Letter

# Exhaled $CO_2$ as a COVID-19 Infection Risk Proxy for Different Indoor Environments and Activities

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**ABSTRACT:** $CO_2$ is co-exhaled with aerosols containing SARS-CoV-2 by COVID-19-infected people and can be used as a proxy of SARS-CoV-2 concentrations indoors. Indoor $CO_2$ measurements by low-cost sensors hold promise for mass monitoring of indoor aerosol transmission risk for COVID-19 and other respiratory diseases. We derive analytical expressions of $CO_2$-based risk proxies and apply them to various typical indoor environments. The relative infection risk in a given environment scales with excess $CO_2$ level, and thus, keeping $CO_2$ as low as feasible in a space allows optimization of the protection provided by ventilation. We show that the $CO_2$ level corresponding to a given absolute infection risk varies by >2 orders of magnitude for different environments and activities. Although large uncertainties, mainly from virus exhalation rates, are still associated with infection risk estimates, our study provides more specific and practical recommendations for low-cost $CO_2$-based indoor infection risk monitoring.
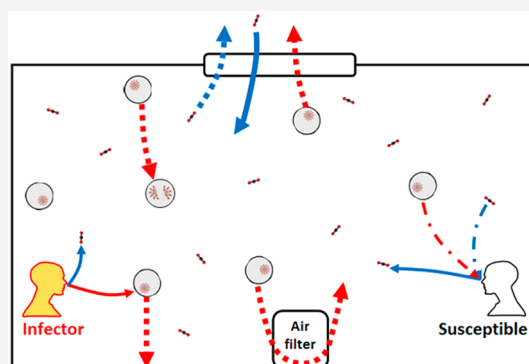
## INTRODUCTION

Coronavirus disease 2019 (COVID-19) is currently sweeping the world and causing major losses of human life.[1] Lockdowns imposed to various extent worldwide for the COVID-19 transmission reduction are not supposed to be long-term measures, otherwise they lead to unaffordable social and economic costs. On the other hand, resumption of social, educational, and business activities raises concerns about transmission resurgence.

In past few months, there has been rapidly mounting evidence for COVID-19 transmission via aerosols,[2−5] i.e., severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2)-containing particles with diameters of <100 $\mu$m that can float in the air for minutes to hours. Such aerosols have been detected in exhaled air of COVID-19 patients[6] and in hospital air,[7,8] and the behaviors of smaller ones out of the proximity of sources have been shown to be similar to those of gas.[9,10] Transmission is much easier indoors than outdoors, which is most consistent with aerosols.[4,11,12] As humans spend most of their time in indoor environments, where air volumes are limited and virus-laden aerosols may easily accumulate, mitigation of indoor COVID-19 transmissions is a subject of great interest[13,14] and is key to a successful societal and economic reopening. Practical, affordable, and widely applicable measures for monitoring and limiting indoor transmission risks are urgently needed.

Direct measurements of virus-containing aerosols are extremely difficult and slow. Indoor $CO_2$ was suggested as an indicator of ventilation of indoor spaces in the 19th century,[15] and more recently as a practical proxy of respiratory infectious

disease transmission risk,[16] as pathogen-containing aerosols and $CO_2$ are co-exhaled by those infected (Figure 1). Because the background (ambient) $CO_2$ level is almost stable and indoor excess $CO_2$ is usually only from human exhalation, measurements of indoor $CO_2$ concentrations by low-cost $CO_2$ sensors can often be good indicators of infection risk and suitable for mass deployment.[17,18] However, the $CO_2$ level corresponding to a given COVID-19 infection risk is largely unknown. A few guideline limit concentrations have been proposed, but without a solid and quantitative basis.[19,20] In particular, only a single $CO_2$ threshold was recommended in each of these proposed guidelines. Whether a single $CO_2$ concentration ensures a low COVID-19 infection risk in all common indoor environments remains an open question but is also critical for effective $CO_2$-based mass risk monitoring.

In this study, we derive the analytical expressions of the probability of indoor COVID-19 infection through room-level aerosol transmission only (i.e., assuming social distance is kept so that close proximity aerosol and droplet pathways are eliminated; fomite transmission is not included), human-exhaled $CO_2$ concentration, and subsequently a few $CO_2$-based quantities as infection risk proxies. On the basis of available
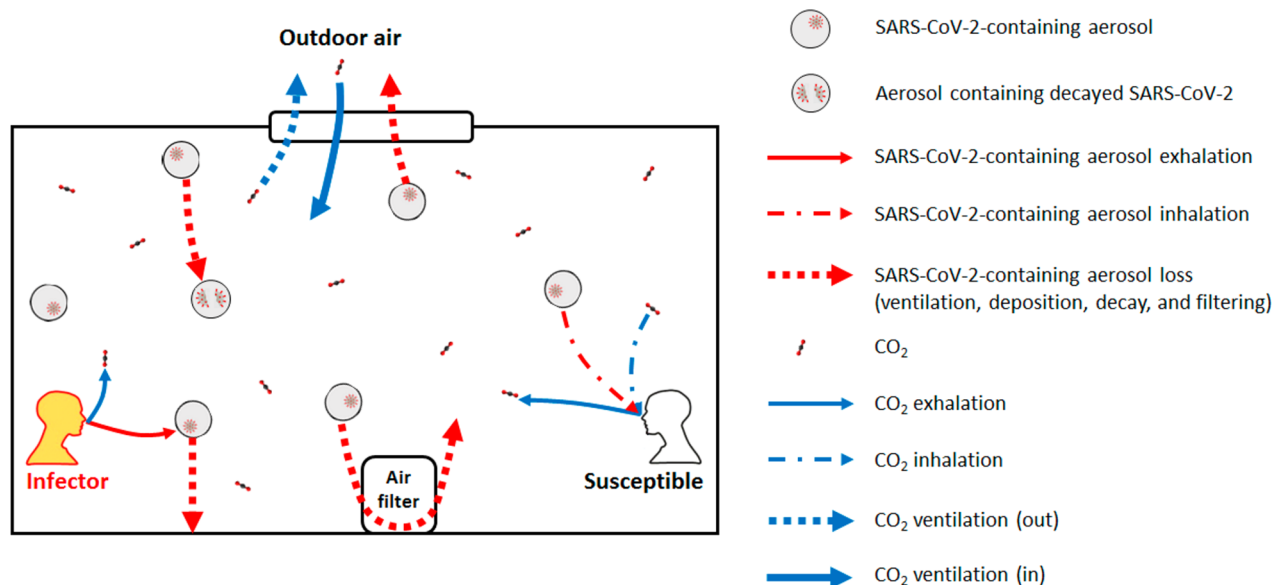
**Figure 1.** Schematic illustrating the exhalation, inhalation, and other loss processes of SARS-CoV-2-containing aerosols and the exhalation, inhalation, and other sources and sinks of $CO_2$ in an indoor environment.

data, we apply these expressions to common indoor settings to answer the open question mentioned above.

### ■ MATERIALS AND METHODS

To derive the SARS-CoV-2 aerosol concentration in indoor air, we assume well-mixed air (Figure 1). The degree of inhomogeneity can be easily quantified with portable low-cost sensors. If significant inhomogeneity in indoor air is present, the indoor space can often be approximated as several compartments, each of them having relatively well-mixed air. Ventilation with outdoor air, decay of virus and deposition of virus onto surfaces, and additional control measures [e.g., air filtration and use of germicidal ultraviolet (UV) radiation] result in losses of infective virus from indoor air. Other sinks (e.g., inhalation by humans and animals indoors) are assumed to be insignificant. This model will underestimate the risk in environments with significant nonrespiratory sources of infective aerosols (e.g., bathrooms due to toilet flushing, resuspension in healthcare facilities due to donning and doffing of personal protective equipment). The amount of the virus infectious doses ($n$, "quanta") inhaled by a susceptible person determines their probability of infection ($P$) (see Table S1 for the list of symbols in this study). According to the Wells−Riley model of aerosol infection[21]

$$P = 1 - e^{-n} \qquad (1)$$

One SARS-CoV-2 quantum corresponds to a probability of infection of $1 - 1/e$ (63%). The expected value of $n$ ($\langle n \rangle$) for an originally uninfected person corresponding to a given level of immunity in the local population (probability of an occupant being immune, $\eta_{im}$) can be calculated as follows

$$\langle n \rangle = (1 - \eta_{im}) c_{avg} BD(1 - m_{in}) \qquad (2)$$

where $c_{avg}$, $B$, $D$, and $m_{in}$ are the average virus concentration (quanta per cubic meter), the breathing rate of the susceptible person (cubic meters per hour), the duration of the event (hours), and the mask filtration efficiency for inhalation, respectively. The term $1 - \eta_{im}$ is included because quanta inhaled by an immune uninfected individual will not lead to

infection and should be excluded. Under the assumption of no occupants and no SARS-CoV-2 in the indoor air at the start of the event, the analytical expression of the expected value of $c_{avg}$ based on the prevalence of infectors in the local population (probability of an occupant being an infector, $\eta_I$, $\langle c_{avg} \rangle$), is (see Section S1 of the Supporting Information for the derivation)

$$\langle c_{avg} \rangle = \frac{\eta_I (N-1) E_p (1 - m_{ex})}{V} \left( \frac{1}{\lambda} - \frac{1 - e^{-\lambda D}}{\lambda^2 D} \right) \qquad (3)$$

where $N$ is number of occupants, $E_p$ is the SARS-CoV-2 exhalation rate by an infector (quanta per hour), $m_{ex}$ is the mask filtration efficiency for exhalation, $V$ is the indoor environment volume (cubic meters), and $\lambda$ is the first-order overall rate constant of the virus infectivity loss (inverse hours) that includes the ventilation with outdoor air and all other virus removal and deactivation processes.

If there are no other significant $CO_2$ sources or sinks (e.g., gas/coal stove and pets/plants), i.e., if indoor excess $CO_2$ (relative to the background outdoor level) production is only due to human exhalation and its loss is ventilation, similar quantities for $CO_2$ can be expressed as follows (see Section S1 for the derivation)

$$n_{\Delta CO_2} = \Delta c_{avg,CO_2} BD \qquad (4)$$

$$\Delta c_{avg,CO_2} = \frac{N E_{p,CO_2}}{V} \left( \frac{1}{\lambda_0} - \frac{1 - e^{-\lambda_0 D}}{\lambda_0^2 D} \right) \qquad (5)$$

where $n_{\Delta CO_2}$, $\Delta c_{avg,CO_2}$, and $E_{p,CO_2}$ are the inhaled excess (human-exhaled) $CO_2$ volume (cubic meters), the excess $CO_2$ volume mixing ratio, and the $CO_2$ exhalation rate per person (cubic meters per hour), respectively, and $\lambda_0$ is the ventilation rate (inverse hours).

When $P$ is low, as it should be for a safe reopening, $P \approx n$. As airborne SARS-CoV-2 and excess $CO_2$ are co-exhaled and co-inhaled, in principle $n_{\Delta CO_2}$ can be a proxy of $\langle n \rangle$, and thus $P$. The ratio of $n_{\Delta CO_2}$ to $\langle n \rangle$ (in cubic meters per quantum)
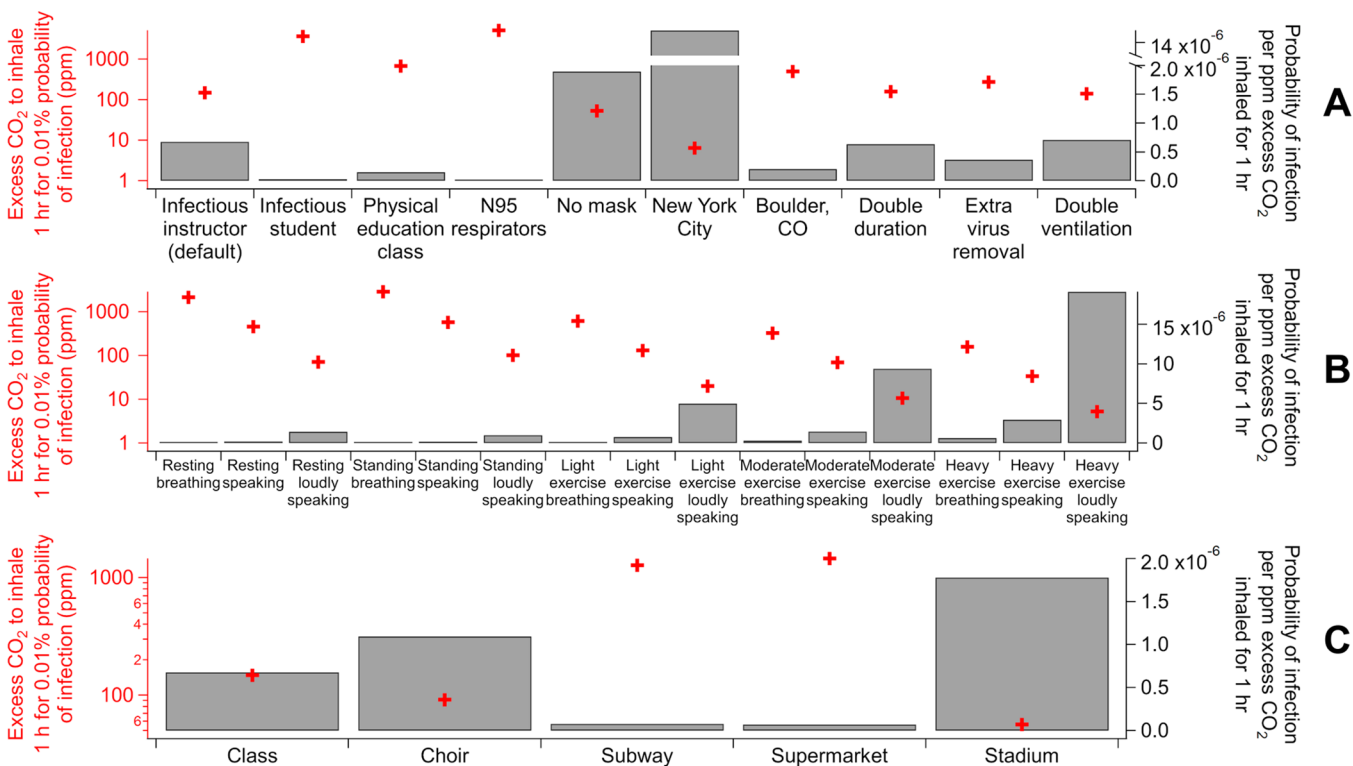
**Figure 2.** Excess $CO_2$ volume mixing ratio (parts per million) that an uninfected individual inhales for 1 h for a probability of infection of 0.01% ($\Delta c_{CO_2}^*$) and probability of infection per parts per million excess $CO_2$ inhaled for 1 h (inversely proportional to $\Delta c_{CO_2}^*$) with a probability of an occupant being an infector of 0.1% (except the New York City and Boulder, CO, cases in panel A and the choir case in panel C) for (A) variants of the university class case (see Table S2 for the case details), (B) various activities (see Table S3 for details of the activities), and (C) several indoor environments (see Table S4 for the case details).

$$\frac{n_{\Delta CO_2}}{\langle n \rangle} = \frac{N E_{p,CO_2}}{(1 - \eta_{im})\eta_I(N-1)E_p(1-m_{ex})(1-m_{in})}$$
$$\times \frac{\frac{1}{\lambda_0} - \frac{1-e^{-\lambda_0 D}}{\lambda_0^2 D}}{\frac{1}{\lambda} - \frac{1-e^{-\lambda D}}{\lambda^2 D}} \tag{6}$$

indicates the volume of inhaled excess $CO_2$ corresponding to a unit inhaled quantum. However, this quantity, involving inhaled $CO_2$ volume that is difficult to measure, is not practical for widespread transmission risk monitoring, which usually requires a fast decision-making process simply based on the indoor $CO_2$ concentration reading (usually in parts per million) of a low-cost sensor. Therefore, we propose, as another proxy of the risk of an environment with $\eta_I = 0.1$%, the reference excess $CO_2$ level ($\Delta c_{CO_2}^*$), i.e., the volume mixing ratio of excess $CO_2$ that an uninfected individual inhales for a typical duration (1 h) in that environment for a typical probability of infection (0.01%).

$$\Delta c_{CO_2}^* = \frac{0.0001/1h \times N E_{p,CO_2}}{(1 - \eta_{im})\eta_I(N-1)E_p(1-m_{ex})(1-m_{in})B}$$
$$\times \frac{\frac{1}{\lambda_0} - \frac{1-e^{-\lambda_0 D}}{\lambda_0^2 D}}{\frac{1}{\lambda} - \frac{1-e^{-\lambda D}}{\lambda^2 D}} \tag{7}$$

This quantity is closely related to the excess $CO_2$ level corresponding to the unity basic reproduction number ($R_0$)[16] (see Section S2) and can be directly and easily compared to

$CO_2$ sensor readings. The ratio of the excess $CO_2$ reading to $\Delta c_{CO_2}^*$ is that of the probability of infection of an originally uninfected person in that environment for 1 h to 0.01%. $\Delta c_{CO_2}^*$ scales (roughly) linearly with most of the parameters in eq 7 (see discussions below). A $P$ of 0.01% being chosen as the reference does *not* imply safety at this $P$ in all situations, because when $N$ and/or $D$ is large, and/or the event is repeated many times (e.g., in school/university settings), the overall probability of infection for one susceptible person and/or total infections may still be significant.

## ■ RESULTS AND DISCUSSION

The reference excess $CO_2$ level is a function of a number of variables. A priori, varying any of them can result in a different value of $\Delta c_{CO_2}^*$ even for similar settings. As an example, we study a set of model cases for a typical university class. The cases are specified in Table S2. The $\Delta c_{CO_2}^*$ and $\frac{n_{\Delta CO_2}}{n}$ in these cases are shown in Figure 2A and Figure S1A, respectively.

In the base class case, the infector is assumed to be the instructor. Compared to the case with a student being the infector, $\Delta c_{CO_2}^*$ in the base case is ~1.5 orders of magnitude lower, just because the vocalization of the instructor, who usually speaks, greatly enhances $E_p$[22,23] while virus exhalation by students, who are assumed here to speak little, is much less efficient. In the case of a physical education (PE) class in the same indoor environment, where occupants are assumed to be doing heavy exercise and no talking, $\Delta c_{CO_2}^*$ is much lower than

for the infected student case in a traditional lecture (Figure 2A). Compared to sitting, heavy exercise increases both occupants' virus and $CO_2$ exhalation rates to similar extents,[22−24] which does not significantly change the reference excess $CO_2$ level. However, breathing rates of occupants doing intense activities are much higher than those sitting.[25] Even if $CO_2$ and SARS-CoV-2 concentrations are the same as in the infected student case, a susceptible person in the PE class case can still inhale a larger dose of SARS-CoV-2 and more excess $CO_2$ and have a remarkably different $P$. As a result, a single recommendation of indoor $CO_2$ threshold is not valid even for a series of school settings. The range of $CO_2$ levels measured in real-world classrooms is very large.[26] The reference excess $CO_2$ level of the infectious student case (relatively safe) is exceeded in some classrooms, while that of the infectious instructor case (relatively risky) is met in other classrooms.

According to eqs 2 and 3, whether occupants wear masks and what masks they wear can make a substantial difference in infection risk through virus filtration in the same indoor setting. However, masks do not filter $CO_2$. The base class case (with surgical masks), that with all occupants wearing N95 respirators, and that with no mask use have identical $CO_2$ mixing ratios, but an up to ~2 order of magnitude different probability of infection (Table S2) due to filtration of virus-containing particles by mask. Therefore, for the same probability of infection of 0.01%, the base class case is estimated by eq 7 to have a corresponding excess $CO_2$ level 30 times lower than the case with all occupants wearing N95 respirators but ~2 times higher than the case with no mask use (Figure 2A).

$\eta_I$ is obviously another important factor governing the infection risk, as $P$ is proportional to it. Again, it has no impact on $CO_2$. Compared to the base class case ($\eta_I$ = 0.001), the estimated situations of similar classes in New York City (NYC) in April ($\eta_I$ = 0.023) and in Boulder, CO, in June ($\eta_I$ = 0.0003) have ~20 times higher and ~2 times lower values of $P$, respectively (Table S2), and hence $\Delta c^*_{CO_2}$ values proportionally lower and higher, respectively (Figure 2A). Note that $\Delta c^*_{CO_2}$ is smaller than the current typical accuracy of low-cost $CO_2$ sensors (±50 ppm)[27] and cannot be meaningfully measured by those sensors in very risky situations such as the NYC case here. Closure of environments with such low permissible $\Delta c^*_{CO_2}$ is likely needed. However, $\eta_{im}$ usually cannot result in a difference in $P$ greater than a factor of 2 under conditions of interest, because if $\eta_{im}$ > 50%, the population has reached or is close to herd immunity[28] and widespread transmission risk monitoring is no longer needed.

According to eq 7, the other variables that can affect $\Delta c^*_{CO_2}$ are $N$, $D$, $\lambda$, and $\lambda_0$. $\Delta c^*_{CO_2}$ is generally not highly sensitive to them, although some of them (e.g., $\lambda$) can have a large impact on $P$. As long as occupants are not only a few, $\frac{N}{(N-1)}$, where $N$ plays a role in eq 7, is close to 1. The fraction term including $D$, $\lambda$, and $\lambda_0$ (after the product sign) in eq 7 usually does not deviate from 1 substantially (Figure S2). It is close to 1 when $\lambda D$ is very small and $\lambda/\lambda_0$ when $\lambda D$ is very large. As long as the indoor environment is not very poorly ventilated or equipped with very strong virus removal setups (e.g., substantial filtering of recirculated air, portable HEPA filters, and germicidal UV), $\lambda/\lambda_0$ is relatively close to 1. Compared to the base classroom case ($\lambda/\lambda_0$ ~ 1.3), doubling the duration or ventilation causes

minimal changes in $\Delta c^*_{CO_2}$. Increasing $\lambda/\lambda_0$ to ~3 by additional virus control measures increases $\Delta c^*_{CO_2}$ more significantly, as those measures do not remove $CO_2$, but this change is still within a factor of 2 for the range of control measures in these examples (Figure 2A).

As discussed above, occupants' activities indoors, to which $E_p$, $E_{p,CO_2}$, and $B$ are all related, are a major or dominant factor governing the infection risk. We thus compile the data of these parameters as a function of activity (intensity and vocalization degree) (Table S3). Note that this compilation has large uncertainties from $E_p$ data[22,23] and matching of activity categories, which are all classified differently for $E_p$, $E_{p,CO_2}$, and $B$ (see Section S3 for details). These uncertainties are currently difficult to quantify but likely large enough to be the dominant uncertainty sources for the model output. Other sources of uncertainty are thus not discussed. Further systematic uncertainty analyses would be of interest. However, the trends shown are clear and thus able to reveal the *relative* risk of these activities with confidence. Simply, the stronger the vocalization, the higher the risk, and the more intense activity, the higher the risk. We calculate $\Delta c^*_{CO_2}$ for these activities when $N$ is large, $D$ = 1 h, $\eta_I$ = 0.001, $\lambda_0$ = 3 h$^{-1}$, $\lambda$ = 4 h$^{-1}$, and no mask is used (Figure 2B), a setting similar to the class case. Three class cases, i.e., base, infected student, and PE cases, can be easily related to the activity categories of "standing−loudly speaking", "resting−breathing", and "heavy exercise−breathing", respectively. The related pairs have $\Delta c^*_{CO_2}$ values within a factor of ~2, and their mask use setting and close but different $E_p$, $E_{p,CO_2}$, and $B$ values can largely explain the differences in $\Delta c^*_{CO_2}$.

Then we apply this analysis to a range of real-world settings, in addition to the class case, i.e., the Skagit County choir superspreading event,[5] a subway car, a supermarket (focused on a worker), and an event in a stadium, which, though outdoors, often has somewhat stagnant air allowing virus-laden aerosols to build up and thus can be treated like an indoor environment (see Table S4 for the specifications of these cases). Figure 2C and Figure S1B show their $\Delta c^*_{CO_2}$ and $\frac{n_{\Delta CO_2}}{n}$, respectively. Again, these values span orders of magnitude. We can still relate these cases to the activity categories of "standing−loudly speaking", "resting−breathing", "light exercise−breathing" (or "light exercise−speaking"), and "light exercise−speaking" (or "light exercise−loudly speaking").

For the actual choir case, its $\eta_I$ is an order of magnitude lower than 0.1% while the estimated $E_p$ is an order of magnitude higher (20), resulting in a reference excess $CO_2$ level similar to that of "standing−loudly speaking" shown in Figure 2B. $\Delta c^*_{CO_2}$ in the stadium case is between those of "light exercise−speaking" and "light exercise−loudly speaking", as both activities may happen during the event. The difference in $\Delta c^*_{CO_2}$ between the supermarket case and its related activities shown in Figure 2B is mainly due to the long duration of the event (8 h). The $\Delta c^*_{CO_2}$ of the supermarket case divided by the duration leads to the excess $CO_2$ threshold for the worker to inhale over 8 h between those of "light exercise−breathing" and "light exercise−speaking". The $\Delta c^*_{CO_2}$ of the subway case is ~$^1/_3$ lower than that of "resting−breathing" in Figure 2B

because of the short duration (0.33 h) and mask use (universal use of surgical masks or equivalent).

As shown above, the infection risk analysis for various settings can be based on the relevant activities with adjustments for $\eta_I$, $D$, mask use, etc. For policy making concerning an acceptable indoor $CO_2$ level, we also recommend an activity-dependent approach. Reference excess $CO_2$ levels for indoor environments with certain types of activities mainly involved can be found in Figure 2B. Then, this mixing ratio can be scaled for typical $D$ (by multiplying it) and target $P$ (by multiplying its ratio to 0.01%) to obtain an excess $CO_2$ threshold, which may be relaxed a little further depending on the local mask policy. The sum of this value and the local outdoor $CO_2$ concentration, the latter of which we recommend measuring regularly due to possible variations,[29] is the final recommended indoor $CO_2$ concentration threshold. For more complex setups (e.g., with many $CO_2$ meters in a company or school), a meter should be located outdoors to measure $CO_2$ concentration continuously. To the best of our knowledge, $CO_2$ is the only quantity that can be easily measured by fast low-cost sensors as an infection risk proxy. The relative risk of infection in a given situation has been shown to scale with the excess $CO_2$ concentration. The absolute risk can be estimated when the parameters needed are known. Calculations for various scenarios can be easily performed with the online COVID-19 aerosol transmission estimator.[30] Then this method can provide a stronger scientific basis for using $CO_2$ than having one threshold for all situations. However, it may still not be trivial for the general public to estimate the parameters used in our model and implement it. Regulatory authorities may derive the $CO_2$ thresholds for different types of indoor spaces or provide more assistance for businesses to do so. Even if the parameters are unknown, our study suggests that simply keeping the $CO_2$ level and the physical intensity and vocalization level of the activities as low as practically feasible in indoor environments will still reduce the risk.

## ASSOCIATED CONTENT

### Supporting Information

The Supporting Information is available free of charge at https://pubs.acs.org/doi/10.1021/acs.estlett.1c00183.

> Discussions about the relationship between the reference excess $CO_2$ level and the Rudnick–Milton model, details of the mathematical derivations and of how the values of major parameters are assigned, a list of the cases of interest, values of the large fraction term in eqs 6 and 7, and a table of the symbols used in this study (PDF)

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