Association of the infection probability of COVID-19 with ventilation rates in confined spaces

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Abstract
A growing number of cases have proved the possibility of airborne transmission of the coronavirus disease 2019 (COVID-19). Ensuring an adequate ventilation rate is essential to reduce the risk of infection in confined spaces. In this study, we estimated the association between the infection probability and ventilation rates with the Wells–Riley equation, where the quantum generation rate ($q$) by a COVID-19 infector was obtained using a reproductive number-based fitting approach. The estimated $q$ value of COVID-19 is 14–48 h⁻¹. To ensure an infection probability of less than 1%, a ventilation rate larger than common values (100–350 m³/h per infector and 1200–4000 m³/h per infector for 0.25 h and 3 h of exposure, respectively) is required. If the infector and susceptible person wear masks, then the ventilation rate ensuring a less than 1% infection probability can be reduced to a quarter respectively, which is easier to achieve by the normal ventilation mode applied in typical scenarios, including offices, classrooms, buses, and aircraft cabins. Strict preventive measures (e.g., wearing masks and preventing asymptomatic infectors from entering public spaces using tests) that have been widely adopted should be effective in reducing the risk of infection in confined spaces.

Keywords
COVID-19, SARS-CoV-2, ventilation, infection probability, Wells–Riley equation

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1 Introduction
The once-in-a-century coronavirus disease 2019 (COVID-19) pandemic has shown that the airborne transmission of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) cannot be ignored (Morawska and Cao 2020). Although the transmission of COVID-19 occurs mainly via droplets during close contact or via contaminated surfaces, a recent study showed that SARS-CoV-2 remains viable in aerosols for multiple hours (Van Doremalen et al. 2020). A recent report suggested that current research supports the possibility that SARS-CoV-2 can be spread via bioaerosols generated directly by patients’ exhalation (National Research Council 2020). Furthermore, recent visualization studies also support that SARS-CoV-2 may spread between the infector and susceptible person simply by speaking (Anfinrud et al. 2020; Stadnytskyi et al. 2020). A recent field sampling study further indicated that SARS-CoV-2 is widely distributed in the air, and the transmission distance in the air might be up to 4 m (Guo et al. 2020). These results partly explain the easier spread of SARS-CoV-2 in confined spaces. The Wells–Riley equation is a classic model for quantifying the risk associated with airborne transmission of respiratory diseases (Riley et al. 1978). It has been used to demonstrate that some building factors (particularly the ventilation rate) are important removal mechanisms for airborne infectious agents (Escombe et al. 2007). In addition, a report by WHO suggested that insufficient ventilation increases disease transmission (Chartier et al. 2009). Therefore, ensuring a sufficient ventilation rate for offices, classrooms, and public transport is essential to reduce the potential risk of infection in these confined spaces, which is extremely important for daily life.

In this study, we employed the Wells–Riley equation to estimate the association between the infection probability and ventilation rate, where the quantum generation rate ($q$) is a critical parameter. The value of $q$ of a COVID-19 infector is currently not well established, and we obtained this value based on its fitted correlation with the basic reproductive number ($R_0$). We then estimated the association between
the infection probability and ventilation rate for some typical scenarios, including offices, classrooms, buses, and aircraft cabins.

2 Methods

2.1 Wells–Riley equation

The Wells–Riley equation is as follows (Riley et al. 1978):

\[ P = \frac{C}{S} = 1 - e^{-q_{p}/Q} \]  

(1)

where \( P \) is the probability of infection risk; \( C \) is the number of cases that develop infection; \( S \) is the number of susceptible people. \( I \) is the number of source patients (infectors); in this study, we focused on normal scenarios (not health care settings for COVID-19 patients), thus, we assumed \( I = 1 \). \( p \) is the pulmonary ventilation rate of each susceptible person per hour (m³/h); \( p = 0.3 \) m³/h when people are sitting or participating in light activity indoors (Duan et al. 2013). \( Q \) is the room ventilation rate (m³/h); \( q \) is the quantum generation rate produced by one infector (h⁻¹); \( t \) is the exposure time (h).

The unit "quantum" in this risk model is not an actual physical unit; it is a hypothetical infectious dose unit that is typically back-calculated from epidemiological studies. One quantum is defined as a collection of pathogen particles that can infect susceptible people (Riley et al. 1978).

The Wells–Riley equation has two key assumptions, as follows: (1) droplet nuclei are evenly distributed in space, which means that the infection risk predicted by this equation is uniform within the space and (2) the equation neglects the viability and infectivity of the pathogen quanta.

Owing to the filtration effect of the mask, we considered that wearing a mask of infector can dilute exhaled pathogen concentration; while for the susceptible person, it’s equivalent to increasing the room ventilation rate. The filtration efficiency of ordinary medical surgical masks on virus-laden aerosols is about 60% (Hui et al. 2012), which can be set as 50% considering the influence of air leakage (Davies et al. 2013). The modified Wells–Riley equation is as follows:

\[ P = \frac{C}{S} = 1 - e^{-q_{p} / Q} \]  

(2)

where \( q_{p} \) is the exhalation filtration efficiency, \( q_{r} = 50\%; \) \( q_{s} \) is the respiration filtration efficiency, \( q_{s} = 50\% \).

Furthermore, a previous study modified the Wells–Riley equation to include deposition rates \( k_{\text{deposition}} \) and filtration removal rates \( k_{\text{filtration}} \) (Stephens 2012).

The \( k_{\text{deposition}} \) depends primarily on the particle size, density, and room characteristics, such as air speeds and surface areas (Fennelly and Nardell 1998; Fisk et al. 2004). It’s measured (Liu et al. 2020b) that viral RNA in aerosols and found that SARS-CoV-2 aerosol mainly resides in two size ranges, namely 0.25–0.50 μm and 0.50–1.00 μm. It’s indicated that the range of \( k_{\text{deposition}} \) is 0.21–0.63 h⁻¹ with a mean value of 0.36 h⁻¹ for aerosols in such two size ranges (Ji and Zhao 2015). This \( k_{\text{deposition}} \) value has a negligible effect compared with the removal effect of ventilation in this study.

Regarding the \( k_{\text{filtration}} \) value, if we treat the ventilation rate \( Q \) as the equivalent clean air delivery rate (CADR), which is equal to the actual ventilation rate multiplied by filtration efficiency (Foarde 1999), then no modification is required to incorporate it.

Finally, the key input parameter when applying the Wells–Riley equation is \( q \).

2.2 Quantum generation rate with a reproductive number-based fitting approach

So far, there have been no available reported values of \( q \) of SARS-CoV-2. To obtain a reasonable \( q \) value, we collected the known \( q \) value and \( R_{0} \) for other airborne transmitted infectious diseases in previous studies, and fitted the association between them. \( R_{0} \) is the key epidemiological determinant that characterizes the transmission potential of an infectious disease, which is defined as the average number of infectious individuals created by a single infector in a susceptible population (Anderson et al. 1992). As both the \( q \) and \( R_{0} \) essentially determine the rate of spread of an epidemic and are estimated based on actual confirmed cases, we think that it may be logical to obtain the \( q \) value based on its association with \( R_{0} \) as a compromised but relatively reasonable approach.

We then estimated \( q \) with the \( R_{0} \) of COVID-19 based on the fitted equation. Table 1 lists the values of \( q \) and \( R_{0} \) for several typical infectious diseases collected from the references. It’s reported that the transmission of respiratory pathogens such as measles, tuberculosis, middle east respiratory syndrome coronavirus (MERS), severe acute respiratory syndrome (SARS) and influenza in indoor environments are governed by similar physical and biological

| Disease          | \( q \) (h⁻¹) | \( R_{0} \)     | Reference         |
|------------------|---------------|-----------------|-------------------|
| Tuberculosis     | 1–50          | 2.22–5.46       | Stephens 2012; Li et al. 2018 |
| MERS             | 6–140         | 0.50–1.20       | WHO 2019          |
| SARS             | 10–300        | 2.00–5.00       | WHO 2003          |
| Influenza        | 15–500        | 1.60–3.00       | Lee et al. 2012   |
| Measles          | 570–5600      | 11.00–18.00     | Plans Rubió 2012  |
processes to COVID-19 (Stephens 2012; WHO 2003, 2019). With the values listed in this table as inputs, we fitted the association between \( q \) and \( R_0 \) with a least square method implemented in Origin 2019.

3 Results

3.1 Fitted quantum generation rate of COVID-19 with the basic reproductive number

The fitted curve representing the association between \( q \) and \( R_0 \) is shown in Figure 1, and the fitted equation is as follows:

\[
q = A + B_1 \times R_0 + B_2 \times R_0^2
\]  
(3)

where \( A = -30.27958 \pm 26.07 \), \( B_1 = -44.81536 \pm 12.34048 \), and \( B_2 = 19.67934 \pm 0.62317 \).

With the widely used range of \( R_0 \) of 2.0–2.5 (Imai et al. 2020; Li et al. 2020; Majumder and Mandl 2020; Read et al. 2020; Zhao et al. 2020b), we obtained the corresponding range of \( q \) for COVID-19, which was 14–48 h\(^{-1}\).

Considering that the \( R_0 \) of measles is much larger than that of other infectious diseases, we fitted the association between \( q \) and \( R_0 \) with the same method but removed the \( R_0 \) of measles (Figure 2). The corresponding range of \( q \) for COVID-19 is 2–40 h\(^{-1}\), which is comparable to the results obtained from Figure 1 and Eq. (3).

During the preparation of this manuscript, we found a study conducted by Lu et al. (2020) reporting a suspected case of airborne transmission of COVID-19 in Guangzhou, China, which is a suitable epidemiological case for the back-calculation of \( q \). According to the description presented in the article, we back-calculated the value of \( q \) to be 22 h\(^{-1}\). Furthermore, we also noticed another novel method for predicting the value of \( q \) on the basis of the emitted viral load from the mouth (Buonanno et al. 2020). The results show that \( q \) varies significantly as a function of different parameters (including inhalation rate, type of respiratory activity, and activity level). Our aim is to analyze the asymptomatic case in light exercise activity, which represents the typical characteristic of infectors in common public spaces. Considering this point, the corresponding \( q \) reported by Buonanno et al. (2020) is between 10.5 and 33.9 h\(^{-1}\). Such values are in the same magnitude as the results in this study.

Based on the above analysis, it can be considered that the initially estimated value of \( q \) in this study is reliable. Thus, we applied a \( q \) value of 14–48 h\(^{-1}\) for the sedentary state for later analysis.

3.2 Association between infection probability and ventilation rate

3.2.1 Infection probability

Figure 3 shows the estimated association between the infection probability and ventilation rate. For the case with one infector inside of a confined space, a ventilation rate of 100–350 m\(^3\)/h per infector was required to ensure an infection probability of less than 1% for 0.25 h of exposure of a susceptible person. A ventilation rate of 1200–4000 m\(^3\)/h per infector was required to ensure an infection probability of less than 1% for 3 h of exposure of a susceptible person.

3.2.2 Effect of wearing a mask

The estimated association between the infection probability and ventilation rate for the cases with masks is shown in Figure 4. The results indicated that wearing a mask plays an important role in reducing the infection risk. For the case where there was one infector inside of a confined space, if both the infector and susceptible person wore masks, then the required ventilation rate to ensure an infection probability of less than 1% was reduced to 30–90 m\(^3\)/h per infector for 0.25 h of exposure and 300–1000 m\(^3\)/h per infector for 3 h of exposure of a susceptible person, respectively.
3.2.3 Typical scenarios

Figure 5 shows the association of the estimated infection probability with the ventilation rate for some typical scenarios. Table 2 lists the corresponding air change rates (ACHs) for typical volumes of the examined confined spaces. The ACH is a measurement of how much fresh/clean air replaces indoor air in 1 h (Sherman and Wilson 1986), which is more widely used in engineering to determine if ventilation and air conditioning systems can provide a sufficient ventilation rate to ensure a low infection probability.

If people are wearing masks, then natural ventilation or normal mechanical ventilation can provide a sufficient ventilation rate to ensure that the infection probability is less than 1% for most scenarios (except for offices; Table 2). However, not wearing masks may result in a relatively higher infection risk for most cases, especially if $q$ reaches the maximum value. In these scenarios, additional measures could be suggested if the actual ventilation rate is out of a safe range, including UV sterilization, HEPA filtration and increasing outdoor air rate (Zhao et al. 2020a).
**Discussion**

$R_0$ varies due to different methodological issues, including the utilized datasets, confirmed cases, outbreaking region and mathematical models. Several methods have been produced, resulting in different estimations of $R_0$. For example, there are studies showing that the $R_0$ of COVID-19 can be reached a high level of 6.47 (Tang et al. 2020). However, $R_0$ seems to be stabilized at around 2–3, which is obtained from a large database (Liu et al. 2020a) and from more different researches (D’Arienzo and Coniglio 2020; Imai et al. 2020; Li et al. 2020; Majumder and Mandl 2020; Read et al. 2020; Riou and Althaus 2020; Zhao et al. 2020b), the data used includes both the Mainland of China and overseas outbreak up to March, 2020. Thus the $R_0$ of 2–2.5 here can be expected to be reliable, as it builds upon more case data.

Given the large uncertainties associated with each of the parameters, we relied on two assumptions in an attempt to link the indoor ventilation strategy to infectious disease risk models, namely, the ventilation efficiency was assumed to be 1 and the concentration of viruses was assumed to be homogeneous in the indoor environment.

The estimation indicates that when an infector remains in a public confined space, the infection risk is relatively high with an infection probability of approximately 2% at the common ventilation rate (500–2500 m³/h per infector for 0.25 h of exposure; Figure 3). Such findings may partly explain the early large-scale epidemics in China and European and American countries.

Thus, preventing the infectors (especially asymptomatic infectors that have not been confirmed) from entering these types of public spaces is critical to suppress the spread of the virus via airborne transmission. Current measures, strict screening of asymptomatic infectors with wider nucleic acid tests, and home isolation have already ensured that infectors are unlikely to enter public spaces and have ensured a relatively safe environment in confined spaces.

Furthermore, the above findings imply that wearing an ordinary medical surgical mask is effective; thus, it is important to advise people to wear masks when they enter or remain in confined spaces.

It is acknowledged that COVID-19 is transmitted mainly by close contact or droplets deposited on contaminated surfaces. However, the risk of potential or opportunistic airborne transmission cannot be ignored, especially in public confined spaces. The value of $q$ also warrants further study.

Furthermore, strict preventive measures have been widely adopted, such measure provides a prerequisite for common ventilation strategies to ensure a sufficiently low risk of airborne transmitted infection.

**Conclusion**

This study employed Wells–Riley equation to predict infection probability in some typical scenarios based on the estimated quantum generation rate. The main conclusions are as follows:

1. A quantum generation rate of 14–48 h⁻¹ for COVID-19 is obtained with a reproductive number-based fitting approach.
2. The Infection probability less than 1% requires ventilation rate larger than 100–350 m³/h per infector and 1200–4000 m³/h per infector for 0.25 h and 3 h of exposure.
3. Wearing an ordinary medical surgical mask is effective to reduce the infection risk.

*The ACHs in red text indicate that the ventilation rates could be achieved with a normal ventilation system or natural ventilation for the corresponding scenarios.*
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