1	Estimation of airborne viral emission: quanta emission rate of SARS-CoV-2 for
2	infection risk assessment
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12 Abstract

13 Airborne transmission is a pathway of contagion that is still not sufficiently investigated despite the 14 evidence in the scientific literature of the role it can play in the context of an epidemic. While the 15 medical research area dedicates efforts to find cures and remedies to counteract the effects of a 16 virus, the engineering area is involved in providing risk assessments in indoor environments by 17 simulating the airborne transmission of the virus during an epidemic. To this end, virus air emission 18 data are needed. Unfortunately, this information is usually available only after the outbreak, based 19 on specific reverse engineering cases. In this work, a novel approach to estimate the viral load 20 emitted by a contagious subject on the basis of the viral load in the mouth, the type of respiratory 21 activity (e.g. breathing, speaking), respiratory physiological parameters (e.g. inhalation rate), and 22 activity level (e.g. resting, standing, light exercise) is proposed. The estimates of the proposed 23 approach are in good agreement with values of viral loads of well-known diseases from the 24 literature. The quanta emission rates of an asymptomatic SARS-CoV-2 infected subject, with a viral load in the mouth of 10⁸ copies mL⁻¹, were 10.5 quanta h⁻¹ and 320 quanta h⁻¹ for breathing and 25 speaking respiratory activities, respectively, at rest. In the case of light activity, the values would 26 27 increase to 33.9 quanta h⁻¹ and 1.03×10³ quanta h⁻¹, respectively.

The findings in terms of quanta emission rates were then adopted in infection risk models to demonstrate its application by evaluating the number of people infected by an asymptomatic SARS-CoV-2 subject in Italian indoor microenvironments before and after the introduction of virus containment measures. The results obtained from the simulations clearly highlight that a key role is played by proper ventilation in containment of the virus in indoor environments.

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Keywords: SARS-CoV-2 (CoVID19); virus airborne transmission; indoor; ventilation; coronavirus;
 viral load

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37 1. Introduction

38 Expiratory human activities generate droplets, which can also carry viruses, through the atomization 39 processes occurring in the respiratory tract when sufficiently high speeds are reached (Chao et al., 40 2009; Morawska, 2006). Indeed, during breathing, coughing, sneezing or laughing, toques of liquid 41 originating from different areas of the upper respiratory tract are drawn out from the surface, pulled 42 thin, and broken into columns of droplets of different sizes (Hickey and Mansour, 2019). The content 43 of infectious agents expelled by an infected person depends, among other factors, on the location 44 within the respiratory tract from which the droplets originated. In particular, air velocities high 45 enough for atomization are produced when the exhaled air is forced out through some parts of the

respiratory tract which have been greatly narrowed. The front of the mouth is the site of narrowing 46 47 and the most important site for atomization; since most droplets originate at the front of the mouth, 48 the concentration of an infectious agent in the mouth (sputum) is representative of the 49 concentration in the droplets emitted during the expiratory activities (Morawska, 2006). Thus, 50 knowledge of the size and origin of droplets is important to understand transport of the virus via 51 the aerosol route. Contrary to the findings of early investigations (Duguid, 1945; Jennison, 1942; 52 Wells, 1934), subsequent studies involving optical particle detection techniques capable of 53 measurements down to fractions of a micrometer suggested that the majority of these particles are 54 in the sub-micrometer size range (Papineni and Rosenthal, 1997). More recently, the growing 55 availability of higher temporal and spatial visualization methods using high-speed cameras (Tang et 56 al., 2011), particle image velocimetry (Chao et al., 2009) and, above all, increasingly accurate particle 57 counters (Morawska et al., 2009) allowed the detailed characterization and quantitation of droplets 58 expelled during various forms of human respiratory exhalation flows (e.g. breathing, whispering, 59 speaking, coughing). Therefore, in recent years a marked development has occurred both in the 60 techniques for detecting the viral load in the mouth and in the engineering area of the numerical 61 simulation of airborne transmission of the viral load emitted.

However, the problem of estimating the viral load emitted, which is fundamental for the simulation of airborne transmission, has not yet been solved. This is a missing "transfer function" that would allow the virology area, concerned with the viral load values in the mouth, to be connected with the aerosol science and engineering areas, concerned with the spread and mitigation of contagious particles.

67 A novel approach is here presented for estimating the viral load emitted by an infected individual.

68 This approach, based on the principle of conservation of mass, represents a tool to connect the

69 medical area, concerned with the concentration of the virus in the mouth, to the engineering area,

70 dedicated to the simulation of the virus dispersion in the environment. On the basis of the proposed

approach, the quanta emission rate data of SARS-CoV-2 were calculated as a function of different

72 respiratory activities, respiratory parameters, and activity levels.

The quanta emission rate data, starting from the recently documented viral load in sputum (expressed in copies mL⁻¹), were then applied in an acknowledged infection risk model to investigate the effectiveness of the containment measures implemented by the Italian government to reduce the spread of SARS-CoV-2. In particular, airborne transmission of SARS-CoV-2 by an asymptomatic subject within pharmacies, supermarkets, restaurants, banks, and post offices were simulated, and the reduction in the average number of infected people from one contagious person, R₀, was

79 estimated.

80 2. Materials and methods

81 **2.1. Estimation of the quanta emission rate**

The approach proposed in the present work is based on the hypothesis that the droplets emitted by the infected subject have the same viral load as the sputum. Therefore, if the concentration of the virus in the sputum and the quantity of droplets emitted with dimensions less than 10 μ m is known, the viral load emitted can be determined through a mass balance. In particular, the viral load emitted, expressed in terms of quanta emission rate (ER_q, quanta h⁻¹), was evaluated as:

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$$ER_q = c_v \cdot V_{br} \cdot N_{br} \cdot \int_0^{10\mu m} N_d(D) \cdot dV_d(D)$$
(1)

89 where c_v is the viral load in the sputum (RNA copies mL⁻¹), V_{br} is the volume of exhaled air per breath 90 (cm³; also known as tidal volume), N_{br} is the breathing rate (breath h⁻¹), N_d is the droplet number 91 concentration (part. cm⁻³), and $V_d(D)$ is the volume of a single droplet (mL) as a function of the 92 droplet diameter (*D*). Information about the viral load in terms of quanta is essential as the quantum 93 represents the "viral load" considered in engineering science: in other words, an infected individual 94 constantly generates a number of infectious quanta over time, where a "quantum" is defined as the 95 dose of airborne droplet nuclei required to cause infection in 63% of susceptible persons.

96 The volume of the droplet (V_d) was determined on the basis of data obtained experimentally by 97 (Morawska et al., 2009): they measured the size distribution of droplets for different expiratory 98 activities (e.g. breathing, whispering, counting, speaking), recognizing that such droplets present 99 one or more modes occurring at different concentrations. In particular, in the study a particle size 100 distribution with four channels was considered with midpoint diameters of D_1 =0.8, D_2 =1.8, D_3 =3.5, 101 and D_4 =5.5 μ m. As an example, speaking was recognized as producing additional particles in modes 102 near 3.5 and 5.5 µm. These two modes became even more pronounced during sustained 103 vocalization. Details of the aerosol concentrations at the four channels of the size distribution during 104 each expiratory activity are reported in Table 1. The midpoint diameters of each channel were used 105 to calculate the corresponding volume of the droplets.

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 Table 1 - Droplet concentrations (Ni, part. cm⁻³) of the different size distribution channels during each

 expiratory activity measured by (Morawska et al., 2009).

Expiratory activity	<i>D</i> 1 (0.80 μm)	D₂ (1.8 μm)	<i>D</i> ₃ (3.5 μm)	<i>D</i> ₄ (5.5 μm)	
Whispered counting	0.236	0.068	0.007	0.011	
Voiced counting	0.110	0.014	0.004	0.002	
Speaking	0.751	0.139	0.139	0.059	
Breathing	0.084	0.009	0.003	0.002	

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Based on the results obtained by (Morawska et al., 2009), equation (1) can be simplified as: $ER_{q,i} = c_v \cdot IR \cdot \sum_{i=1}^{4} (N_{i,i} \cdot V_i)$ (2)

where *j* indicates the different expiratory activities considered (namely whispered counting, voiced 111 112 counting, speaking, breathing) and IR ($m^3 h^{-1}$) is the inhalation rate, i.e. the product of breathing 113 rate (N_{br}) and tidal volume (V_{br}) , which is a function of the activity level of the infected subject. The quanta emission rate from equation (2) can vary in a wide range depending on the virus 114 115 concentration in the mouth, the activity level, and the different types of expiration. Regarding the 116 inhalation rate effect, the quanta emission rate calculations are shown for three different activity levels (resting, standing, and light exercise) in which the inhalation rates, averaged between males 117 118 and females, are equal to 0.36, 0.54, and 1.16 m³ h⁻¹, respectively (Adams, 1993; International 119 Commission on Radiological Protection, 1994).

120 2.2. A demonstration application: the containment measures for the spread of SARS-CoV-2 in Italy

The pandemic of a novel human coronavirus, now named Severe Acute Respiratory Syndrome 121 122 CoronaVirus 2 (SARS-CoV-2 throughout this manuscript), emerged in Wuhan (China) in late 2019 123 and then spread rapidly in the world (https://www.who.int/emergencies/diseases/novel-124 coronavirus-2019). In Italy, an outbreak of SARS-CoV-2 infections was detected starting from 16 125 cases confirmed in Lombardy (a northern region of Italy) on 21 February. The Italian government 126 has issued government a decree dated 11 March 2020 concerning urgent measures to contain the 127 contagion throughout the country. This decree regulated the lockdown of the country to counteract and contain the spread of the SARS-CoV-2 virus by suspending retail commercial activities, with the 128 129 exception of the sale of food and basic necessities. It represents the starting point of a system with 130 imposed constraints. Among the measures adopted for the containment of the virus in Italy, great importance was placed on the safe distance of 1 m (also known as "droplet distance"). This distance 131

was actually indicated by the World Health Organization as sufficient to avoid transmission by air, 132 133 without any reference to the possibility of transmission over greater distances indoors 134 (https://www.who.int/emergencies/diseases/novel-coronavirus-2019). With this measure, along 135 with the opening of only primary commercial establishments (such as pharmacies, supermarkets, 136 banks, post offices) and the closure of restaurants, the Italian government has adopted the concept of spacing (known as "social distancing") to prevent the spread of the infection. Obviously, this limit 137 138 per se would have no influence on the reduction of airborne transmission of the infection in indoor 139 environments since this distance is compatible with the normal gathering of people in commercial 140 establishments. Actually, on an absolutely voluntary basis, and despite the continuous denials by 141 the government on the risk of indoor airborne transmission, commercial associations have changed 142 the methods of accessing their commercial spaces such as restaurants, pharmacies, supermarkets, 143 post offices, and banks; for example, by forcing customers to queue outside. It is clear that the best 144 choice in containing an epidemic is a total quarantine which, however, appears to have enormous 145 costs and social impacts, especially in Western countries.

To show the possible effect of the measures imposed by the Italian government (i.e. lockdown), the infection risk in different indoor microenvironments for the exposed population due to the presence of one contagious individual was simulated, adopting the infection risk model described in section 2.2.1. In particular, the risk expressed in terms of basic reproduction number (R₀) was derived from the quanta concentration and the infection risk; indeed, R₀ represents the average number of secondary infections produced by a typical case of an infection in a population where everyone is susceptible (Rothman et al., 2008).

153 The indoor microenvironments considered here were a pharmacy, supermarket, restaurant, post 154 office, and bank whose dimensions are summarized in Table 2. Two different exposure scenarios 155 were simulated for each microenvironment: before lockdown (B) and after lockdown (A). In the 156 simulation of the scenario before lockdown, the microenvironments were run with no particular 157 recommendations; thus, people enter the microenvironments and queue indoors, often resulting in 158 overcrowded environments. Since most of the indoor microenvironments in Italy are not equipped 159 with mechanical ventilation systems, the simulations were performed considering two different 160 situations: natural ventilation (a typical value for an Italian building equal to 0.2 h^{-1} was adopted, with reference to (d'Ambrosio Alfano et al., 2012; Stabile et al., 2017)) and mechanical ventilation 161 162 (calculated according the national standard, UNI 10339 (UNI, 1995), as a function of the crowding 163 index and the type of indoor microenvironment). The scenario after lockdown was tested 164 considering the typical solutions adopted (on a voluntary basis) by the owners of stores and offices 165 - reduced personnel, a reduced number of customers inside the microenvironment, customers 166 forced to queue outdoors, and doors kept open. The scenario after lockdown was also tested for 167 both natural ventilation and mechanical ventilation; in this case a slight increase in the air exchange 168 rate (AER) for natural ventilation (0.5 h⁻¹) was considered in order to take into account that the door 169 was always kept open. The restaurant was not tested in the scenario after lockdown since such 170 commercial activity was closed down as a consequence of the lockdown. For all the scenarios 171 considered in the simulations, the infected individual was considered to enter the 172 microenvironment as the first customer (alone or along with other individuals according to the 173 scenarios summarized in Table 2). All the scenarios were simulated taking into account that the virus 174 is able to remain viable in the air for up to 3 hours post aerosolization as recently detected by (van 175 Doremalen et al., 2020); thus, if the infected individual remained inside the environment for 10 minutes (e.g. pharmacy), the calculation of the quanta concentration, infection risk, and Ro was 176 177 performed for up to 3 hours and 10 minutes (named "total exposure time" in Table 2). For 178 restaurants the calculation was performed for 3 hours considering that after 3 hours (i.e. two groups

remaining inside for 1 hour and 30 minutes one after the other) the microenvironment was leftempty.

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Table 2 - Summary of the exposure scenarios tested for the different microenvironments under investigation: dimensions, ventilation conditions, number of workers and customers.

		Pharmacy	Supermarket	Restaurant	Post office	Bank
	Floor area (A, m²)	25	600	100	100	50
Dimensions	Height (h, m ²)	3	3	3	3	3
	Volume (V, m ³)	75	1800	300	300	150
	Number of workers	5 (always present)	10 (always present)	4 (just the waiters, always present)	8 (always present)	4 (always present)
Exposure scenario	Number and activity of the customers	 1 new customer per min entering the pharmacy, every customer remains 10 min inside (including waiting time), thus, 10 customers are simultaneously present 	 1 new customer every 30 s entering the supermarket, every customer remains 30 min inside, thus, 60 customers are simultaneously present 	 80 costumers every 1.5 hours, restaurant working for 3 hours (evening), thus, 80 customers are simultaneously present for a total number of 160 customers per evening. 	 1 new customer every 30 s entering the post office, every customer remains 15 min inside (including waiting time), thus, 30 customers are simultaneously present 	 1 new customer per min entering the bank, every customer remains 15 min inside (including waiting time), thus, 15 customers are simultaneously present
before lockdown (B)	Air exchange rate (AER, h ⁻¹) for natural ventilation (NV)	0.2	0.2	0.2	0.2	0.2
	Air exchange rate (AER, h ⁻¹) for mechanical ventilation (MV)	2.2	1.1	9.6	2.4	2.4
	Total exposure time	3 hours and 10 minutes	3 hours and 30 minutes	3 hours	3 hours and 15 minutes	3 hours and 15 minutes
	Number of workers	3 (always present)	10 (always present)	-	4 (always present)	4 (always present)
Exposure scenario after lockdown (A)	Number and activity of the costumers	 2 new customers every five min entering the pharmacy, every customer remains 5 min inside, people forced to queue outside the pharmacy, thus, 2 customers are simultaneously present 	 1 new customer per min entering the supermarket, every customer remains 10 min inside, people forced to queue outside the supermarket, thus, 10 customers are simultaneously present 	-	 4 new customers every five min entering the post office, every customer remains 10 min inside, people forced to queue outside the post office, thus, 4 customers are simultaneously present 	 4 new customers every five min entering the bank, every customer remains 10 min inside, people forced to queue outside the bank, thus, 4 customers are simultaneously present
	Air exchange rate (AER, h ⁻¹) for natural ventilation (NV)	0.5	0.2	-	0.5	0.5
	Air exchange rate (AER, h ⁻¹) for mechanical ventilation (MV)	2.2	1.1	-	2.4	2.4
	Total exposure time	3 hours and 5 minutes	3 hours and 10 minutes	3 hours	3 hours and 10 minutes	3 hours and 10 minutes

183 2.2.1. The infection risk model

184 The simulation of airborne transmission of SARS-CoV-2 was performed adopting the infection risk assessment typically implemented to evaluate the transmission dynamics of infectious diseases and 185 to predict the risk of these diseases to the public. The model considered here to quantify the 186 187 airborne transmitted infection risk was carried out by Gammaitoni and Nucci (Gammaitoni and 188 Nucci, 1997) which represents an upgrade of an earlier model provided by Wells-Riley (Riley et al., 189 1978). This model was successfully adopted in previous papers estimating the infection risk due to 190 other diseases (e.g. influenza, SARS, tuberculosis, rhinovirus) in different indoor microenvironments 191 such as airplanes (Wagner et al., 2009), cars (Knibbs et al., 2011), and hospitals. The Gammaitoni 192 and Nucci model is based on the rate of change in quanta levels through time; in particular, the 193 differential equations for the change of quanta in a control volume as well as the initial conditions 194 (here not reported for the sake of brevity) allowed to evaluate the quanta concentration in an 195 indoor environment at the time t, n(t), as:

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$$n(t) = \frac{ER_q \cdot I}{AER \cdot V} + \left(n_0 + \frac{ER_q \cdot I}{AER}\right) \cdot \frac{e^{-AER \cdot t}}{V} \qquad (\text{quanta m}^{-3})$$
(3)

197 where AER (h^{-1}) represents the air exchange rate of the space investigated, n_0 represents the initial 198 number of quanta in the space, *I* is the number of infectious subjects, *V* is the volume of the indoor 199 environment considered, and ER_q is the abovementioned quanta emission rate (quanta h^{-1}) 200 characteristic of the specific disease/virus under investigation.

- The equation was derived considering the following simplifying assumptions: the quanta emission 201 202 rate is considered to be constant, the latent period of the disease is longer than the time scale of 203 the model, and the droplets are instantaneously and evenly distributed in the room (Gammaitoni 204 and Nucci, 1997). The latter represents a key assumption for the application of the model as it 205 considers that the air is well-mixed within the modelled space. The authors highlight that in 206 epidemic modeling, where the target is the spread of the disease in the community, it is impossible 207 to specify the geometries, the ventilation, and the locations of the infectious sources in each 208 microenvironment. Therefore, adopting the well-mixed assumption is generally more reasonable 209 than hypothesizing about specific environments and scenarios because the results must be 210 interpreted on a statistical basis (Sze To and Chao, 2010).
- To determine the infection risk (*R*, %) as a function of the exposure time (*t*) of susceptible people,
 the quanta concentration was integrated over time through the Wells–Riley equation (Riley et al.,
 1978) as:

$$R = \left(1 - e^{-IR \int_0^T n(t)dt}\right) \tag{4}$$

215 where IR is the inhalation rate of the exposed subject (which is, once again, affected by the subject's 216 activity level) and T is the total time of exposure (h). From the infection risk R, the number of 217 susceptible people infected after the exposure time can be easily determined by multiplying it by 218 the number of exposed individuals. In fact, equations (3) and (4) were adopted to evaluate the 219 infection risk of different exposure scenarios of Italian microenvironments hereinafter reported. 220 The quanta emission rate used in the simulation of the scenario represents the average value 221 obtained from the four expiratory activities (whispered counting, voiced counting, speaking, and breathing); the data are reported and discussed in the result sections. 222

223 3. Results and discussions

224 **3.1. The quanta emission rate**

As discussed in the Materials and methods section, the quanta emission rate, ER_q, depends on several parameters. In Figure 1 the ER_q (quanta h⁻¹) trends are reported as a function of the viral load in the sputum (c_v, RNA copies mL⁻¹) for different expiratory activities (whispered counting, voiced counting, speaking, breathing) and different activity levels (resting, standing, light exercise). To represent the large variabilities (over several orders of magnitude) of ER_q as a function of c_v, the graph is reported on a bi-logarithmic scale.

231 To benchmark the proposed approach for the estimation of the quanta emission rate, we 232 considered the case of seasonal influenza for which more data are available in terms of both viral 233 load in sputum and quanta emission rate. As an example, (Hirose et al., 2016) found an average 234 value of RNA concentration in sputum for influenza equal to 2.38×10⁷ copies mL⁻¹. Thus, applying 235 the findings of the proposed approach in the case of a standing subject, a corresponding ER_a varying between 3.7 (breathing) and 114 quanta h⁻¹ (speaking) is estimated: this value is in good agreement 236 with the guanta emission rates for influenza found in the scientific literature, from 2 to 128 237 238 guanta h⁻¹ with a most frequent value of 67 guanta h⁻¹ (Knibbs et al., 2012). Such variability in the 239 quanta emission rates for influenza is due both to the method used to calculate it (Rudnick and 240 Milton, 2003) and, especially, the viral load of the subject and the type of respiratory activity, which 241 is typically not reported and discussed.

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With reference to SARS-CoV-2 infection, researchers have recently found values for the viral load in the mouth between 10² and 10¹¹ copies mL⁻¹, also variable in the same patient during the course of the disease (Pan et al., 2020; To et al., 2020; Woelfel et al., 2020). (Rothe et al., 2020) reported a case of SARS-CoV-2 infection acquired outside Asia in which transmission appears to have occurred during the incubation period in the index patient. A high viral load of 10⁸ copies mL⁻¹ was found, confirming that asymptomatic persons are potential sources of SARS-CoV-2 infection; this may warrant a reassessment of the transmission dynamics of the current outbreak. Table 3 lists the quanta emission rates (ER_q) for a SARS-CoV-2 infected asymptomatic subject as a function of activity level (resting, standing, and light exercise) and respiratory activity (voiced counting, whispered counting, speaking, breathing). The data confirm the huge variations in the quanta emission rate, with the lowest value being for breathing during resting activity (10.5 quanta h⁻¹) and the highest value being for speaking during light activity (more than 1000 quanta h⁻¹).

260**Table 3** – Quanta emission rates (ER_q) for a SARS-CoV-2 infected asymptomatic subject ($c_v=10^8$ copies mL⁻¹)261as a function of the activity level and respiratory activity.

Activity loval	Respiratory activity				
Activity level	Voiced counting	Whispered counting	Speaking	Breathing	Avg
Resting	49.9	12.1	320	10.5	98.1
Standing	74.8	18.1	480	15.7	147
Light exercise	161	39.1	1.03×10 ³	33.9	317

262 **3.2. Results of the demonstration application**

In this section, the results of the simulations performed for the microenvironments and exposurescenarios described in section 2.2 and summarized in Table 2 are reported.

265 3.2.1. Infection risk and R₀ for different indoor environments and exposure scenarios

266 As an illustrative example, Figure 2 shows the quanta concentration (n(t)) and infection risk (R) 267 trends as a function of time for two different exposure scenarios simulated for the pharmacy, i.e. 268 before lockdown (B) in natural (NV) and mechanical ventilation (MV) conditions. The trends clearly 269 highlight that the presence of the infected individual remaining inside for 10 minutes leads to an 270 increase in the quanta concentration in the volume: in particular, a higher peak of quanta 271 concentration was recognized, as expected, for reduced ventilation (NV) with respect to the 272 mechanical ventilation (MV). People entering the pharmacy after the infected individual are 273 exposed to a certain quanta concentration during their 10-min time, and the resulting risk for their 274 exposure (evaluated through equation (4)) is just a function of the quanta concentration trend. For 275 example, people entering the microenvironment around the quanta concentration peak are at a 276 higher risk than people entering the pharmacy later. Figure 2 shows an example of a customer 277 entering at min 26 and leaving at min 36: the risk for this 10-min exposure is 2.4% in natural 278 ventilation conditions and 1.0% in mechanical ventilation conditions. During the entire exposure 279 time of such a scenario (3 hours and 10 minutes), 179 customers (after the infected individual) enter 280 the pharmacy and each of them receive their own risk. In particular, the average risk of the 179 281 customers is 2.0% for NV conditions and 0.4% for MV conditions, then leading to a R_0 (among the customers) of 3.52 and 0.68, to which must be added the R₀ of the five pharmacists exposed for the 282 283 entire period. Similar trends, not shown here graphically for the sake of brevity, were obtained for 284 all the scenarios investigated, then leading to the evaluation of the R₀ for each of them as described 285 in the methodology section.

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Figure 2 - Details of application of the proposed approach in the calculation of quanta concentrations, n(t),
 and infection risks, R, in the pharmacy environment for the exposure scenarios before lockdown (B) in
 natural (NV) and mechanical ventilation (MV) conditions. The graph shows the entry of the infected
 individual (first 10 minutes) and the risk for a customer entering the microenvironment at min 26 and
 remaining inside for 10 minutes. The trends are shown for up to 100 minutes to highlight the peaks of the
 n(t) and R values.

294 Figure 3 shows the reproduction number (R_0) data calculated for all the exposure scenarios and 295 microenvironments under investigation (summarized in Table 2). The R₀ data were calculated for an 296 asymptomatic SARS-CoV-2 infected subject (c_v=1×10⁸ copies mL⁻¹) while standing; in particular, the average ER_q value among the different respiratory activities was considered (147 quanta h⁻¹, Table 297 298 **3**). The exposed subjects were also considered to be standing (IR=0.54 m³ h⁻¹). The graph clearly 299 highlights some critical exposure scenarios and microenvironments. Indeed, in all the 300 microenvironments, a $R_0>1$ was estimated for all the exposure scenarios before lockdown (B) when 301 the ventilation relied only upon the building being airtight (i.e. natural ventilation conditions): R_0 302 was equal to 5.55, 3.51, 59.3, 5.59, and 6.04 for pharmacy, supermarket, restaurant, post office, 303 and bank, respectively. The huge value for the restaurant is obviously due to the simultaneous co-304 presence of many people (80 customers and 4 waiters) and to the long exposure time (1 h and 305 30 min in the current simulations). This situation is obviously improved if mechanical ventilation 306 systems are adopted, but the R₀ is still higher than 1 (R₀=3.40). Similar results are obviously expected 307 for all the indoor environments characterized by high crowding indexes and long-lasting exposures 308 such as schools, swimming pools, gyms - venues that, in fact, were also concomitantly locked down 309 by the government. Actually, adopting mechanical ventilation solutions that purportedly provide an 310 adequate indoor air quality (i.e. providing AER values suggested by the standards (UNI, 1995)) did 311 not satisfactorily reduce the R₀ in the other microenvironments investigated. Indeed, the R₀ values 312 obtained from the simulations performed for the pharmacy, supermarket, post office, and bank 313 equipped with mechanical ventilation systems in the conditions before lockdown, with mechanical 314 ventilation in operation, were still >1.

The new regulations and methods of accessing the indoor environments that were applied in the conditions after lockdown (i.e. queuing outside, limited time spent in the environments, lower crowding index) were very effective; indeed, the R₀ values were reduced by roughly 80%–90% (for both natural and mechanical ventilation conditions) with respect to the corresponding prelockdown scenarios.

320 As an example, for the natural ventilation scenario, the only critical microenvironment was the bank, 321 since the R₀ was >1; this was due to a crowding index that was higher than the post-office, which 322 had a larger floor area but same number of customers. In contrast, all the R_0 values for indoor environments equipped with mechanical ventilation systems were much lower than 1 (0.18, 0.12, 323 324 0.15, and 0.30 for pharmacy, supermarket, post office, and bank, respectively). Therefore, if in a 325 single day the infected individual visited different environments, the resulting R₀ would be lower 326 than 1 only if all the microenvironments were equipped with mechanical ventilation systems. Once 327 again, these results highlight the importance of proper ventilation of indoor environments and are 328 in line with the scientific literature that recognizes the importance of ventilation strategies in 329 reducing indoor-generated pollution (Stabile et al., 2017)(Stabile et al., 2019).

The values obtained with this approach could vary significantly as a function of (i) the activity levels of both the infected subject and the exposed subjects; and (ii) the viral load in the sputum of the infected subject; therefore, in future studies, more specific exposure scenarios could be simulated on the basis of the findings proposed and discussed in this study.

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Figure 3 - R₀ calculated for all the exposure scenarios (natural ventilation, mechanical ventilation; before
 lockdown, after lockdown) and microenvironments (pharmacy, supermarket, restaurant, post office, bank)
 under investigation considering an asymptomatic SARS-CoV-2 infected subject (c_v=1×10⁸ copies mL⁻¹) while
 standing (IR=0.54 m³ h⁻¹; ER_q=147 quanta h⁻¹) and the exposed population, also standing.

340 4. Conclusions

341 The present study proposed the first approach aimed at filling the gap of knowledge still present in 342 the scientific literature about evaluating the viral load emitted by infected individuals. This 343 information could provide key information for engineers and indoor air quality experts to simulate 344 airborne dispersion of diseases in indoor environments. To this end, we have proposed an approach 345 to estimate the quanta emission rate (expressed in quanta h⁻¹) on the basis of the emitted viral load 346 from the mouth (expressed in RNA copies in mL⁻¹), typically available from virologic analyses. Such 347 approach also takes into account the effect of different parameters (including inhalation rate, type 348 of respiratory activity, and activity level) on the quanta emission rate. The suitability of the findings 349 was checked and confirmed as it was able to predict the values of quanta emission rates of previous 350 well-known diseases in accordance with the scientific literature. The proposed approach is of great 351 relevance as it represents an essential tool to be applied in enclosed space and it is able to support 352 air quality experts and epidemiologists in the management of indoor environments during an 353 epidemic just knowing its viral load, without waiting for the end of the outbreak. 354 For this purpose, it has been applied to the Italian case which, at the time of writing, represents the 355 country with the highest number of deaths from SARS-CoV-2 in the world, highlighting the great 356 importance of ventilation in indoor microenvironments to reduce the spread of the infection.

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359 References

- Adams, W.C., 1993. Measurement of Breathing Rate and Volume in Routinely Performed Daily Activities.
 Final Report. Human Performance Laboratory, Physical Education Department, University of
 California, Davis. Human Performance Laboratory, Physical Education Department, University of
 California, Davis. Prepared for the California Air Resources Board, Contract No. A033-205, April 1993.
- Chao, C.Y.H., Wan, M.P., Morawska, L., Johnson, G.R., Ristovski, Z.D., Hargreaves, M., Mengersen, K., Corbett,
 S., Li, Y., Xie, X., Katoshevski, D., 2009. Characterization of expiration air jets and droplet size
 distributions immediately at the mouth opening. Journal of Aerosol Science 40, 122–133.
 https://doi.org/10.1016/j.jaerosci.2008.10.003
- d'Ambrosio Alfano, F.R., Dell'Isola, M., Ficco, G., Tassini, F., 2012. Experimental analysis of air tightness in
 Mediterranean buildings using the fan pressurization method. Building and Environment 53, 16–25.
 https://doi.org/10.1016/j.buildenv.2011.12.017
- Duguid, J.P., 1945. The numbers and the sites of origin of the droplets expelled during expiratory activities.
 Edinburgh Medical Journal LII (II), 385–401.
- Gammaitoni, L., Nucci, M.C., 1997. Using a mathematical model to evaluate the efficacy of TB control
 measures. Emerging Infectious Diseases 335–342.
- Hickey, A.J., Mansour, H.M., 2019. Inhalation Aerosols: Physical and Biological Basis for Therapy, Third
 Edition. Taylor & Francis Ltd.
- Hirose, R., Daidoji, T., Naito, Y., Watanabe, Y., Arai, Y., Oda, T., Konishi, H., Yamawaki, M., Itoh, Y., Nakaya, T.,
 2016. Long-term detection of seasonal influenza RNA in faeces and intestine. Clinical Microbiology
 and Infection 22, 813.e1-813.e7. https://doi.org/10.1016/j.cmi.2016.06.015
- International Commission on Radiological Protection, 1994. Human respiratory tract model for radiological protection. A report of a Task Group of the International Commission on Radiological Protection.
 Annals of the ICRP 24, 1–482. https://doi.org/10.1016/0146-6453(94)90029-9
- Jennison, M.W., 1942. Atomizing of mouth and nose secretions into the air as revealed by high speed
 photography. Aerobiology 17, 106–128.
- Knibbs, L.D., Morawska, L., Bell, S.C., 2012. The risk of airborne influenza transmission in passenger cars.
 Epidemiology and Infection 140, 474–478. https://doi.org/10.1017/S0950268811000835
- Knibbs, L.D., Morawska, L., Bell, S.C., Grzybowski, P., 2011. Room ventilation and the risk of airborne infection
 transmission in 3 health care settings within a large teaching hospital. American Journal of Infection
 Control 39, 866–872.
- Morawska, L., 2006. Droplet fate in indoor environments, or can we prevent the spread of infection? Indoor
 Air 16, 335–347. https://doi.org/10.1111/j.1600-0668.2006.00432.x
- Morawska, L., Johnson, G.R., Ristovski, Z.D., Hargreaves, M., Mengersen, K., Corbett, S., Chao, C.Y.H., Li, Y.,
 Katoshevski, D., 2009. Size distribution and sites of origin of droplets expelled from the human
 respiratory tract during expiratory activities. Journal of Aerosol Science 40, 256–269.
 https://doi.org/10.1016/j.jaerosci.2008.11.002
- Pan, Y., Zang, D., Yang, P., Poon, L.M., Wang, Q., 2020. Viral load of SARS-CoV-2 in clinical samples Yang Pan
 Daitao Zhang Peng Yang Leo L M Poon Quanyi Wang. The Lancet.
- Papineni, R.S., Rosenthal, F.S., 1997. The size distribution of droplets in the exhaled breath of healthy human
 subjects. Journal of Aerosol Medicine.
- Riley, C., Murphy, G., Riley, R.L., 1978. Airborne spread of measles in a suburban elementary school. American
 journal of epidemiology 431–432.
- Rothe, C., Schunk, M., Sothmann, P., Bretzel, G., Froeschl, G., Wallrauch, C., Zimmer, T., Thiel, V., Janke, C.,
 Guggemos, W., Seilmaier, M., Drosten, C., Vollmar, P., Zwirglmaier, K., Zange, S., Wölfel, R.,
 Hoelscher, M., 2020. Transmission of 2019-nCoV Infection from an Asymptomatic Contact in
 Germany. N Engl J Med 382, 970–971. https://doi.org/10.1056/NEJMc2001468
- 406 Rothman, K.J., Greenland, S., Lash, T.L., 2008. Modern Epidemiology, 3rd ed. Lippincott Williams & Wilkins.
- Rudnick, S.N., Milton, D.K., 2003. Risk of indoor airborne infection transmission estimated from carbon
 dioxide concentration. Indoor Air 13, 237–245. https://doi.org/10.1034/j.1600-0668.2003.00189.x

- Stabile, L., Buonanno, G., Frattolillo, A., Dell'Isola, M., 2019. The effect of the ventilation retrofit in a school
 on CO2, airborne particles, and energy consumptions. Building and Environment 156, 1–11.
 https://doi.org/10.1016/j.buildenv.2019.04.001
- Stabile, L., Dell'Isola, M., Russi, A., Massimo, A., Buonanno, G., 2017. The effect of natural ventilation strategy
 on indoor air quality in schools. Science of the Total Environment 595, 894–902.
 https://doi.org/10.1016/j.scitotenv.2017.02.030
- Sze To, G.N., Chao, C.Y.H., 2010. Review and comparison between the Wells–Riley and dose-response
 approaches to risk assessment of infectious respiratory diseases. Indoor Air 20, 2–16.
 https://doi.org/10.1111/j.1600-0668.2009.00621.x
- 418 Tang, J.W., Noakes, C.J., Nielsen, P.V., Eames, I., Nicolle, A., Li, Y., Settles, G.S., 2011. Observing and 419 quantifying airflows in the infection control of aerosol- and airborne-transmitted diseases: an 420 overview of approaches. Journal of Hospital Infection 77, 213-222. 421 https://doi.org/10.1016/j.jhin.2010.09.037
- To, K.K.-W., Tsang, O.T.-Y., Leung, W.-S., Tam, A.R., Wu, T.-C., Lung, D.C., Yip, C.C.-Y., Cai, J.-P., Chan, J.M.-C.,
 Chik, T.S.-H., Lau, D.P.-L., Choi, C.Y.-C., Chen, L.-L., Chan, W.-M., Chan, K.-H., Ip, J.D., Ng, A.C.-K., Poon,
 R.W.-S., Luo, C.-T., Cheng, V.C.-C., Chan, J.F.-W., Hung, I.F.-N., Chen, Z., Chen, H., Yuen, K.-Y., 2020.
 Temporal profiles of viral load in posterior oropharyngeal saliva samples and serum antibody
 responses during infection by SARS-CoV-2: an observational cohort study. The Lancet Infectious
 Diseases. https://doi.org/10.1016/S1473-3099(20)30196-1
- 428 UNI, 1995. UNI 10339 Impianti aeraulici al fini di benessere. Generalità, classificazione e requisiti. Regole
 429 per la richiesta d'offerta, l'ordine e la fornitura.
- van Doremalen, N., Bushmaker, T., Morris, D.H., Holbrook, M.G., Gamble, A., Williamson, B.N., Tamin, A.,
 Harcourt, J.L., Thornburg, N.J., Gerber, S.I., Lloyd-Smith, J.O., de Wit, E., Munster, V.J., 2020. Aerosol
 and Surface Stability of SARS-CoV-2 as Compared with SARS-CoV-1. N Engl J Med.
 https://doi.org/10.1056/NEJMc2004973
- Wagner, B.G., Coburn, B.J., Blower, S., 2009. Calculating the potential for within-flight transmission of
 influenza A (H1N1). BMC Medicine 7, 81. https://doi.org/10.1186/1741-7015-7-81
- Wells, W.F., 1934. On airborne infection: study II. Droplets and Droplet nuclei. American Journal of
 Epidemiology 20, 611–618. https://doi.org/10.1093/oxfordjournals.aje.a118097
- Woelfel, R., Corman, V.M., Guggemos, W., Seilmaier, M., Zange, S., Mueller, M.A., Niemeyer, D., Vollmar, P.,
 Rothe, C., Hoelscher, M., Bleicker, T., Bruenink, S., Schneider, J., Ehmann, R., Zwirglmaier, K., Drosten,
 C., Wendtner, C., 2020. Clinical presentation and virological assessment of hospitalized cases of
 coronavirus disease 2019 in a travel-associated transmission cluster. medRxiv 2020.03.05.20030502.
 https://doi.org/10.1101/2020.03.05.20030502
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