	Air samples detected positive (18/28)
Horve et al. (2020)	Hospital HVAC systems	Viral RNA detected at the supply air damper (5/15)
Jiang et al. (2020)	Hospital rooms	Air samples detected positive (1/28)
Jin et al. (2021)	Hospital rooms	Air samples detected positive (1/1)
Kenarkoohi et al. (2020)	Hospital rooms	Air samples detected positive (2/14)
J. Y. Kim et al. (2020)	Hospital rooms	Air samples detected negative (0/52)
Lane et al. (2020)	Hospital rooms	Air samples detected negative in the rooms of patients on ventilators (0/30)
Lednicky et al. (2020)	Hospital rooms	Viable (infectious) virus detected in aerosols (4/6) (6–74 viruses/L)
Lednicky et al. (2021)	Car	Viable (infectious) virus detected in airborne particles of the size range of
		0.25 to 0.5 μm in a car driven by a COVID-19 patient
Y. Li et al. (2020)	Hospital rooms	Aerosol samples detected negative (0/135)
Liu et al. (2020)	Hospital rooms	Air samples detected positive (21/35) (1–42 RNA copies/m ²);
Ma et al (2020)	Hosnital rooms or anarantined hotel	Air complex detected nocitive in an unventilated hotel toilet (176)
Maestre et al. (2021)	Quarantined household with infected occupants	Viral RNA detected on the filter of the HVAC system (approximately $69+43$ RNA conjector ³).
		The detected viral RNA does not imply infectivity;
Masoumbeigi et al. (2020)	Hospital rooms	Air samples detected negative (0/31)
Moore et al. (2021)	Hospital rooms	Air samples detected positive (4/55) (<10-460 RNA copies/m ³);
Moreno et al. (2021)	Dublic transmontations	Viable (infectious) virus was not recovered from any positive air sample; Air camples detected negitive (3/12):
141010110 of al. (2021)	i done transportations	The potential infectivity of the positive samples is considered to be
MC	11	extremely low;
Mouchiouri et al. (2020)	nosputat rooms, nursing nome, terry ooat, tong-term care facility	An samples detected positive $(1/12)$; Viral RNA detected on air exhaust duct surface and the filter of the
Makamina at al (2020)	Homital rooms	HVAC system;
nakannia et al. (2020)	nospital tooms	All samples detected negative (0/11)

.(
Reference ^a	Settings	Findings
Nissen et al. (2020) Ong et al. (2020)	Hospital rooms Hospital rooms	Virus detected in ventilation systems Air samples detected negative (0/3 rooms); Surface complete on oir exhauter detected negatives.
Razzini et al. (2020) Santarpia et al. (2020)	Hospital rooms	Air samples detected positive (100% in contaminated areas) Air samples detected positive (11/18 rooms/patients) (2.42–8.34 RNA copies/L for air samples in patient rooms, 2.08–8.69 RNA copies/L for hallway air samples, 5.37–48.22 RNA copies/L for personal air samples);
Setti et al. (2020a) Stern et al. (2021) Tan et al. (2020) Zhou et al. (2021)	Outdoor environments Hospital rooms Hospital rooms Hospital rooms	Viable (Infectious) virus detected in some samples; Air samples detected positive (20/34) Air samples detected positive (9%) Air samples detected positive during an intubation procedure (1/12) Air samples detected positive (3/44) (9–219 viruses/m ³); Virus observed in exhaled breath (2/9) (\sim 10 ⁵ RNA copies/m ³)
Laboratory experiments		
Dabisch et al. (2020)	Experimental conditions	The time needed for a 90% decrease in infectious virus ranged from 4.8 min under simulated high-intensity sunlight, to greater than 2 h without sunlight
Fears et al. (2020) Schuit et al. (2020)	Experimental conditions Experimental conditions	Infectivity and virion integrity maintained for up to 16 h in aerosols Mean decay rates of aerosols under simulated sunlight were 0.121–0.306 min ⁻¹ (90% loss, 8–19 min) and mean decay rate without simulated sunlight was 0.008 min ⁻¹ (90% loss 286 min)
Smither et al. (2020)	Experimental conditions	Half-lives of infectious virus ranged from 30 to 177 min for the different conditions
van Doremalen et al. (2020)	Experimental conditions	Half-lives of infectious virus in aerosols were around 1.1h
Retrospective analysis on real outbreak events	outbreak events	
Almilaji and Thomas (2020)	Cruise ship	Airborne transmission of COVID-19 through the ventilation system on the Diamond Princess cruise ship could explain the higher-than-expected virus spread into cabins without previously confirmed cases during the diagraphic period
Azimi et al. (2021)	Cruise ship	Aerosol inhalation was likely the dominant contributor to viral transmission on the Diamond Princess cruise shin
Buonanno, Morawska, and Stabile (2020)	Restaurant and choir in a church	Airborne transmission was likely the main route for spreading the virus during the outbreaks in a restaurant in Guangzhou and a choir in
J. Cai et al. (2020)	Shopping mall	Transmission through aerosols in the confined space was likely associated with a cluster of COVID-19 cases in a showning mall in Wenzhou
Chau et al. (2021)	Bar	The superspreading event at a bar in Ho Chi Minh City indicated that crowds in enclosed indoor settings with poor ventilation may be
Crema (2021)	Regions around the Amazon rainforest	considered at figh fisk for transmission Airborne transmission was likely the key to explain the remarkable viral spreading around the Amazon rainforest

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Aerosols were the probable mode of transmission in a quarantined hotel in New Zealand, and the enclosed and unventilated space in the hotel corridor probably facilitated the spread	Transmission of SARS-CoV-2 can occur over distances of at least 8 m in confined spaces in a German meat processing plant under conditions without sufficient ventilation	Airborne transmission through the air duct in the restroom was likely the primary reason to explain the virus spreading among the infectors in an anartment building in Seoul	Vertical spread of virus-laden fecal aerosols via stacks and vents was likely the reason for the outbreak in a high-rise flat building in Guangzhou,	Supported by the positive environmental samples. Airborne transmission was suggested since the case patients seated up to 15 m from the index patient without close physical contact during church cinging in Sydney.	Transmission through aerosols was likely occurred in the business class area during a long-haul flight to Vietnam, where seats are already more widely spaced than in economy class, and infection spread much further	than the existing 2-row or 2-meter rule recommended for COVID-19	Aerosol transmission due to poor ventilation may explain the viral spreading in a restaurant in Guanozhou	Airborne transmission may explain the high attack rate on a tour coach and a minibus in Hunan province	Transmission by the aerosol route was likely; it appeared unlikely that either fomite or ballistic droplet transmission could explain a substantial fraction of the cases in a choir in Mount Vernon	Airborne spread of the virus may at least partially explain the markedly high attack rate observed on a tour coach in Zheiiano province	Aerosols were observed in the restrooms on 25th and 27th floors after flushing the toilet in the 15-floor restroom, which possibly explained the virus spreading between the studied rooms in an apartment building in Guangzhou	Airborne transmission was likely the primary reason to explain the virus	Airborne transmission was highly virulent and represented the dominant route to spread the disease		Airborne spread observed SARS-CoV-2 is transmitted through the air	SARS-CoV-2 is transmitted efficiently via the air Viruses were transmissible between cats via the airborne route	
Quarantined hotel	Meat processing plant	Apartment building	Apartment building	Choir in a church	Aircraft		Restaurant	Tour coach and minibus	Choir in a church	Tour coach	Onsite tracer simulation experiment in an apartment	Apartment building	Outbreaks in China, Italy, and NYC		Experiments on ferrets Experiments on ferrets	Experiments on ferrets Experiments on domesticated animals	
Eichler et al. (2021)	Günther et al. (2020)	Hwang et al. (2021)	Kang et al. (2020)	Katelaris et al. (2021)	Khanh et al. (2020)		Y. Li et al. (2020)	Luo et al. (2020)	Miller et al. (2020)	Y. Shen et al. (2020)	S. Tang et al. (2020)	Wang and Du (2020)	R. Zhang et al. (2020)	Animal experiments	Y. I. Kim et al. (2020) Kutter et al. (2021)	Richard et al. (2020) Shi et al. (2020)	^a Listed in alphabetical order

^aListed in alphabetical order.

^bProportion of positive samples in all available samples (i.e. positive samples).

^cAir samples in two out of three rooms detected positive. Totally 18 samples were collected (six for each room), but the positive sample number was not reported.

^dMeasured viral concentration.

half-lives of infectious virus in aerosols were around 1.1 h, while the viral half-lives measured by Smither et al. (2020) ranged from half-hour to nearly 3 h for different humidity conditions. Fears et al. (2020) observed that infectivity and virion integrity were maintained for up to 16h in aerosols. Dabisch et al. (2020) measured the time required for a 90% decrease in infectious viruses greater than 2 h. The 90% viral infectivity loss time measured by Schuit et al. (2020) was more than 4h. Rapid decay of viral infectivity was also observed when the virus was exposed to simulated sunlight. The 90% infectivity loss time was greatly reduced to between 4.8 and 19 min (Dabisch et al. 2020; Schuit et al. 2020). However, the results of laboratory experiments were questioned by some experts since the aerosol-generating procedure in laboratory conditions was likely stronger than normal human cough conditions as high-powered machines were used to generate particles (WHO 2020c). WHO (2020c) believed that this kind of experimentally induced aerosol-generating procedure cannot reflect the actual setting and therefore was not sufficient for suggesting airborne transmission.

Other studies conducted retrospective analyses on real outbreak events and indicated that airborne transmission was likely to occur in these outbreaks since the other routes were not able to explain the viral spreading in these outbreaks. For example, airborne transmission through the air was likely the key to explain the remarkable viral spreading around the Amazon rainforest (Crema 2021). The outbreaks in apartment buildings in Seoul (Hwang et al. 2021), Guangzhou (Kang et al. 2020; S. Tang et al. 2020), and Ordos (Wang and Du 2020) were believed to be highly relevant with the airborne transmission as no other contacts were reported between some patients. The viral spreading among people during the church singing in Mount Vernon (Buonanno, Morawska, and Stabile 2020; Miller et al. 2020) and Sydney (Katelaris et al. 2021), during the traveling in a coach/bus in Hunan (Luo et al. 2020) and Zhejiang (Y. Shen et al. 2020), during a long-haul flight to Vietnam (Khanh et al. 2020), and during working in a German meat processing plant (Günther et al. 2020), was highly suggested to be associated with the transmission through airborne route since infection occurred over distances of at least 2 m. The outbreaks in a Guangzhou restaurant (Buonanno, Morawska, and Stabile 2020; Y. Li et al. 2020) and a bar in Ho Chi Minh City (Chau et al. 2021) were likely due to the poorventilated air in the room. The airborne transmission was also likely a possible viral spreading route for the outbreaks on the Diamond Princess cruise ship (Azimi et al. 2021) and in a shopping mall in Wenzhou (J. Cai et al. 2020). R. Zhang et al. (2020) analyzed some outbreak events in China, Italy, and New York City, and suggested that airborne transmission was highly virulent and represented the dominant route to spread the disease. Evidence revealed that airborne transmission can partially explain the high attack rate in many superspreading events (Buonanno, Morawska, and Stabile 2020; D. Lewis 2021a). The airborne transmission was also observed between some animals like ferrets and

cats through animal experiments (Y. I. Kim et al. 2020; Kutter et al. 2021; Richard et al. 2020; Shi et al. 2020).

Based on research evidence, the airborne route of viral transmission has been widely recognized by scientists and experts (Aghalari et al. 2021; E. L. Anderson et al. 2020; Greenhalgh et al. 2021; Morawska and Cao 2020). In July 2020, 239 scientists from more than 30 countries highlighted the possible role of aerosols in the transmission of SARS-CoV-2 (Morawska and Milton 2020). Greenhalgh et al. (2021) recently presented ten scientific reasons in support of the airborne transmission of SARS-CoV-2. Many scientific journals, e.g. Nature, The Lancet Respiratory Medicine, The BMJ, and Indoor Air, have published their editorials to suggest the potential of airborne transmission of COVID-19 (A. K. Melikov 2020; Nature 2021; Rijn et al. 2020; J. W. Tang et al. 2021; The Lancet Respiratory Medicine 2020b; N. Wilson, Corbett, and Tovey 2020b; J. Zhang 2020). Most public health and environment agencies, including WHO, U.S. CDC, China NHC, Europe CDC, Canada PHAC, and ASHRAE, have also declared the possibility of airborne transmission in their guidelines. The aforementioned studies on airborne transmission were listed in Table 1. A brief timeline of studies on airborne transmission and responses from public agencies was illustrated in Figure 1. Based on the existing evidence, it is increasingly clear and accepted that airborne transmission is an important contributor to the rapid and long-distance spreading of SARS-CoV-2.

Mechanism of airborne transmission

Virus-containing body secretions and excreta can be aerosolized into infectious virus-laden droplets or particles in a variety of ways (S. Tang et al. 2020). Respiratory secretions can be aerosolized through daily activities (e.g. breathing, talking, coughing, and sneezing) and medical procedures (e.g. intubation) (Zietsman, Phan, and Jones 2019), while excreta can be aerosolized through toilet flushing (Johnson et al. 2013). Considering that respiratory activities are performed all the time, this study mainly focuses on aerosol transmission due to daily respiratory activities. But the SARS-CoV-2 transmission by aerosols generated through medical procedures or toilet flushing is still a potential route. Medical procedures were likely related to the vastly detected viral RNA in hospitals (Tan et al. 2020). Fecal aerosol transmission due to toilet flushing was suspected to be associated with some reported cases in apartment buildings (Hwang et al. 2021; Kang et al. 2020; Ma et al. 2020).

The virus-carrying droplets are formed in the respiratory tract and released from the mouth and nose to the environment through breathing, talking, coughing, and sneezing (Dhand and Li 2020; Wei and Li 2016). The number and size of expelled droplets depend largely on the locations within the respiratory tract where they originate. The oral cavity usually produces larger droplets ($\geq 100~\mu m$ in size) during speech and coughing, whereas smaller droplets originate in the larynx during voicing and coughing ($\geq 1~\mu m$) and the bronchioles during normal breathing ($\leq 1~\mu m$) (Dhand and Li 2020). The time and distance expelled

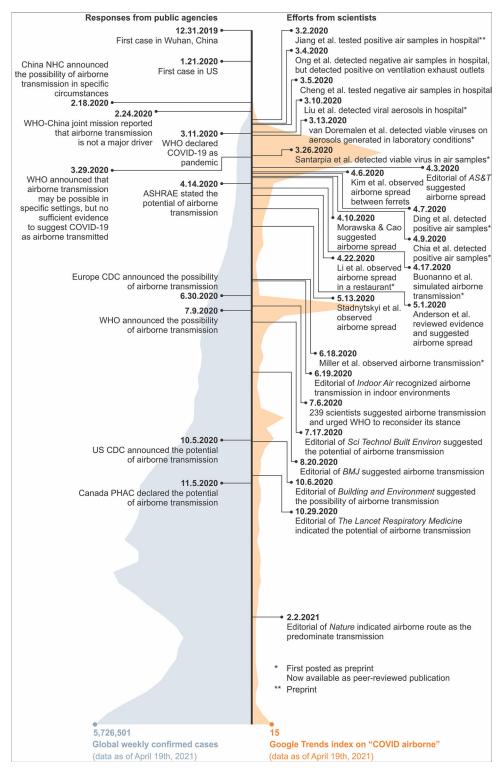


Fig. 1. Timeline of major studies on the airborne transmission of COVID-19 and responses from public agencies (data as of April 19th, 2021 (WHO 2021a)) (Google Trends index indicates the worldwide query data on "COVID airborne" on Google (2020). The index is calculated to be ranged from 0 to 100 (Choi and Varian 2012)).

droplets travel in indoor environments greatly depend on the expiration air jet, particle size, and ambient airflow field. Typically, coughing and sneezing have a higher exhalation

velocity and droplet concentration but a lower event frequency, while breathing and talking have a lower velocity and droplet concentration but a higher event frequency.

0.003 (3.1%)

0.002 (2.0%)

0.098

0.012 (1.8%)

0.006 (0.9%)

0.678

 3.5 ± 0.7

 5.5 ± 1.0

Total

	(/)					
	Numb	er concentration [cn	n^{-3}] and proportion [[%] in different sizes	s of expelled partic	eles
Particle diameter [μm]	Voiced counting	Whispered counting	Unmodulated vocalization	Unmodulated whisper	Breathing	Coughing
$\leq 0.80 \pm 0.08$	0.236 (73.3%)	0.110 (84.6%)	0.751 (69.0%)	0.636 (94.6%)	0.084 (85.7%)	0.567 (83.6%)
1.8 ± 0.3	0.068 (21.1%)	0.014 (10.8%)	0.139 (12.8%)	0.037 (5.5%)	0.009 (9.2%)	0.093 (13.7%)

0.139 (12.8%)

0.059 (5.4%)

1.088

0.004 (3.1%)

0.002 (1.5%)

0.130

Table 2. Number concentration and proportion in different sizes of expelled particles during different activities (adapted from Morawska et al. (2009)).

Chao et al. (2009) measured an average expiration air velocity of 11.7 m/s for coughing and 3.9 m/s for speaking. Large droplets usually deposit or evaporate into small nuclei quickly in indoor environments. Deposition occurs either by gravitational sedimentation, turbulent eddy impaction, or diffusional deposition. The deposited particles can be resuspended to the air due to occupant activities such as walking and hence become a potential source (Y. Li et al. 2020). It was reported that expelled large droplets can typically travel around 2 m before falling on the floor or being evaporated (Wei and Li 2016). Therefore, most droplet transmission likely occurs at close range and the 2-meter social distance has been widely recommended by various guidelines. However, for small droplets, gravitational sedimentation is relatively negligible, whereas turbulent eddy impact or diffusional deposition may occur. Small droplets are subject to the ambient velocity field, and their evaporation (into small nuclei) clearly assists in transporting them a significant distance (Dhand and Li 2020). Therefore, smaller droplets can typically remain suspended in the air for a longer time and travel with the ambient airflow for a farther distance (over 2 m). Evidence showed that particles of diameters 1-3 μm remained suspended almost indefinitely (Wei and Li 2016). Therefore, expelled aerosols (<5 µm in diameter) are believed to play an important role in SARS-CoV-2 transmission.

0.007 (2.2%)

0.011 (3.4%)

0.322

The susceptible individuals' inhalation of the contaminated air that carries infectious aerosols expelled by the infected subjects, is referred to as the airborne route (Wei and Li 2016). Aerosols that carrying SARS-CoV-2 can remain infectious for hours in the air, depending on temperature, humidity, and sunlight (Azuma et al. 2020; Dabisch et al. 2020; Fears et al. 2020; Schuit et al. 2020; van Doremalen et al. 2020). The susceptible individual who breathes in these infectious aerosols can become infected with COVID-19. Inhaled particles can deposit in the respiratory tract and cause infection. The infection risk depends on the quantity of virus inhaled and the site at which the virusladen particles deposit within the respiratory tract. Therefore, the viral transmission through the airborne route generally includes three phases, i.e. the release of respiratory particles, the transport and dispersion of aerosols in the indoor environment, and the ultimate exposure of the susceptible individual. Understanding the mechanism of airborne transmission

can be beneficial in reducing the infection risk in indoor environments.

0.000 (0%)

0.000 (0%)

0.672

Viral load in mouth

The viral load of the particles exhaled by the infector is closely associated with the infection risk of airborne transmission. However, it is still difficult to directly determine the viral load of the exhaled particles. Considering that exhaled particles are formed in the respiratory tract and released through respiratory activities, it is possible to derive the exhaled viral load based on the viral load in the respiratory tract, which can be detected more easily (Buonanno, Stabile, and Morawska 2020). Walsh et al. (2020) performed a detailed review on studies in viral load of SARS-CoV-2 and observed no significant difference of viral loads between asymptomatic and symptomatic patients. Evidence also suggests that the viral load of SARS-CoV-2 peaks around symptom onset or a few days thereafter, and becomes undetectable about two weeks after symptom onset. Higher viral load is likely associated with more severe symptoms (Walsh et al. 2020). Buonanno, Stabile, and Morawska (2020) presented an approach (Equation 1) to estimate the viral load emitted by a contagious subject based on the viral load in the mouth (or sputum), the type of respiratory activity (e.g., breathing, speaking, or coughing) and activity level (e.g., resting, standing, light exercise). The emission rate of virus quanta (ER) can be determined by the viral load in the sputum (c_v) , the conversion factor (c_i) , inhalation rate (IR), and total particle volume (relies on particle number N_D and volume V_D of each size in diameter D). One quantum in the model represents an infectious dose that would infect 63% of the population with the exposure, i.e. the infection dose ID63 (E. C. Riley, Murphy, and Riley 1978). The infectious quanta generation rate has a unit of quanta per hour (h⁻¹) and describes the number of infectious particles/pathogens in a way that implicitly includes both the number of virus particles generated in time and the infectivity of particles (which also inherently captures particle size effects and probability of deposition in appropriate regions of the respiratory system) (Sze To and Chao 2010).

$$ER = c_v \cdot c_i \cdot IR \cdot \int (N_D \cdot dV_D) \tag{1}$$

The viral load in sputum has been measured in some studies. Fajnzylber et al. (2020) measured 1.8–9.0 log₁₀

Table 3. Concentration and distribution of airborne SARS-CoV-2 across particle diameters.

	SARS-CoV-2 concentr	ration [RNA copies/m ³]	
Scenarios	Size distribution	Total	Reference
Protective-apparel removal room	0–0.25 μm: 0	42	Liu et al. (2020)
•	0.25–0.5 μm: 40		
	0.5–1 μm: 0		
	1–2.5 μm: 1		
	>2.5 μm: 1		
Protective-apparel removal room	0–0.25 μm: 1	20	
	0.25–0.5 μm: 2		
	0.5–1 μm: 9		
	1–2.5 μm: 1		
	>2.5 μm: 7		
Medical staff's office	0–0.25 μm: 3	20	
	0.25–0.5 μm: 2		
	0.5–1 μm: 2		
	1–2.5 μm: 3		
	>2.5 μm: 10	_	
Patient room with a symptomatic infector	<1 μm: 0	0	Chia et al. (2020b)
(sampled near the exhaust, >1 m from	1–4 μm: 0		
the patient)	>4 μm: 0	2224	
Patient room with a symptomatic infector	<1 μm: 0	3384	
(sampled within 1 m from the patient)	1–4 μm: 1384		
	>4 µm: 2000	10.42	
Patient room with an asymptomatic infector	<1 μm: 0	1843	
(sampled within 1 m from the patient)	1–4 μm: 916		
D (') () () () () ()	>4 μm: 927	1057	7 F (1 (2021)
Patient room with a symptomatic infector	<1 μm: 1112	1857	Z. Feng et al. (2021)
(sampled at the bedside, 0.2 m from the bed)	1–4 μm: 0		
Hospital corridor	>4 μm: 745	9	Zhou et al. (2021)
Hospital ICU room with infector(s)	/	21	Zilou et al. (2021)
Patient room with infector(s) (sampled	/	219	
at bedside)	1	219	
Two four-bed cohort bays with symptomatic	/	<10-460	Moore et al. (2021)
infectors (sampled <1 m from four			
different patients)			
Patient rooms with symptomatic infector(s)	/	$2.70 - 8.34 \times 10^3$	Santarpia et al. (2020)
(sampled > 1 m from the patient)			
Patient rooms with symptomatic infector(s)	/	$2.42-2.48 \times 10^3$	
(sampled >2 m from the patient)		2	
Patient room with symptomatic infector(s)	/	4.07×10^{3}	
(sampled near the patient)		2	
Hospital hallways	/	$2.08-8.69\times10^{3}$	
Personal air samples	/	$5.37-48.22 \times 10^3$	
Unventilated quarantine hotel toilet room	/	6.07×10^3	Ma et al. (2020)
Patient room with two infectors (sampled 2 m	/	74×10^3	Lednicky et al. (2020)
from patient 1 and 4.6 m from patient 2)	,	10 103	
Patient room with one infector (sampled 2 m	/	18×10^3	
from the patient)	,	3	
Patient room with two infectors (sampled 4.8 m	/	27×10^3	
from patient 1 and 3 m from patient 2)	ı	6 103	
Patient room with one infector (sampled 4.8 m	/	6×10^3	
from the patient)			

RNA copies/mL in sputum with a median level of 4.4 log₁₀ RNA copies/mL. Wölfel et al. (2020) measured the average viral load in sputum of 6.85 log₁₀ RNA copies/mL, with a maximum of 9.37 log₁₀ RNA copies/mL. J. Yoon et al. (2020) reported the viral load in the sputum of two patients between 4.86 and 8.22 log₁₀ RNA copies/mL. J. Y. Kim et al. (2020) detected the viral load in the sputum of the first two patients in South Korea roughly between 5 and 9 log₁₀ RNA copies/mL. Pan et al. (2020) collected sputum samples from 80 patients and observed a median viral load of 5.88 log₁₀ RNA copies/mL with a maximum of 11.13 log₁₀ RNA copies/mL. S. Zheng et al. (2020) collected 1846 respiratory samples (including sputum and saliva samples) and detected the viral load approximately between 3 and 8 log₁₀ RNA copies/mL. To et al. (2020) measured the peak viral load in the respiratory tract between 3.91 and 7.56 log₁₀ RNA copies/mL with a median of 5.29 log₁₀ RNA copies/mL for patients with mild disease, and the peak viral load between 4.27 and 7.40 log₁₀ RNA copies/mL with a median of 6.91 log₁₀ RNA copies/mL for patients with severe disease. Han et al. (2020) detected the viral load in the sputum of a patient roughly between 4 and 6 log₁₀ RNA copies/mL. It can be observed that the viral load in sputum can reach a level as high as 11 log₁₀ RNA copies/mL, while most of the existing data is approximately below 9 log₁₀ RNA copies/ mL. A recent study (J. Li et al. 2021) reviewed and analyzed the existing data of viral load in sputum in literature (397 samples) and observed a normal distribution of viral load with a mean value of 4.87 log₁₀ RNA copies/mL and a standard deviation of 1.90, indicating 99% of the viral load in sputum is below 9.30 log₁₀ RNA copies/mL. Some variants may have higher viral loads than the original strain. A study reported over 1000 times greater viral loads in oropharyngeal swabs for the Delta variant than the original viral lineage (B. Li et al. 2021).

Virus-laden particle concentration and size distribution Morawska et al. (2009) measured the size distribution of expelled aerosols during different expiratory activities (Table 2), including breathing, whispered counting, voiced counting, unmodulated vocalization, unmodulated whispering, and coughing. It recognized that most expelled droplets are in the smaller size, with diameters below 0.8 µm at average concentrations up to 0.75 cm⁻³ (accounting for around 70-80% of total particle numbers), followed by droplets at 1.8 µm at lower concentrations up to 0.14 cm⁻³ (around 10-20%), while few droplets near 3.5 μm and 5.5 μm generated (typically less than 6%). However, speech produced additional particles near 3.5 μm and 5.5 μm , at average concentrations of 0.04 cm⁻³ and 0.16 cm⁻³ (18% of total particles), respectively. Stephens (2013) conducted a systematic review for the particle size distribution of droplet nuclei and came up with a similar conclusion that the majority (typically 80-90%) of particles expelled during various human activities are smaller than 1-2 µm in diameter. It also suggested that the size distribution did not change significantly between activities, although coughing emitted a larger number of particles than breathing and talking, on average. Variations in size between age groups and gender groups were not significant either.

Although the majority of expelled droplets distribute in a smaller size range, droplets in smaller diameters may contain less virus due to the smaller particle volume, which means the pathogen in the air does not distribute in the same way as the size-resolved particle number distribution. When considering the droplet volume and droplet size distribution, larger droplets contain more pathogens, typically 80–90% for particles $>3~\mu m$ (J. Shen et al. 2021). It can also explain why some non-HEPA (i.e. high-efficiency particulate air) filters, such as filters with minimum efficiency reporting value (MERV) of 13, can still be effective in removing pathogens, as those filters are typically more effective in removing larger particles (J. Shen et al. 2021).

Some studies have conducted measurements of virus distribution in indoor environments rather than directly on exhaled droplets. Stephens (2013) reviewed such studies for influenza and estimated the size-resolved distribution of pathogen using the data in the literature (Lindsley et al. 2010), i.e. 15% of pathogens in the 0.3–1 µm size range, 25% in the 1–3 μ m size range, and 60% in the 3–10 μ m size range. A report from CDPH (2020) suggested the distribution of 20%, 30%, and 50% for infectious particles of SARS-CoV-2 in 0.3-1 µm, 1-3 µm and 3-10 µm, respectively. It was assumed that those indoor measurements account for a combination of all human respiratory activities that occur indoors (Stephens 2013). The published data regarding the measured viral concentration and size distribution of airborne SARS-CoV-2 aerosols are summarized in Table 3. The proportion of pathogens in larger droplets is lower than the proportion in expelled droplets, which is likely due to more deposition and evaporation for larger particles. The present of viral aerosols in protective-apparel removal room, staff's office and corridor suggested the possibility of infection beyond patient rooms in hospitals. Usually, higher viral concentrations appeared within closer distances to patients. Both symptomatic and asymptomatic patients can generate considerable viral aerosols.

Natural inactivation rate for infectious aerosols

Virus-carrying aerosols expelled by the COVID-19 patients have been detected to be infectious in some hospital settings (Lednicky et al. 2020; Santarpia et al. 2020). However, the field data are not designed to determine how long the virus can survive. Some experimental studies have also been conducted in laboratory environments to study the viability of virus-laden aerosols in the air under different environmental conditions. The infectivity of viral aerosols can typically remain for up to hours in the air under typical indoor environment conditions. But a rapid decay was observed when the aerosols were exposed to sunlight (Dabisch et al. 2020; Schuit et al. 2020). Temperature and humidity may also influence the persistence of viable SARS-CoV-2 in aerosols.

van Doremalen et al. (2020) tested the half-lives of SARS-CoV-2 in aerosols and estimated an approximation of 1.1 h (95% credible intervals (CI) 0.64–2.64 h) at 21–23 °C and 40% RH, which indicates an inactivation rate of 0.63 h⁻¹ (95% CI 0.26–1.08 h⁻¹). This value has been widely used in some modeling works (Buonanno, Morawska, and Stabile 2020; Buonanno, Stabile, and Morawska 2020; Rothamer et al. 2020). Fears et al. (2020)

observed no decay in virus-containing aerosol for 16h at 23 ± 2 °C and $53 \pm 11\%$ RH, indicating a nearly zero inactivation rate. Smither et al. (2020) suggested an average inactivation rate of $0.75\,h^{-1}$ in aerosols at $19-22\,^{\circ}\mathrm{C}$ and medium relative humidity (40–60%) in the dark, and 0.80 h⁻¹ at high relative humidity (68–88%) for the SARS-CoV-2 Alpha variant. Schuit et al. (2020) suggested that relative humidity alone did not affect the inactivation rate but can have minor interactions with other factors. The inactivation rate without sunlight across all relative humidity levels (20-70%) was $0.48 \pm 0.66 \,\mathrm{h}^{-1}$ (90% loss, 286 min). The natural inactivation rate under simulated sunlight levels representative of late winter/early fall and summer were $7.26 \pm 1.02 \,\mathrm{h^{-1}}$ (90% loss, 19 min) and $18.36 \pm 5.82 \,\mathrm{h^{-1}}$ (90% loss, 8 min), respectively. Dabisch et al. (2020) observed that temperature, sunlight, and humidity are all significant factors influencing the persistence of viable SARS-CoV-2 in aerosols, while sunlight and temperature have a greater influence on decay than humidity across the tested range (10-40 °C, 20-70% RH, 0-1.9 W/m² UVB irradiance). Higher temperature and sunlight intensity can generally increase the inactivation of viral infectivity. An average inactivation rate of 0.69 h⁻¹ was observed at the typical indoor temperature (20 °C) without sunlight. Elevated inactivation rates were observed at 20 °C under low-intensity (representative of late winter/early fall, 7.26 h⁻¹) and high-intensity (representative of summer, 18.09 h⁻¹) sunlight conditions.

The regression analysis between the number of confirmed cases and the ambient environmental conditions (i.e. temperature, RH, and ozone concentration) of Chinese cities between January and March in 2020, also supported the impacts of environmental conditions on virus infectivity (Yao et al. 2020). The result showed that the spread of SARS-Cov-2 was reduced by increasing ambient ozone concentration level from 48.83 to 94.67 μ g/m³, and decreasing relative humidity from 82.67 to 23.33% and temperature from 19 to -13.17 °C (Yao et al. 2020).

In conclusion, the typical inactivation rate of viable SARS-CoV-2 aerosols at typical indoor temperature (20–23 °C) and relative humidity (20–70%) without sunlight is generally between 0 and 1 h⁻¹. Sunlight can contribute greatly to the inactivation of the viability of SARS-CoV-2. The inactivation rate of the viable aerosols is generally over 7 h⁻¹ under low-intensity sunlight and over 18 h⁻¹ under high-intensity sunlight. It has the potential as a passive strategy for mitigating virus transmission indoors. Temperature and humidity may also affect the inactivation of the infectivity of virus-laden aerosols.

Transmissibility of SARS-CoV-2 variants

Genetic variants of SARS-CoV-2 have been emerging and circulating around the world throughout the COVID-19 pandemic (U.S. CDC 2021m). Evidence has shown that some variants are associated with an increase in transmissibility and more severe disease, e.g. increased hospitalizations or deaths, and may also reduce the effectiveness of previous treatment measurements (U.S. CDC 2021m). At the current stage, the main SARS-CoV-2 variants of concern (VOCs) include the Alpha variant (B.1.1.7, first detected in the U.K.), the Beta variant (B.1.351, first detected in South

Africa), the Gamma variant (P.1, first detected in Brazil), the Epsilon variant (B.1.427 and B.1.429, first detected in California) and the Delta variant (B.1.617.2, first detected in India). Studies have revealed that these variants generally have higher transmissibility than the original lineage, therefore, cause even more concerns about their spreading. The Alpha variant was observed 43 to 90% more transmissible than the predecessor lineage (N. Davies et al. 2021). The Beta variant was estimated to be 1.56 (95% CI 1.50-1.74) times transmissible as previously circulating lineages (Pearson et al. 2020). The Epsilon variant was detected and estimated to have an 18.6-24% increase in transmissibility (Deng et al. 2021). The latest and prevailing variant so far, the Delta variant, is responsible for over 83% of new infections in the U.S. as of July 2021 (Fry and Rapp 2021). It is highly contagious, nearly twice as transmissible as the original strain or previous variants (with an odds ratio of 1.88 (95% CI 0.95–3.76) compared with the original type (Ong et al. 2021) or an odds ratio of 1.64 (95% CI 1.26-2.13) compared to the Alpha variant (Hester Allen et al. 2021)). A recent report indicated that the viral loads of Delta infections were on average around 1000 times greater compared to the original strain of COVID-19 (B. Li et al. 2021). In many countries, the Delta variant has become the major challenge of COVID-19. Thus, the transmission of the airborne route for the Delta variant requires certain attentions and studies.

Outbreaks in different scenarios and the role of airborne transmission

Due to the huge difference in occupancy characteristics (e.g. density, age group, and behavior), room configurations (e.g. dimension and furniture layout), and ventilation settings (e.g. fresh and total supply airflow rate, air filtration, air distribution, and natural ventilation), the viral transmission in various indoor scenarios can vary greatly. The intervention for mitigating infection risk in various scenarios, therefore, can be very different. Public health agencies have provided guidelines for some key scenarios, including school, college, and workplace (U.S. CDC 2021a). Some reports have counted the COVID-19 cases in different scenarios. But the role of the airborne route in transmitting the virus in these scenarios has not been analyzed yet.

A report regarding the total COVID-19 outbreaks in Colorado since the pandemic revealed the proportion of outbreaks in various settings (data as of May 26th, 2021 (CDPHE 2020)). The documented outbreaks were associated with more than 86,000 infection cases, accounting for around 16% of total cumulative infection cases in Colorado. The majority of outbreaks were related to long-term care facilities, K-12 schools, restaurants and bars, retail facilities, and offices (Figure 2). In addition, the outbreaks in long-term care facilities (25 cases/outbreak on average), correctional facilities (123 cases/outbreak), colleges and universities (90 cases/outbreak), and transportations (24 cases/outbreak) typically infected more people. The data in Michigan (MDHHS 2020) only reported the new and ongoing

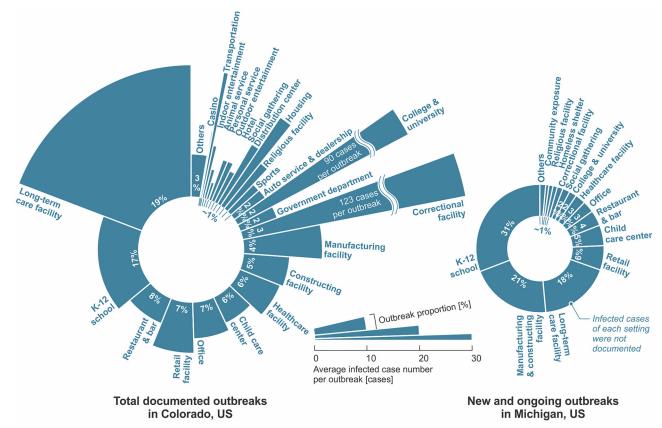


Fig. 2. COVID-19 outbreaks in various settings in Colorado (data as of May 26th, 2021 (CDPHE 2020)) and Michigan (data as of May 20th, 2021 (MDHHS 2020)).

outbreaks in different settings. The documented data showed that K-12 schools, manufacturing and constructing facilities, long-term care facilities, and retail facilities are facing higher risks (Figure 2). The definition of building/facility settings may have a few differences between various data. But it can be observed that long-term care facilities, manufacturing facilities, correctional facilities, K-12 schools, colleges, healthcare facilities and hospitals, retail facilities, restaurants, and offices are generally the scenarios with high outbreak probabilities, although the transmission modes of these documented cases were complicated. The data in Illinois (IDPH 2020) and Louisiana (LDHH 2020) revealed similar results. J. Shen et al. (2021) simulated the airborne transmission in various scenarios with typical settings and estimated that long-term care facilities, colleges, meat plants, hotels, restaurants, casinos, and cruise ships are facing considerably higher infection probabilities and have a higher potential to cause serious outbreaks or even superspreading events, which is generally consistent with the reported data. This section reviewed the hotspot scenarios of COVID-19 outbreaks based on the documented data and analyzed the role of airborne route in viral transmission.

Long-term care facility

Long-term care facilities provide a variety of services, both medical and personal care, to people who are unable to live independently, including facilities like nursing homes, and assisted living facilities (U.S. CDC 2019). There are

currently more than 5 million people who reside or work in long-term care facilities in U.S. (Chidambaram 2020). The COVID-19 has had a disproportionate effect on the people in long-term care facilities, which makes long-term care facilities the major risk for COVID-19 infection (The New York Times 2020a, 2020b). The fatality rate of the infection cases among nursing home residents in the U.S. is nearly 11% (U.S. CMS 2021) (by May 16th, 2021), far exceeding the average nationwide fatality rate of 1.8% (WHO 2021a). A worldwide study reviewed the fatalities of COVID-19 in a range of high-income countries and found that 46% of reported nationwide COVID-19 fatalities took place in nursing homes (data as of June 26th, 2020) (Comas-Herrera et al. 2020; Gregg Girvan 2020). In some countries, this proportion can even reach 80% (Comas-Herrera et al. 2020; Gregg Girvan 2020). In 13 states in the U.S., at least half of deaths are linked to nursing homes (The New York Times 2020b). The residents in long-term care facilities are typically elderly people (83.5% of residents in nursing homes are aged 65 and older (Harris-Kojetin et al. 2019)), who are probably more vulnerable to COVID-19 and may face more significant risks of developing severe illness once get infected (F. Li et al. 2021). Therefore, the transmission and prevention of SARS-CoV-2 in long-term care facilities require more attentions (D. C. Anderson et al. 2020).

The airborne transmission of SARS-CoV-2 in long-term care facilities has been suggested as a potential route.

Mouchtouri et al. (2020) detected viral RNA on the filter of the air-conditioner device in a patient room in a nursing home. It suggested that the viral aerosols expelled by the infected patients can be displaced by the airflow. de Man et al. (2020) suggested that an outbreak in a nursing home is caused by aerosol transmission in a situation of inadequate ventilation, considering the detected viral RNA in the HVAC system and the unfiltered air recirculation of the HVAC system. Besides, asymptomatic and pre-symptomatic transmission has been vastly reported in long-term care facilities (Dora et al. 2020; Kimball et al. 2020), which is supportive of the potential of airborne transmission (Greenhalgh et al. 2021).

A typical long-term care facility usually includes several single- or double-occupancy resident suites (including bedroom and bathroom) and other shared common areas, such as dining room, living room, activity room, or lounge. A nursing home may contain extra medical spaces. The size of a long-term care facility can vary from 20 or fewer residents to over 120 residents (National Institute on Aging 2017). A national survey for nursing homes in the U.S. suggested that the average number of beds per nursing home is around 107.6, of which an average of 86.3% are occupied (National Center for Health Statistics 2004), indicating an average of 93 residents per nursing home. The high occupancy and shared common areas elevate the occupant density in longterm care facilities. Considering the frequently reported asymptomatic cases (Dora et al. 2020; Kimball et al. 2020), the transmission through the airborne route in common areas is likely to happen if the space is insufficiently ventilated and the intervention measurements are not well implemented (de Man et al. 2020). J. Shen et al. (2021) modeled the infection risk through airborne route in a long-term care facility with typical settings and suggested that virus is likely transmitted via airflow in double-resident bedrooms and common areas such as dining room and therapy room during communal dining and group activities. U.S. CDC (2021i) suggested the residents wear PPE in common areas to control potential sources and recommended placing quarantined residents in single-resident bedrooms instead of double-resident rooms. Indoor ventilation should be improved to minimize the viral airborne transmission, e.g. increasing outdoor air intake of the HVAC system, and using a filter with MERV 13 or higher for recirculated air (Government of Canada 2021; MDH 2021). It is also recommended that the door to the room remain closed to reduce the transmission of SARS-CoV-2 through airflow between rooms (U.S. CDC 2021i). Lynch and Goring (2020) introduced practical steps to improve airflow to reduce the infection risk, using strategies adapted from negative-pressure isolation rooms in acute care facilities.

K-12 school

The COVID-19 outbreaks in K-12 schools have been vastly documented around the world (D. Lewis 2020b; Otte Im Kampe et al. 2020; Pray et al. 2020; Stein-Zamir et al. 2020). Previous studies indicated that few outbreaks were

reported in schools, and most infections reported in children were actually acquired at their homes (ECDC 2020; D. Lewis 2020b; WHO 2020g). Even in school outbreaks, it was more likely that the virus was introduced by the employees (ECDC 2020; WHO 2020g). Many studies have suggested that children are generally less susceptible and less infectious, and less likely to show severe symptoms than adults, particularly for younger ones (<10 years old) (ECDC 2020; Goldstein, Lipsitch, and Cevik 2020; Ludvigsson 2020; Viner et al. 2020; WHO 2020g). But recent research suggested that children can spread the emerging variants of SARS-CoV-2 as possibly as adults (D. Lewis 2021b). Therefore, although the fatalities among children and adolescents were generally lower than other populations, large outbreaks and severe diseases even deaths will still occur in schools, particularly considering the emergence of variants with higher transmissibility such as the Delta variant (Mallapaty 2020; Stein-Zamir et al. Szablewski et al. 2020; WHO 2020d). Data suggested that 31% of new and ongoing outbreaks in Michigan (MDHHS 2020) and 17% of total outbreaks in Colorado (CDPHE 2020) were tied to K-12 schools, which makes it another hotspot for virus spreading. But transmissions through other routes (contacts or droplets) cannot be ruled out based on existing data.

Evidence suggests that many K-12 schools that have strictly implemented prevention strategies have been able to safely open for in-person instruction and remain open (U.S. CDC 2021j). Previous studies in South Korea (Y. Yoon et al. 2020) and Spain (Catala et al. 2020) observed no significant increase in COVID-19 infection after schools reopened. Actually, school closure can introduce certain risks and negative impacts to students and their families, such as additional childcare expenses, and loss of access to education (Public Health Ontario 2020; United Nations 2020; WHO 2020g). WHO suggested that school closure should be considered only if there is no other alternative (WHO 2020g). Therefore, viral transmission in schools needs to be controlled and minimized to create a healthy and safe environment for students and instructors.

A study on 101 classrooms in 19 public schools in New York City estimated the airborne transmission risk and revealed that the probability of student-to-student transmission is lower than teacher-to-student transmission (Pavilonis et al. 2021), which is consistent with the reported situation (ECDC 2020; WHO 2020g) and other modeling results (J. Shen et al. 2021). The high occupant density, long exposure time, and insufficient ventilation are likely the main factors that enhanced the viral transmission through the airborne route. A previous review revealed that most schools did not provide sufficient ventilation in practice (Fisk 2017). Poor ventilation in classrooms can elevate the infection risk (Jones et al. 2021). Other common areas in schools like cafeterias, dining rooms, and gyms may face high risks as well (J. Shen et al. 2021). Therefore, prevention strategies should be strictly implemented in schools. Curtius, Granzin, and Schrod (2021) tested portable air cleaners in a school classroom and suggested that air cleaners can efficiently reduce

the airborne transmission risk for SARS-CoV-2. J. Zhang (2020) introduced multiple IAQ control strategies, including source control, ventilation, and disinfection, to reduce the risk of asymptomatic SARS-CoV-2 infections in classrooms. A. K. Melikov, Ai, and Markov (2020) presented an intermittent occupancy strategy combined with ventilation to reduce the airborne transmission risk in classrooms, which can reduce the risk significantly. Rothamer et al. (2020) suggested that viral particles can travel a long distance with the ventilation airflow in the classroom and also recommended using mask fitters to further improve the filtration efficiency of masks. Mask wearing and enhanced ventilation for reducing risk have been highlighted by U.S. CDC (2021j) as well. Besides, it is suggested to move the activities with high exhalation rates (such as singing, shouting, band, or sports and exercise) to larger and well-ventilated spaces or outdoors, when possible (U.S. CDC 2021j). The shared common areas, such as cafeterias, are suggested to be closed for communal use (U.S. CDC 2021j).

College and university

By the end of 2020, more than 1,800 colleges and universities in the U.S. have recorded over 397,000 COVID-19 confirmed cases, accounting for around 2.4% of the country's total confirmed cases (W. Cai et al. 2020) while students and employees in colleges and universities represent around 7% of the total population in the U.S. (Hussar et al. 2020; Univstats 2020). But few of these cases have resulted in deaths. Most of the college-related deaths were tied to staff or faculty members (Hubler and Hartocollis 2020). Many colleges and universities in the U.S. have reopened their campuses partially, but strict measures were taken to prevent the COVID-19 spreading on campuses, including online teaching, social distancing, mask-wearing requirement, partition installation, and ventilation system upgrading (U.S. CDC 2020b).

The strict prevention measures taken in studying areas on campus, such as classrooms, offices, and libraries, were believed to be efficient since the outbreaks that occurred in these spaces were rarely reported (Diep 2020). The outbreaks were more frequently reported during off-campus social gatherings, as well as in resident halls and athletic departments, where people were less likely to wear masks or keep physical distancing (Blinder, Higgins, and Guggenheim 2020; Diep 2020; Hubler and Hartocollis 2020; M. Lewis 2020; Teran et al. 2020; E. Wilson et al. 2020). Inadequate ventilation in these dense spaces enhanced cross-contamination through airborne transmission. It is consistent with the simulation result by Gressman and Peck (2020), indicating that the risk through academic contacts (directly tied to classes) is significantly lower than the risk through residential contacts and other large-scale social contacts. J. Shen et al. (2021) also indicated high airborne transmission risk in dining halls, gyms, resident halls, and Greek houses based on simulation, while the risk in studying areas is relatively lower. Therefore, the shared areas like dining halls, lounges, and gyms are suggested to be closed for communal use, if possible. Otherwise, strict prevention strategies should be implemented during gatherings in these scenarios. Masks are always required to wear. Ventilation system upgrades and improvements, e.g. increased outdoor air, elevated air supply, and higher-efficiency filters, are encouraged to increase the delivery of clean air and dilute potential contaminants in campus buildings (U.S. CDC 2020b). When the weather conditions allow, windows are suggested to open to introduce more outdoor air, incorporating with the use of fans to improve the effectiveness of opening windows (U.S. CDC 2020b). Air cleaning and disinfection strategies are also recommended to be used. When the ventilation is inadequate, the occupants in the room need to be decreased to maintain sufficient ventilation rate per person (U.S. CDC 2020b).

Manufacturing facility

The COVID-19 outbreaks in manufacturing facilities have been widely reported across the world, which makes it a hotspot of COVID-19 spreading (BBC News 2020a, 2020b; Guenther et al. 2020; Günther et al. 2020; Middleton, Reintjes, and Lopes 2020; Platsky et al. 2020; Richards and Vassalos 2020; Steinberg et al. 2020; Stewart, Kottasova, and Khaliq 2020; Waltenburg et al. 2020). According to the data in U.S., manufacturing facility has become one of the major risks for COVID -19 infection in the country throughout the pandemic (CDPHE 2020; IDPH 2020; MDHHS 2020). Particularly, a great number of outbreaks have been documented in meat processing plants, and some of them have been reported as superspreading events, which could get many hundreds of people infected (Waltenburg et al. 2020). For example, an outbreak among employees at a meat processing plant in South Dakota resulted in a total of 929 (25.6%) confirmed COVID-19 cases among 3,635 employees (Steinberg et al. 2020). At a meat processing plant in North Carolina, a total of 570 (25.4%) were tested positive among 2,244 employees (Tyson Foods Inc 2020). More than half of the 300 workers at a chicken plant in Delaware were confirmed as infected (Sternberg 2020). Among 1,216 employees working at a meat processing plant in Nebraska who were tested, 375 (31%) had positive results (Donahue et al. 2020). In July 2020, U.S. CDC reported more than 16,000 infections and 86 deaths tied to meat processing plants in U.S. (Waltenburg et al. 2020). But those numbers highly underestimated the actual condition since only 28 states provided data. According to the data of the last two weeks in 2020, around 20.8% of the reported cases in Louisiana were related to meat processing plants, which is a scenario with highest infection risk (LDHH 2020). But airborne transmission is not the only reason for these superspreading outbreaks as other transmissions cannot be ruled out.

Various reasons have been proposed for the outbreaks in meat processing plants, including the crowded working conditions, the cold working environment, the poor indoor ventilations, workers not wearing masks properly or not at all, and the need for workers to talk loudly over the background noise of machinery (Donaldson 2020; Nabarro, Deland, and

Lasbennes 2020; Stewart, Kottasova, and Khaliq 2020). Some studies believed that the frequent use of high-pressure water for brushing the floors and surfaces in meat processing plants would produce large quantities of droplets and aerosols, which may increase the risk of inhalation exposure if the floors and surfaces were contaminated by the virus (Donaldson 2020; Nabarro, Deland, and Lasbennes 2020). The COVID-19 outbreak in a German meat processing plant provides evidence that transmission occurred likely via airborne route over long distances (over 8 m) in the confined area (Günther et al. 2020). The environmental conditions in the meat plant, including low temperature, low air exchange rates, and constant air recirculation, together with the relatively close distance between workers and demanding physical work, likely created an unfavorable mix of factors promoting the efficient aerosol transmission of SARS-CoV-2 particles (Günther et al. 2020). It was believed that these factors are also responsible for worldwide ongoing outbreaks in other meat processing facilities (Günther et al. 2020). The high attack rate in meat plants was also supported by the modeling of airborne transmission (J. Shen et al. 2021).

To prevent the outbreaks in meat processing plants, prevention strategies are necessary, including reducing the occupant density, wearing masks properly, separating workers from each other to maintain social distancing, installing partitions between workers, introducing enhanced cleaning and disinfection strategies, and improving ventilation systems (U.S. CDC 2020f).

Retail facilities

Retail facilities usually include various categories, including grocery stores, supermarkets, home improvement retailers, and shopping malls. Many retail facilities, especially supermarkets, were not closed during the pandemic, but most of them shortened the working hours or limited the in-store customer numbers. The high volume and liquidity of in-store customers would make retail facilities vulnerable to COVID-19, particularly for the employees considering their long working time and close contacts with customers (Chang et al. 2020). Among 104 employees in a grocery store in Boston, 21 (20.2%) were tested positive, which is higher than the average risk in the community (Lan et al. 2020). In Colorado, around 7% of the infection cases were associated with retail facilities, of which most cases are the employees in the retail facilities (CDPHE 2020).

The airborne route is a possible route of virus transmission in retail facilities. A cluster of COVID-19 cases in a shopping mall in China was likely associated with the transmission through aerosols (Shao et al. 2021). Vuorinen et al. (2020) suggested that the airborne exposure during a supermarket visit even with relatively high customer densities would be quite low, partially due to the short residence time, which is consistent with the reported data (CDPHE 2020). However, considering a large number of supermarket visits daily, the number of exposed customers can become significant in large populations. It was also observed that the inhaled virus-carrying aerosols increase rapidly with the

residence time in the market. Therefore, for customers, it was recommended to limit the frequency and duration of visiting supermarkets, avoid visits during rush hours, and reduce the time spent at the cashier (Vuorinen et al. 2020). For employees, strict prevention strategies should be implemented to reduce the potential airborne risk.

Healthcare facility

Healthcare workers in healthcare facilities usually face higher COVID-19 exposure through their work since they are more likely to have close contact with the COVID-19 patients. The infection events among healthcare workers have been frequently reported across the world (Khurana et al. 2020; Koh 2020; Lai et al. 2020). Data suggested that over 328,000 healthcare workers in the U.S. have been infected by the end of 2020, accounting for over 17% of nationwide infection cases (U.S. CDC 2020a). Due to the significant gaps in data collection, this estimation probably underestimates the actual number of infected healthcare workers. Thus, viral transmission in healthcare facilities requires more attentions and studies. More efficient prevention strategies should be performed to reduce the high attack rate.

Healthcare facilities can be categorized into inpatient and outpatient healthcare facilities, typically including hospitals and various clinics (U.S. EIA 2020). Viral RNA has been vastly detected in the air of hospital settings, such as patient rooms, ICUs, restrooms, corridors, and even protectiveapparel removal rooms and medical staff's offices (Binder et al. 2020; Chia et al. 2020a; Ding et al. 2021; B. Feng et al. 2021; Z. D. Guo et al. 2020; Jiang et al. 2020; Jin et al. 2021; Kenarkoohi et al. 2020; Lednicky et al. 2020; Y. Li et al. 2020; Moore et al. 2021; Razzini et al. 2020; Santarpia et al. 2020; Stern et al. 2021; Tan et al. 2020; Zhou et al. 2021), and viable viruses were observed in some samples (Lednicky et al. 2020; Santarpia et al. 2020), indicating strong evidence for supporting the viral transmission through airborne route. The high contaminant concentrations were probably due to the high density of infectors in hospitals and the aerosol-generating procedures for treatment (e.g. intubation), which can generate considerable virus-carrying aerosols. Airborne precautions are recommended for circumstances and settings in which aerosol-generating procedures are performed (WHO 2020c). Airborne transmission is also possible to occur in some outpatient facilities, such as dental clinics, although no clear evidence has been reported (Melo et al. 2021). Dental procedures, such as the use of drills, saws and high-pressure water sprays, can generate considerable particles within close proximity of the patient, which puts the dentists at high exposure risks (Checchi et al. 2021). Masks and air disinfection strategies are suggested to prevent the potential airborne transmission.

Office

The COVID-19 outbreaks in office buildings have been reported across the world (CDPHE 2020; Loop News 2020; Park et al. 2020; Weissberg et al. 2020). Data suggested that

approximately 7% of the documented outbreaks in Colorado occurred in office scenarios (CDPHE 2020). Although the lack of clear evidence, the high attack rate in a call center in South Korea that caused 94 out of 216 employees infected (43.5%), can be partially explained by the transmission through airborne route (Park et al. 2020). It served to highlight that being in an enclosed space, sharing the same air for a prolonged period increases the risk of exposure and infection (ASHRAE 2021). Therefore, the virus spread in office settings is likely affiliated with the long working hours that employees spend together in the dense and confined space, where social distancing is less likely to maintain, and ventilation may be inadequate. In order to reduce the risk in office settings, enhanced ventilation, such as increased outdoor air, elevated air supply, and high-efficiency filter, are recommended to be implemented. Natural ventilation can further increase the delivery of clean air. Standalone air cleaning technologies like portable air cleaners or upper-room UVGI systems are also suggested. Besides, a staggering shift of working time that can reduce the number of employees in common areas like conference rooms and lounges is probably an efficient administrative control strategy for reducing airborne transmission (U.S. CDC 2021f).

Correctional facility

The COVID-19 outbreaks affiliated with correctional facilities have been widely reported across the world, which makes it another key hotspot of COVID-19 spreading (Council of Europe 2020; Davlantes et al. 2020; Hagan et al. 2020; Njuguna et al. 2020; Wallace et al. 2020a, 2020b). Data in the U.S. revealed that over 433,000 COVID-19 cases have been confirmed in 1,960 correctional facilities across the country by the end of 2020, accounting for approximately 22% of total infection cases in the U.S. (The New York Times 2020a; U.S. CDC 2020a). But the high attack rate of COVID-19 in correctional facilities cannot be attributed solely to airborne transmission.

The majority (82.4%) of the confirmed cases were the inmates (U.S. CDC 2020a) since they were essentially confined together in crowded spaces for a prolonged period. Superspreading COVID-19 outbreaks have been vastly reported in correctional facilities across the world (Swinkels 2020). For example, around 10% of inmates and 21% of staff in a correctional facility in Michigan tested positive for COVID-19 in June 2020 (Detroit Free Press 2020). Data in Colorado revealed that an average of 123 people can get infected in each outbreak in correctional facilities (CDPHE 2020). There is a high risk of rapid transmission in correctional facilities due to high occupant density and turnover, overcrowding, and frequent movements within and between establishments (Beaudry et al. 2020). In addition, a high proportion of inmates may be more vulnerable to severe disease due to underlying health conditions and generally older ages (Beaudry et al. 2020). A correctional facility typically consists of multiple housing units and common areas. A housing unit usually includes multiple sleeping areas (cells or rooms)

and an open dayroom, which is usually contiguous to the sleeping areas and inmates may be admitted to the dayroom for daytime activities such as dining, bathing, and selected recreation or exercise (U.S. NIC 2011). The modeling of airborne transmission revealed that multi-resident dormitories and dayrooms can readily spread the disease among inmates (J. Shen et al. 2021). Therefore, control strategies like masks, enhanced ventilation, regular cleaning and disinfection, and preventing overcrowding, are required for reducing the infection risk (U.S. CDC 2021g).

Hotel

U.S. CDC suggested a high risk staying in hotels or other multi-unit guest lodgings due to the potential contacts with other people (U.S. CDC 2020e). However, the COVID-19 outbreaks in hotels have not been frequently reported. It is likely because many consumers have yet to resume traveling due to the ongoing pandemic and relevant traveling restrictions, which had resulted in less than half the occupancy rate in most hotels compared to a normal condition, particularly for hotels in urban areas (AHLA 2020).

Viral RNA was detected in the air in an unventilated toilet of a quarantined hotel guest room in China (Ma et al. 2020). Besides, a retrospective study (Eichler et al. 2021) suggested that aerosols were likely the mode of transmission in a quarantined hotel in New Zealand, and the enclosed and unventilated space in the hotel corridor probably facilitated the spread. Therefore, airborne transmission is a potential route for SARS-CoV-2 spreading in hotels. Except for the guest rooms, the common areas of the hotel, such as the lobby and banquet rooms, are also facing high airborne transmission risk (J. Shen et al. 2021). Strict prevention strategies should be implemented in these common areas.

Restaurant and bar

There have been quantities of COVID-19 cases reported in restaurants, bars, or coffee shops (ABC News 2020; Y. Li et al. 2020; Lu et al. 2020; NBC News 2020). A study found that people who tested positive for COVID-19 are more likely tied to dining in restaurants, bars, or coffee shops before becoming ill, which suggests higher risks in these places (Fisher et al. 2020). Data revealed that 8% of the outbreaks in Colorado (CDPHE 2020) were reported in restaurants and bars. Around 19.3% of the COVID-19 outbreaks reported within the last two weeks of 2020 in Louisiana were tied to restaurants or bars, which is the place with the second primary risk in the state (LDHH 2020). Therefore, restaurants and bars have become a hotspot of COVID-19 outbreaks. Many restaurants and bars have fully or partially reopened at the current stage. Although prevention measures may have been taken, the dense customers, close contacts, and unabundant ventilation in restaurants and bars will still likely elevate the infection risk, particularly considering that people are unlikely to wear masks when they are dining or drinking. On-site dining with indoor seating has been evaluated as higher and highest risks by U.S. CDC (2021c).

Food services like a drive-through, delivery, take-out, and curb-side pick-up, are still preferred.

Airborne transmission has been suspected to be affiliated with the outbreaks in a restaurant in Guangzhou (Y. Li et al. 2020) and a bar in Ho Chi Minh City (Chau et al. 2021). The insufficient clean air delivery and recirculated contaminated air increased the infection risk through airborne transmission. Masks, ventilation improvements, air cleaning, and disinfection technologies are required to be used in restaurants and bars. Partitions are also suggested to be installed, particularly in areas where it is difficult for individuals to remain at least 2 m apart (U.S. CDC 2021c).

Religious place

Places in which people gather for religious activities, such as chapels, churches, mosques, synagogues, and temples, are also facing high risks of COVID-19 spreading. The COVID-19 infection has been widely reported in these places (Conger, Healy, and Tompkins 2020; James et al. 2020; Lee et al. 2020). Around 8% of the COVID-19 outbreaks reported within the last two weeks in 2020 in Louisiana were related to religious places (LDHH 2020). The crowded people in the confined religious building with poor ventilation are exposed to high risks, particularly considering the elevated respiratory rates during praying and singing.

Some outbreaks during activities in religious places have been suspected to be associated with the transmission through aerosols. Miller et al. (2020) revealed that the transmission by the aerosol route is likely to occur in the Skagit Valley Chorale superspreading event, as either fomite or droplet-borne transmission could unlikely explain the substantial proportion of the cases in the choir. Katelaris et al. (2021) suggested airborne transmission as the probable route for virus spreading during church singing in a choir in Sydney because some infected individuals seated up to 15 m from the index patient without close physical contact. Therefore, it is encouraged to use masks and intensify cleaning, disinfection, and ventilation in religious places to mitigate the airborne transmission risk (U.S. CDC 2021b).

Transportations

A study suggested that transport-based COVID-19 outbreaks are the second dominant category of outbreaks, behind the building-based ones (Qian et al. 2020). The outbreaks tied to transportation spaces, such as airplanes, cruise ships, and tour coaches, have been widely reported, particularly at the early stage of pandemic (before traveling restrictions) (ASHRAE 2020c; Kasper et al. 2020; Khanh et al. 2020; Luo et al. 2020; Moriarty et al. 2020; Y. Shen et al. 2020; Xu et al. 2020; S. Zhao et al. 2020; R. Zheng et al. 2020). A study found that the modes of transportation that keep passengers in the same confined cabin over a long period, such as tour coaches or airplanes, could spread the virus significantly (Sam Schwartz Consulting 2020). But in spaces where passengers spend a short period and tend not to talk, such as the mass transit systems, risks are relatively low, as long as the spaces are well-ventilated, and passengers wear masks properly. However, if prevention measures like a mask, social distancing, or enhanced ventilation are not taken, the mass transit systems, such as the subway and bus, will readily spread the virus as well. A study suggested that the mass transit system was a major disseminator of SARS-CoV-2 infection in New York City during the initial takeoff of the pandemic (Harris 2020). The major hotspots of transportation reported during the pandemic generally include aircraft, cruise ships, buses (e.g. transit buses, tour coaches, and school buses), subways, and cars.

Airborne transmission of the virus is less likely to occur on aircraft, considering the high ventilation rate and highefficiency filters for recirculating air (Gupta, Lin, and Chen 2012). According to a survey on aircraft HVAC systems, the typical air exchange rate in the aircraft cabin during cruising is approximately between 20 and 25 h⁻¹ (National Research Council 2002). For most modern aircraft, the air passes through HEPA filters before reentering the passenger cabin (WHO 2020a), which can remove at least 99.9% of particles (ASHRAE 2017). However, if the airflow is not well organized, the air in the aircraft cabin can be greatly mixed, which may cause cross-contamination and increase the exposure risk, particularly considering the high passenger density in the cabin (Gupta, Lin, and Chen 2012; Poussou et al. 2010). Therefore, airborne transmission is still a potential route for SARS-CoV-2 spreading on aircraft. For example, transmission through aerosols was likely occurred in the business class area during a long-haul flight to Vietnam, where seats are already more widely spaced than in economy class, and infection spread much further than the 2-row or 2-meter rule recommended for COVID-19 prevention (Khanh et al. 2020). Studies suggested that optimized ventilation systems, such as displacement ventilation, personal ventilation, and local air exhaust, can efficiently reduce the potential exposure risk in the cabin (Dygert and Dang 2012; You et al. 2019). Masks are always required for better protection. In addition to aircraft cabins, air traveling passengers also need to spend considerable time in security lines and airport terminals during stops or layovers, which may bring passengers in close contact with other people. This may elevate the risk during air travel. Viral RNA was detected in the air of some airports (U.S. CDC 2021j). Therefore, it is recommended to take flights with the fewest stops or layovers, if possible (U.S. CDC 20211).

Cruise ships were a hotspot of COVID-19 outbreaks in the early days of the pandemic. The outbreaks on the Diamond Princess cruise ship and the Grand Princess cruise ship caused more than 800 people infected in total (Moriarty et al. 2020). It is believed that cruise ships bring diverse populations into proximity for many days, facilitating the transmission of SARS-CoV-2. The gatherings in common areas on the cruise ship, such as bistros, cafeterias, ballrooms, casinos, and theaters (Princess Cruises 2020), may exacerbate the viral transmission among the passengers (Xu et al. 2021). A study revealed that aerosol inhalation was likely the dominant contributor to viral transmission on the Diamond Princess cruise ship, even considering a conservative assumption of high ventilation rates and no air

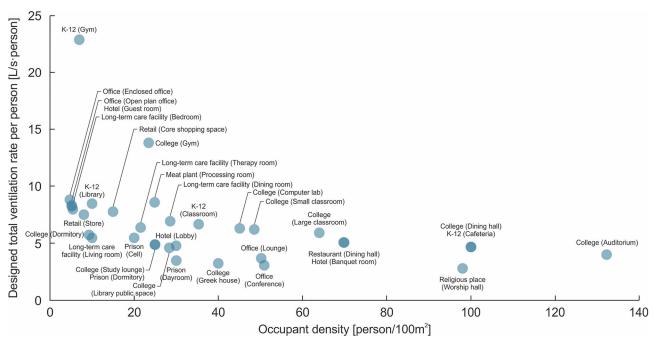


Fig. 3. Relationship between occupant density and designed total ventilation rate per person in different scenarios based on ASHRAE 62.1 (data adapted from J. Shen et al. (2021)).

recirculation for the cruise ship (Azimi et al. 2021). The higher-than-expected virus spread into cabins without previously confirmed cases during the quarantine period on the Diamond Princess cruise ship can be explained by the airborne transmission through the ventilation system (Almilaji and Thomas 2020), although another study suggested that the central ventilation system of cabins did not assist in virus transmission between cabins during the quarantine period as no cross-room transmission observed and most infections occurred before quarantine (Xu et al. 2021). Considering the high infection risk on cruise ships, it is recommended that all people should avoid travel on cruise ships, including river cruises, worldwide (U.S. CDC 2021o). Intervention in common areas on cruise ships, such as enhanced ventilation, is required for mitigating infection risks.

Airborne transmission can likely explain the markedly high attack rate of COVID-19 on the buses (tour coach and minibus) in Hunan (Luo et al. 2020) and Zhejiang (Y. Shen et al. 2020) in China. Besides, viral RNA was tested in airborne aerosols collected from transit buses (Hadei et al. 2021; Moreno et al. 2021). Therefore, the passengers on the buses, including tour coaches, transit buses, and school buses, may face a certain risk of exposure to the virus. Generally, a tour coach likely has a higher risk due to the long traveling time, while a transit bus and a school bus have a relatively lower risk because of the typically shorter commuting time (J. Shen et al. 2021). Enhanced ventilation is suggested on the buses as it can greatly reduce the risk (Moreno et al. 2021). Air supply mode plays an important role in airflow pattern and virus-carrying particle dispersion in the bus (Yang et al. 2020). Backward supplied air is more

suggested for reducing the transmission of viral droplets (Yang et al. 2020). Windows can be open to improve the incabin ventilation when the weather conditions allow (U.S. CDC 2021j), although its benefit does not uniformly impact all passengers on the bus (depending on the exact airflow patterns) (Z. Zhang et al. 2021). Masks are always recommended when using public transit. Passengers are not suggested to sit at nonadjacent seats (Yang et al. 2020). Besides, passengers should avoid eating or drinking on public transit as it requires people to remove their masks for a while (U.S. CDC 2021k). In addition, people are suggested to reduce the frequency of using public transit like buses.

Airborne viral RNA was detected in both subway stations and trains (U.S. CDC 2021j). However, airborne transmission is less likely to occur on subway trains because the ventilation system can usually supply considerable clean air to the passengers (J. Shen and Gao 2019) and people typically spend a short commuting time on subway trains. But the air mixing in the train may result in cross-contamination and increase the exposure risk. Passengers should avoid taking public transits and always ensure mask use during the entire trip.

Travels on personal cars, taxis, and rideshares can have a high exposure risk to SARS-CoV-2 due to airborne transmission. Infectious virus was detected in airborne particles of the size range of 0.25 to 0.5 µm in a car driven by a COVID-19 patient (Lednicky et al. 2021). The compact space of the car cabin with dense passengers can contaminate and mix the air easily. The passengers can be exposed to highly contaminated air. Therefore, masks are necessary for reducing exposure. It is also suggested to improve the ventilation in the vehicle if possible, e.g. opening the windows or setting the air conditioning on non-recirculation mode (U.S.

Table 4. Quanta emission rates (ER) of SARS-CoV-2 particles in different studies.

	,		
Activity	ER $[h^{-1}]$	Introduction	Reference
Estimated from retrospective analysis on real outbreak events	s on real outbreak ev	ents	
Standing + singing Standing + singing	970 ± 390 341	Skagit Valley Chorale superspreading event Skagit Valley Chorale superspreading event	Miller et al. (2020) Buonanno, Morawska, and
Standing + singing Seated + vocalization	870 61	Skagit Valley Chorale superspreading event Guangzhou restaurant outbreak event	Stabile (2020) Bazant et al. (2021) Buonanno, Stabile, and Moreoveka (2020)
Resting + breathing Resting + breathing Resting + breathing Resting + breathing	45 30 29 185.63	Zhejiang tour coach outbreak event Diamond Princess cruise ship outbreak event Wuhan city outbreaks Diamond Princess cruise ship outbreak event	Bazant et al. (2021) Bazant et al. (2021) Bazant et al. (2021) Chen et al. (2021)
Estimated using the viral load model			
Resting Intermediate Light activity + vocalization	<pre></pre>	Estimated based on the viral load in the sputum	Buonanno, Stabile, and Morawska (2020)
Resting + breathing/whispering ^a Standing + breathing/whispering Light activity + breathing/whispering Resting + speaking ^b Standing + speaking Light activity + speaking	3 3 3 50 56 142°	Estimated based on $c_{\nu}=10^9$ RNA copies/mL and $c_{i}=0.02$	Buonanno, Morawska, and Stabile (2020)
Estimated using statistical ineurous Sedentary state	14-48	Estimated based on the fitting curve between ER and R_0 from the data of other respiratory diseases (e.g. influenza and SARS-CoV-1)	Dai and Zhao (2020b)

^aBreathing/whispering represents the mean value between whispering and breathing (Buonanno, Stabile, and Morawska 2020).

^bSpeaking is considered as the mean value between unmodulated vocalization and voiced counting (Buonanno, Stabile, and Morawska 2020).

^cThis value was adopted by many studies (Harrichandra, Ierardi, and Pavilonis 2020; S. Zhang and Lin 2020).

CDC 2021k). The best configuration is to have all windows fully open if possible. If this is not practical, it would be good to have two windows open, preferably one in the rear and one in the front (Mathai et al. 2021). The transmission through aerosols in surface parking lots is likely very low as they are outdoor environments unless close contact with the infected person exists. But the airborne transmission is likely to happen in underground parking spaces equipped with jet fans (Nazari et al. 2021). The areas closer to the fresh air supplier are safer than other areas. Equipping UVGI systems and HEPA filters inside the jet fans can eliminate the viruses.

The above analysis of the outbreaks in different indoor scenarios shows that airborne transmission has likely played a major role. The infection risk due to airborne transmission in various scenarios depends on the occupancy characteristics, room configurations, ventilation settings, air distributions, and interventions. Interventions that reduce airborne transmission would be effective in reducing the infection risk. Indoor ventilation is a key factor impacting infection risk. The vastly used indoor ventilation design guideline, ASHRAE 62.1 (2019), regulates the minimum ventilation rates of different building settings based on the estimation of required ventilation per person and floor area. Therefore, when ASHRAE standard 62.1 is used, it penalizes smaller rooms so that high occupant density per floor area means less ventilation comparing to larger rooms (with the same occupants). Figure 3 shows the total ventilation rate per person estimated from ASHRAE 62.1 in typical scenarios with different occupant densities (J. Shen et al. 2021). Ventilation delivered to each person in scenarios with higher occupant density is generally lower than the ventilation in scenarios with lower occupant density. Therefore, the ventilation designed based on ASHRAE 62.1 may not always be sufficient for occupants, particularly considering the requirements for mitigating the infection risk of SARS-CoV-2. To address this issue, a pathogen-source-based or health-based design criteria for indoor ventilation is probably more applicable for infection prevention. More efforts are required for indoor ventilation design to protect against infection transmission (Morawska et al. 2021).

Model for estimating the risk of airborne transmission

Estimating the risk of airborne transmission in indoor environments is essential for designing control strategies to reduce the risk. The most commonly used model for estimating the airborne transmission risk in a confined space is the Wells-Riley model, which assumes the air in the space is well-mixed in steady-state (E. C. Riley, Murphy, and Riley 1978; Wells 1955). The model has also been extended by some researchers to include unsteady exposure (Gammaitoni and Nucci 1997) and imperfect mixing (Ko et al. 2001, Ko, Thompson, and Nardell 2004), and incorporate with stochastic modeling (Noakes and Sleigh 2009). The infection risk determined by the Wells-Riley model depends on the

susceptible individual's inhalation exposure to the suspended pathogen generated by the infectors (Stephens 2013; Sze To and Chao 2010).

The model has been widely adopted to estimate the airborne transmission risk of many respiratory diseases, such as influenza, tuberculosis, severe acute respiratory syndrome (SARS-CoV-1), middle east respiratory syndrome (MERS), and measles (S. C. Chen and Liao 2008; Qian et al. 2009; Stephens 2013; Yates et al. 2016; Zemouri et al. 2020). Recent studies also applied the Wells-Riley model in estimating the airborne transmission risk of SARS-CoV-2 in indoor environments. For example, Dai and Zhao (2020b) used the model to estimate the required ventilation rate in four typical scenarios ensuring a less than 1% infection probability. Harrichandra, Ierardi, and Pavilonis (2020) estimated the airborne SARS-CoV-2 infection transmission risk in New York City nail salons using the Wells-Riley model and evaluated the impacts of wearing face masks on the risk. J. Shen et al. (2021) applied the model in evaluating the risk in 18 scenarios with typical settings and observed that long-term care facilities, colleges, meat plants, hotels, restaurants, casinos, and cruise ships have higher infection risks. Pavilonis et al. (2021) estimated the aerosol transmission risk of SARS-CoV-2 in New York City public schools during the reopening and revealed that the risk of transmission is higher during the heating season and in newer buildings. Stabile et al. (2021) estimated the airborne transmission risk in schools and proposed a novel feedback control strategy using carbon dioxide concentrations to monitor and adjust the ventilation procedure for minimizing the airborne transmission. Peng and Jimenez 2021) developed the original model and derived analytical expressions of CO2-based risk proxies for estimating COVID-19 infection risks, and applied them to various typical indoor environments.

The Wells-Riley model estimates the risk with the wellmixing assumption. However, the air in indoor environments is typically imperfectly mixed. Close contact with the infected persons can typically inhale more virus-carrying aerosols, while a longer distance from the infectors can likely decrease the exposure due to the dilution by the ventilation and the deposition and natural inactivation of the viral particles. But the air recirculation in space can take the virus-laden aerosols to farther locations (Y. Li et al. 2020). Therefore, the airflow patterns in the space, as well as the locations of the infected and susceptible subjects, can significantly impact the spatial distribution of viruses. To address the spatial distribution of exposure risk, the rebreathed fraction can be incorporated in the Wells-Riley model, which is the fraction of inhaled air that was exhaled by the infector (Rudnick and Milton 2003). A similar approach was also conducted by other researchers for estimating the infection risk of COVID-19 (J. Shen et al. 2021; Sun and Zhai 2020). When the quantity of exhaled aerosols generated by the infectors and inhaled by a susceptible person in a particular spatial location is known, the susceptible person's exposure risk can be estimated based on the Wells-Riley model (Sze To and Chao 2010). It can be done by

Table 5. IAQ control strategies for mitigating the risk of airborne transmission.

Category	Strategy	Effectiveness	Effective scale	Capital cost ^a	Durability	Accessibility
PPE	Cloth mask	30% particle filtration efficiency for purchased masks (considering leakage from gaps caused by improper fit) (Konda et al. 2020); 20% particle filtration efficiency for homemade masks (cotton and nylon materials) (M. Zhao et al. 2020)	Breathing zone	US\$3/count (purchased); US\$0/count (homemade)	Reusable	Easy ^b ; Homemade (using household materials like cotton and nylon (A. Davies et al. 2013)); Comfort should be considered
	Surgical mask	50% particle filtration efficiency (considering leakage from gaps caused by improper fit) (Konda et al. 2020; Rothamer et al. 2020)	Breathing zone	US\$0.2/count	Disposable ^c	Depending on supply capacity (including conventional, contingency, and crisis situations (U.S. CDC 2021d)); Comfort should be considered
	N95 mask	filtration efficiency ps) ^d (Konda	Breathing zone	US\$2/count	Disposable ^c	Depending on supply capacity (including conventional, contingency, and crisis situations (U.S. CDC 2021d)); Comfort should be considered;
	Face shield	23% reduction on aerosol inhalation under long-time use (more aerosol reduction under short-distance use) (Lindsley et al. 2014)	Breathing zone	US\$2/count	Reusable	Easy; Homemade (using plastic sheets and other materials);
	Mask fitter/ sealer/brace ^e	Greatly improving the efficiency Breathing zone of masks by reducing the leakage through gaps (e.g. improve the particle filtration efficiency of surgical masks to over 90% (Rothamer et al. 2020))	Breathing zone	US\$9/count (purchased (UW Madison 2020)); US\$0/count (homemade)	Reusable	Easy; Homemade (using rubber or other materials (Fix The Mask 2020)); Increased discomfort
Ventilation	Upgrading filters of building ventilation systems (e.g. HEPA)	99.9% removal efficiency for particles >0.3 µm (HEPA filter)	Building	US\$280/count (with an increase in fan power due to the increased pressure rise through higher-rating filters (Risbeck et al. 2021))	Replaced every 6–12 months	Easy
	Increasing outdoor air supply of building ventilation	Largely increasing the ventilation rate of the room (e.g. providing 60% more clean air compared to a reference system with 25%	Building	US\$5/m²-year for extra energy consumption (central system); US\$10/m²-year for extra energy consumption	Long-term use ^f	Easy (switching the damper to increase outdoor air supply)
						(Continued)

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Category	Strategy	Effectiveness	Effective scale	Capital cost ^a	Durability	Accessibility
	systems (e.g. 100% outdoor air)	outdoor air and MERV 8 filter)		(individual unit) ^g (U.S. EIA 2018)		
	Personal ventilation (for fresh	Increasing clean air supply to the personal microenvironment with	Personal environment	Varying greatly (depending on system scale)	Long-term use (filters need regular	Need professional design and installation
	air supply)	typical ventilation effectiveness of 1.7, depending on diffuser location			replacements)	
		and airnow rate (A. Melikov, Grønbaek, and Nielsen 2007)				
	Local	Reducing the cross-	Personal .	Varying greatly (depending	Long-term use	Need professional design and
	air exhaust	contamination with typical ventilation effectiveness of	environment	on system scale)		ınstallatıon
		1.4 to 10, depending on the local partition configuration,				
		exhaust location, and airflow rate (Dygert and Dang 2012)				
	Displacement ventilation	Improving the clean air delivery	Room	Varying greatly (depending	Long-term use	Need professional design and
	VOITINIALIOII	ventilation effectiveness of		on system searc)	regular	шэганацоп
		Glicksman 2003)			repracements)	
	Natural ventilation	Providing more outdoor air to the room, e.g. $> 10 h^{-1}$ air	Room	0 \$ \$0	Long-term use	Depending on weather and outdoor air quality conditions
		changes in some scenarios (Qian et al. 2010) (depending				
		on window opening				
		conditions, while condition, and building geometry) ^h				
	Closing doors between	Reducing cross contamination	Building/room	US\$0	Long-term use	Easy (need to coordinate with
	rooms	contamination risk can be				
	(separating rooms and	fully closed)				
	blocking airflow	•				
	across rooms)			,		
Partition	Partition screens	Depending on airflow patterns in the room (well-designed partitions can reduce the risk of cross-contamination while	Personal environment	US\$100/m² material	Long-term use	Easy
		bad designs may increase the				

Need professional design and installation (incorporating with ventilation)	Need professional design and installation (incorporating with ventilation)	Need professional design and installation (incorporating with ventilation)	Easy; Potential emissions should be considered (some air cleaners can generate hazardous emissions like ozone (Guo, Gao, and Shen 2019))	Depending on weather conditions and building design ^k	Easy (need to coordinate with occupant schedules)	(Continued)
Long-term use	Long-term use	Long-term use	Long-term use (filters need regular replacement)	Long-term use	Long-term use	
US\$2000 for a 2×2 m cubicle	US\$3000 for a 2×2 m seat Long-term use	US\$1200–2500 (US\$40–90/m² floor area) (VirusLights 2021)	US\$493/count (i.e. US\$1.32 per m³/h CADR, median price from a market survey (B. Zhao, Liu, and Chen 2020)) (less than US\$50/count for HEPA filter replacement)	US\$0j	US\$01	
Personal environment	Personal environment	Room	Room	Room	Building/room	
risk in some locations) (Rooney et al. 2021) Reducing cross-contamination between cubicles with typical ventilation effectiveness of 1.1 to 3.6 (require well- designed ventilation and air distribution) (Haghighat et al. 1996)	Enclosed modular office walls can fully avoid cross-contamination between modular spaces (require sufficient ventilation supplied to each space)	Proper use can largely improve the inactivation rate of virus (e.g. 16 h ⁻¹ for M. parafortuitum) ¹	Typically supplying 361 m ³ /h CADR (median CADR from a market survey (B. Zhao, Liu, and Chen 2020))	Increasing the natural inactivation of SARS-CoV-2 on aerosols (decay rate: <1 h ⁻¹ without sunlight, >7 h ⁻¹ with low-intensity sunlight, >18 h ⁻¹ with high-intensity sunlight)	Reducing the potential emissions of pathogens to the air and reducing the risk of cross-contamination	
Cubicle workstation	Enclosed/ semi- enclosed modular office walls	Upper-room UVGI system	Portable air cleaners	Sunlight	Occupancy density restriction (limiting occupant number and social distance in the building/room ^m)	
		Air cleaning and disinfection			Occupancy control	

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Accessibility	Easy (need to coordinate with occupant schedules)
Durability	Long-term use
Capital cost ^a	0\$\$0
Effective scale	Building/room
Effectiveness	Reducing the potential emissions of pathogens to the air and decreasing the pathogen inhalation by the susceptible people (e.g. asking students to leave the classroom during the 15 min break after a 35 min class can reduce 35% inhaled pathogen (A. K. Melikov, Ai, and Markov 2020))
Strategy	Intermittent occupancy (A. K. Melikov, Ai, and Markov 2020) or staggered scheduling (U.S. CDC 2021j)
Category	

^{&#}x27;Capital costs are approximately estimated based on a brief market survey in the U.S. (data obtained from search results on Google). The actual costs may vary a lot in

The strategy can be accessed easily through in-store/online purchase or easy operation.

^cSurgical and N95 masks are designed for disposable use but can be reused a couple of times after decontamination under certain conditions (U.S. CDC 2021e). ⁴N95 masks can generally fit better than cloth and surgical masks. Proper use of N95 masks can minimize air leakage through gaps.

Other strategies that can maximize fit for surgical masks include (1) covering the surgical mask with a cloth mask, and (2) knotting ear loops of the surgical mask, which can improve the particle filtration efficiency to around 80% (Brooks et al. 2021).

Long-term use indicates a duration of typically over one year.

^gThe annual average cost of air conditioning with central and individual units in the U.S. is US\$0.31 and US\$0.15 per square foot, respectively (U.S. EIA 2018). Assuming the outdoor air fraction of the HVAC system is 25%, applying 100% outdoor air will simply cost 3× more energy for pre-heating or pre-cooling the extra outdoor air.

^hlt can improve the natural ventilation efficiency and introduce more outdoor air to the room incorporating fans.

The effectiveness of virus inactivation by the upper-room UVGI system also relies on room air distribution. Displacement ventilation may reduce the efficiency as it decreases the residence time of the virus in the irradiated zone (Kanaan and Abou Moughlbay 2018)

Introducing more sunlight to the room may impact the heating and cooling energy consumption for the HVAC system. 'Building design like window/wall ratio and building geometry can impact the incident sunlight.

The potential economic loss due to occupancy restriction is not considered.

mThe safe social distance is recommended as 1.6–3.0 m considering the aerosol transmission of exhaled large droplets from talking (Sun and Zhai 2020)

conducting the tracer gas measurements. Some other studies performed CFD simulations to figure out the spatial concentration distribution of viruses and incorporated them with the Wells-Riley model to estimate the spatial exposure risk distribution in the space (Gao et al. 2008; Qian et al. 2009; Tung and Hu 2008). CFD simulations have been performed in scenarios like restaurant, classroom, grocery store, bus, car, elevator, hospital ward, and underground parking space for estimating the spatial risk distribution of SARS-CoV-2 (Dbouk and Drikakis 2021; Y. Guo et al. 2021; Y. Li et al. 2020; H. Liu et al. 2021; Mathai et al. 2021; Nazari et al. 2021; Shao et al. 2021; Yang et al. 2020; Z. Zhang et al. 2021).

Wells-Riley model

The Wells-Riley model is a classical model for estimating the steady-state infection risk through airborne transmission in the perfectly mixed space (E. C. Riley, Murphy, and Riley 1978; Wells 1955). The infection possibility (P), which can be used to estimate the new infection cases (N_C) based on the susceptible cases (N_S), is calculated as a function of the inhalation exposure dose (E. C. Riley, Murphy, and Riley 1978), which depends on the number of infectors (I), the emission rate of infectious quanta per infector (ER), the fraction of infectious particle penetration through the face mask (R), inhalation rate (IR), exposure time (I) and the room ventilation rate with clean air (I):

$$P = \frac{N_C}{N_S} = 1 - e^{-R\frac{J-ER-JR-t}{Q}} \tag{2}$$

One quantum in the model represents an infectious dose that would infect 63% of the population with the exposure per the Wells-Riley model (E. C. Riley, Murphy, and Riley 1978). It describes the number of infectious particles in a way that implicitly includes both the number and the infectivity of virus particles (which also inherently captures particle size effects and probability of deposition in appropriate regions of the respiratory system) (Stephens 2012). The quanta emission rate has a unit of quanta per hour (h⁻¹), and the magnitude relies on disease species, infector activities (e.g. breathing, coughing) and interventions, and may vary significantly case by case (Buonanno, Stabile, and Morawska 2020; Stephens 2012).

The quanta emission rate can be estimated based on the viral load in the mouth of the infected person (Buonanno, Stabile, and Morawska 2020), which can be calculated through Equation 1. Another approach to estimating the quanta generation rate is through the retrospective analysis of real outbreak events. The quanta generation rate can be backward calculated when the attack rate, occupant activities, room configurations, and ventilation settings of the studies outbreak are known. In addition, statistical analysis was performed to estimate the quanta generation rate of SARS-CoV-2 based on the statistical relationship between the basic reproduction number (R_0) and ER of other respiratory diseases. The estimated quanta generation rates of SARS-CoV-2 using different approaches in previous studies are listed in Table 4.

The room ventilation rate with clean air in the Wells-Riley model indicates the total fresh/virus-free air delivered to the room, including the outdoor air and filtrated/disinfected recirculated air of the ventilation system, the outdoor air supplied through natural ventilation, the clean air supplied by the portable air cleaners, the virus-laden particle removal through UVGI systems, deposition and natural inactivation. The outdoor air is clean and safe for indoor ventilation, except when strong emission sources exist outdoors (Setti et al. 2020b).

Available Wells-Riley model-based tools for estimating the airborne risk of COVID-19

Multiple tools for estimating the airborne infection risk of COVID-19 have been developed based on the Wells-Riley model. For example, Kasibhatla et al. (2020) developed a Web-based tool COVID Exposure Modeler for estimating the COVID-19 infection risk from airborne transmission during classroom teaching incorporating a Monte Carlo approach. Allen, Cedeno-Laurent, and Miller (2020) developed a Google sheet to calculate the risk reduction by using portable air cleaners. Corsi, Van Den Wymelenberg, and Parhizkar (2020) established a Web-based platform named SafeAirSpaces COVID-19 Aerosol Relative Risk Estimator for evaluating the airborne infection risk and tracking the infectious particles. Dols et al. (2020) and William et al. (2020) developed a Web-based tool Fate and Transport of Indoor Microbiological Aerosols (FaTIMA), which allows for the determination of the indoor fate of microbiological aerosols associated with ventilation, filtration, deposition, and inactivation mechanisms. Riediker and Monn (2020) built a spreadsheet tool Indoor Scenario Simulator to estimate the virus concentration in the room and the inhaled dose of other people in the same room. REHVA (2020) developed a spreadsheet tool COVID-19 Ventilation Calculator to estimate the effect of ventilation on COVID-19 airborne transmission. Jimenez and Peng (2020) established a COVID-19 Aerosol Transmission Estimator on Google sheet and evaluated the infection risks in typical indoor scenarios including classrooms, supermarkets, and stadiums. Bazant (2021) developed a Web-based application COVID-19 Indoor Safety Guideline that can estimate the infection risk using the Wells-Riley model and indicate the safe level of occupancy or duration time based on the input configurations. It can also assess the risk based on indoor carbon dioxide concentrations. J. Shen et al. (2021) developed a spreadsheet-based calculator for estimating the infection risk in several space types including ballrooms, conference rooms, bistro/cafeteria, hotel guest rooms, offices, and classrooms. Both steady-state and stochastic simulations can be performed with the calculator to evaluate the effectiveness of various mitigation strategies in reducing the infection risks from baseline conditions established by ASHRAE 62.1 (2019).

Mitigation strategies on airborne transmission

Public health agencies have introduced multiple control strategies for mitigating the airborne transmission risk of COVID-19 in their guidelines (ASHRAE 2020a; Osha n.d.; REHVA 2021; U.S. CDC 2021h; U.S. EPA 2020; WHO 2021b). A recent review introduced and compared the HVAC operating guidelines in different countries during the pandemic (M. Guo et al. 2021). Many researchers also presented different IAQ control strategies (Bazant and Bush 2021; A. K. Melikov, Ai, and Markov (2020; Lidia Morawska et al. 2020; J. Shen et al. 2021; J. Zhang 2020). The IAQ control strategies across multiple scales (from a whole building to rooms, and to personal microenvironments and breathing zone) can be generally divided into three categories, i.e. source control, ventilation, and air cleaning. The commonly used strategies in indoor environments for reducing the infection risk through airborne transmission typically include masks (Brooks et al. 2021; Cook 2020; A. Davies et al. 2013; Liao et al. 2021; Tabatabaeizadeh 2021; Ueki et al. 2020), improved ventilation, and air distribution (Ahlawat et al. 2020; Almilaji and Thomas 2020; Blocken et al. 2021; Borro et al. 2021; CDPH 2020; Correia et al. 2020; Dai and Zhao 2020a; Dillon and Sextro 2020; Dygert and Dang 2012; Masotti et al. 2021; Lidia Morawska et al. 2021; Stabile et al. 2021; Sun and Zhai 2020), standalone air cleaners and upper-room UVGI systems (Beggs and Avital 2020; Blocken et al. 2021; Z. Feng et al. 2021; He et al. 2021; Ko, First, and Burge 2002; Xu et al. 2003), and occupancy control and restriction (Bazant et al. 2021; A. K. Melikov, Ai, and Markov (2020; Sun and Zhai 2020). Detailed reviews of mitigation strategies can be found in several published papers (M. Guo et al. 2021; Kong et al. 2020; J. Shen et al. 2021; S. Tang et al. 2020; J. Zhang 2020). However, most available studies only discussed the performance of mitigation strategies on reducing infection risks but did not consider other factors that can likely affect the use of mitigation strategies in practice, such as cost, durability, and accessibility. A review of both benefits and costs of control strategies can help design the control strategies in indoor environments.

The effectiveness and costs of various control strategies can vary greatly on different scales. A medical mask can typically protect the susceptible person effectively by reducing the inhalation of virus-carrying aerosols by around 50% with a very low cost (around US\$0.2 per mask). However, long-term use of masks can still be costly to sustain since medical masks are disposable. Upgrading filters in the HVAC system can benefit lots of occupants in multiple rooms. A HEPA filter can generally remove over 99.9% particles for the recirculated air of the building ventilation system, but it is also relatively expensive to purchase one (typically US\$280 per filter). However, considering the amount of benefited people and the long duration of the filter (typically 6-12 months), the general cost of using HEPA filters in the ventilation system is actually not so high. But upgrading building ventilation filters cannot completely avoid airborne transmission in short distances while wearing

masks can efficiently reduce the exposure to aerosols in short distances as masks can protect people's breathing zone. Therefore, both benefits and costs should be considered in designing the control strategies for a specific scenario. J. Shen et al. (2021) quantitatively analyzed the effectiveness and costs of some IAO control strategies in mitigating the airborne transmission risks of COVID-19. Besides, the accessibility of strategy should also be considered. Some strategies like cloth masks are relatively easy to access and can even be homemade using household materials during the shortages of masks (A. Davies et al. 2013), which makes it a strategy that can be applied widely. For PPE, the comfort of wearing, which can be quantified by the pressure drop across the PPE material (A. Davies et al. 2013; M. Zhao et al. 2020), should also be considered as it may also impact the use of PPE among the population. Some other control strategies may require professional designs before being implemented to maximize the potential of risk mitigation, e.g. displacement ventilation, personal ventilation, and local air exhaust. Inappropriate ventilation may even elevate risk (Y. Li et al. 2020; Shao et al. 2021). Another example is the implementation of the upper-room UVGI system. A well-designed upper-room UVGI system can typically provide an equivalent 6.7 to 33 air changes per hour to the room, depending on its operating conditions and circumstance (R. Riley and Nardell 1989; U.S. CDC and NIOSH 2009; Xu et al. 2003). But the performance of virus inactivation by the upperroom UVGI system also relies on the room air distribution. Displacement ventilation may reduce the efficiency (around 78% of the efficiency incorporating with mixing ventilation (Kanaan and Abou Moughlbay 2018)) as it decreases the residence time of the virus in the irradiated zone. Besides, the accessibility of some control strategies may vary at different stages of the pandemic. U.S. CDC defined three different strategies for PPE usage guidance, i.e. conventional, contingency, and crisis strategies, for different PPE supply capacity situations (U.S. CDC 2021d). If PPE supply shortages are anticipated or exist, medical-grade PPE like surgical and N95 masks should be reserved for healthcare workers (U.S. CDC 2020c). NIOSH (2015) defined the hierarchy of controls for eliminating the exposure to occupational hazards, such as respiratory pathogen, which has been used as a means of determining how to implement feasible and effective control solutions.

Some intervention strategies for mitigating airborne transmission in different scenarios have been introduced in "Outbreaks in different scenarios and the role of airborne transmission" section. This section reviewed some widely used control strategies considering their benefits and costs, including the effectiveness of risk reduction, effective scale, capital cost, durability, and accessibility (Table 5). It aims to help select the best control strategies (both reliable and affordable) in different scenarios.

Summary of major findings

There has been substantial evidence supporting the airborne transmission of SARS-CoV-2, including the viable virus-laden aerosols detected in the air and the supports from

retrospective analyses on actual outbreaks. In many outbreak events, the airborne route has been considered the predominant route for virus transmission. The viral loads in the respiratory tract are related to the emission rate of pathogens through respiratory activities. The viral load in the mouth of COVID-19 patients is typically below 10⁹ RNA copies/mL. The majority (typically 80–90%) of particles expelled during various human activities are smaller than 1-2 µm in diameter. However, considering that droplets in smaller diameters may contain less virus due to the smaller particle volume, larger expelled particles usually contain more viral RNA. The typical inactivation rate of viable SARS-CoV-2 aerosols at typical indoor temperature (20-23 °C) and relative humidity (30-70%) without sunlight is generally between 0 and 1 h⁻¹, while sunlight can contribute greatly to the inactivation of the viability of SARS-CoV-2 ($>7 h^{-1}$ and $>18 h^{-1}$ under low- and high-intensity sunlight). The majority of outbreaks reported in indoor scenarios were related to long-term care facilities, K-12 schools, restaurants and bars, retail facilities, and offices. The outbreaks in correctional facilities, colleges and universities, long-term care facilities, and transportations typically infected more people according to the reported data. Indoor ventilation is a key factor impacting infection risk. A pathogen-source-based or health-based design criteria for indoor ventilation is probably more applicable to protect against infection transmission. More efforts regarding the indoor ventilation design criteria can be conducted in the future. Besides, both benefits and costs should be considered in designing the control strategies for mitigating infection risk, including PPE, ventilation strategies, partitions, air cleaning and disinfection technologies, and occupancy control strategies.

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Data availability statement

The data that support the findings of this study are available from the corresponding author upon reasonable request.

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