

1 **This is the accepted version of the manuscript.**

2 **SARS CoV-2 Aerosol: How far it can travel to the**  
3 **lower airways?**

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18 **Abstract**

19 The recent outbreak of the SARS CoV-2 virus causes a significant effect on human respiratory  
20 health around the world. The contagious disease infected a large proportion of the world  
21 population resulting in long-term health issues and an excessive mortality rate. The SARS  
22 CoV-2 virus can spread as small aerosols and enters into the respiratory systems through the

23 oral (nose or mouth) airway. The SARS CoV-2 particle transport to the mouth-throat and upper  
24 airways is analysed by the available literature. Due to the tiny size, the virus can travel to the  
25 terminal airways of the respiratory system and form a severe health hazard. There is a gap in  
26 the understanding of the SARS CoV-2 particle transport to the terminal airways. The present  
27 study investigated the SARS CoV-2 virus particle transport and deposition to the terminal  
28 airways in a complex 17-generation lung model. This first-ever study demonstrates how far  
29 SARS CoV-2 particle can travel in the respiratory system. ANSYS Fluent solver was used to  
30 simulate the virus particle transport during sleep, light and heavy activity conditions.  
31 Numerical results demonstrate that a higher percentage of the virus particles are trapped at the  
32 upper airways when sleeping and in a light activity condition. More virus particles have lung  
33 contact in the right lung than the left lung. A comprehensive lobe specific deposition and  
34 deposition concentration study was performed. The results of this study provide a precise  
35 knowledge of the SARS CoV-2 particle transport to the lower branches and could help the lung  
36 health risk assessment system.

37 **Keywords:** SARS CoV-2, Terminal airways, Virus transport, 17-Generation lung.

## 38 **Introduction**

39 thoracic cavity and consists of inhalation and exhalation processes. During inhalation,  
40 airborne pollutants, as for example, particulate matter, dust, smoke, pollens, viruses, or  
41 allergens, often in the form of liquid droplets and aerosols are ingested into the airways.  
42 Aerosol is a term first introduced in the 1920s and was initially used in the context of  
43 therapeutic inhalation (Anderson, 2005).

44 The inhaled air is ingested into the respiratory tract, commonly termed as human airways,  
45 which has a complicated geometry and hence is difficult to reconstruct even using  
46 computerised modelling. Due to lack of CT-scan images, earlier researchers (Weibel, 1963;

47 Philips and Kaye, 1995; Kitaoka et al., 1999) used simplified (often with a regular cross-  
48 sectional shape) geometry of human airways. With the sophisticated imaging techniques, the  
49 researchers started developing more realistic geometry of human airways incorporating lung  
50 intricate shapes and minuscule anatomical features (Martonen et al., 1995; Lizal et al., 2012;  
51 Srivastav et al., 2013; Frederix et al., 2018) for conducting various computational studies.  
52 These studies include the effects of breathing through nasal and oral passages (Fitzpatrick et  
53 al., 2003), transmission and deposition of inhaled aerosols, fine and ultrafine particles etc.  
54 within the both upper and tracheobronchial airways (Fernández Tena and Casan Clarà, 2012;  
55 Mortazavi et al., 2020). In the last two decades, respiratory fluid dynamics has matured enough  
56 to allow multiscale, multiphysics modelling (Burrowes et al., 2008; Pozin, 2017) to analyse the  
57 various aspects of respiratory mechanics, starting from the inhalation mechanism (Islam et al.,  
58 2020) to aerosol (Xi and Longest, 2007; Islam et al., 2017; Lizal et al., 2020) and drug delivery  
59 through pulmonary routes (Heyder, 2004; Kleinstreuer et al., 2008) and even diseased airways  
60 (Martonen et al., 2003; Srivastav et al., 2014).

61 Refer to the term of aerosol, it is the combination between solid or liquid particles which are  
62 suspended in gas. Airborne transmission of many viral diseases is caused due to propagation  
63 of such airborne particles containing saliva, mucus, salts, cells and even infectious pathogens-  
64 viral and/or bacterial particles (Wells, 1955). The droplets are often originated from the viral  
65 infected inner epithelial layers of the respiratory tract surfaces (Mason, 2020) through  
66 exhalation, coughing, sneezing, talking, or vomiting by an infected person (Atkinson et al.,  
67 2009). The ongoing COVID-19 pandemic has triggered the publication of numerous research  
68 articles encompassing various aspects and behaviour of the SARS CoV-2 virus (Zhou and Zou,  
69 2021). The virus mainly attacks human lung airways and eventually damages lung capacity of  
70 gas exchange (Mason, 2020). Hence, the study of the transport of SARS CoV-2 aerosol  
71 to the tracheobronchial airways (termed as lower airways) is important.

72 Prather et al. (2020) suggested that pulmonary infections are caused by small aerosols. Virus  
73 particle can suspend in air for longer period and can be transmitted from an infected person to  
74 a non-infected person. Larger virus particle, on the other hand, can survive on the surfaces for  
75 longer period and can be transmitted through contacts (Bhardwaj and Agrawal, 2020a,  
76 2020b). Kumar and Lee (2020) adopted continuous phase modelling for smaller aerosols,  
77 while discrete phase simulation was conducted for the larger aerosols. Diffusion equation-  
78 based Monte-Carlo modelling was used by Vuorinen et al. (2020) for the transport of virus  
79 aerosols. Appropriate source and sink terms are added to the diffusion transport equations to  
80 represent the variable location of the infected people and source of ventilation, respectively.  
81 Xie et al. (2009) stressed the importance of including droplet distribution (i.e. variation in size)  
82 for discrete phase modelling since it is connected to the travelling path as well as the chances  
83 of viral infections (called viral load) in case of SARS CoV-2. Liquid particles evaporation is  
84 another important phenomenon while transmitting the virus-laden aerosols, which is  
85 dependent on ambient temperature and saturation pressure. A recent study reported in the  
86 literature that small droplets released from the exhalation may be laden with Covid-19 virus has very  
87 short evaporation time scale ( $\leq 1s$ ) and hence are evaporated as soon as it is ejected. Hence, the virus is  
88 considered as a particle (Chaudhuri et al. 2020). The larger droplet or virus-laden particles are  
89 larger and cannot travel to the lower airways, as larger particles usually deposit to the upper  
90 airways. A recent study (Kwee and Kwee 2020a) showed that nano-sized SARS CoV-2  
91 Aerosol are deposited to various lobes of the lung in their radiographic images, which supports  
92 the assumption of this study. However, after inhalation the evaporation is largely regulated by  
93 the body temperature. Recently, a number of researchers (Feng et al., 2020; Chaudhuri et al.,  
94 2020, de Oliveira et al., 2021) investigated various aspects of liquid particles evaporation and  
95 transmission in light of the COVID-19 pandemic. Smaller aerosols generated from continuous  
96 speech in a poorly ventilated room increased the infection risk by 11% as revealed in a recent

97 study (de Oliviera et al. 2021) and hence reiterates the importance of maintaining proper  
98 ventilation and physical distancing to avoid infection transmission.

99 With the outbreak of Covid-19 pandemic, a few researchers have also worked on the virus  
100 transmission in human airways. Balázs et al. (2020) employed a lung model based on a  
101 stochastic deposition, which was developed by Koblinger and Hofmann (1990) to find out the  
102 deposition of viral loads and revealed that over 60% of the inhaled viral masses were  
103 deposited in the extrathoracalis (upper) which are the portion of the human lung airways, and  
104 suggested to affect the upper airways and if not diagnosed, could eventually develop into  
105 pneumonia. Other researchers focused on aerosol behaviour in the intra-distal region of a  
106 simplistic lung model in the presence of different breathing conditions (Ciloglu, 2020), gravity  
107 and surface tension effects on micro-bubbles in simplistic bifurcated airways (Munir and Xu,  
108 2020), mask-wearing effects in upper respiratory geometry (Xi et al., 2020), aerosol transport  
109 in phantom lung bronchioles (Mallik et al., 2020), cough exhalation from a 18-generation  
110 simplistic airways (Si et al., 2021) etc.

111 The review of literature reveals a plethora of works on the transmission of infections  
112 and exhalation behaviour originated from oral and nasal openings of the human airways.  
113 However, the transportation of virus- aerosols to the tracheobronchial human airways involving  
114 a realistic and detailed geometry of the airways has not yet discussed and analysed in detail.  
115 Since empirical evidence of SARS CoV-2 attacking the respiratory organ in the COVID-19  
116 infected population exists, a CFD investigation of virus aerosol transport in a realistic human  
117 airways up to the 17th generation would help medical practitioners and inform further  
118 diagnosis and prognosis of the COVID-19 disease. The CFD studies of airflow and CoV-2  
119 virus deposition in a digital reference model of the 17-generation airway were based on the  
120 anatomical model of an adult male, free of pathological alterations by Schmidt et al. (2004).  
121 The lung model consists 1453 bronchi up to the 17<sup>th</sup> Horsfield order.

122 **Numerical Methods**

123 The numerical study solved the air and particle transport equations and analysed the particle  
124 flow in first 17-bifurcations of the human lung. The Lagrangian scheme and Finite Volume  
125 based discretisation techniques are used for this study. The numerical calculation method in  
126 this study is performed and conducted based on ANSYS 19.2 (FLUENT). The governing  
127 equations are solved as following;

128 
$$\nabla \cdot (\rho \vec{v}) = 0 \tag{1}$$

129 
$$\nabla \cdot (\rho \vec{v} \vec{v}) = -\nabla p + \nabla \cdot (\mu (\nabla \vec{v} + \nabla \vec{v}^T)) + \rho \vec{g} \tag{2}$$

130 where, static pressure is  $p$ , gravitational body force  $\rho \vec{g}$ , molecular viscosity  $\mu$ .

131 The internal energy equation is

132

133 
$$\nabla \cdot (\rho \vec{v} e) = -\nabla \cdot \vec{J} \tag{3}$$

134 where  $e$  is the specific internal energy. The heat flux vector  $\vec{J}$  is the sum of contributions due  
135 to heat conduction and enthalpy diffusion effects.

136 The inlet condition and velocity profiles are highly complex and irregular for person to person.  
137 No proper velocity profiles are established by the available literature. However, the flow inside  
138 the airway is similar to internal pipe flow and the flow become parabolic at the tracheal area of  
139 the airway. This study only considers the simulation which starts from trachea to lower  
140 generations of the human lung as well as the trachea wall which is considered as the inlet of  
141 the airway. Therefore, a fully developed parabolic inlet condition (White 2003) is used

142 
$$u(r) = 2u_{av} \left(1 - \frac{r^2}{R^2}\right) \tag{4}$$

143 where  $R$  is the pipe radius. The corresponding velocity for 7.5 lpm, 15 lpm and 30 lpm cases  
144 are 0.4916 m/s, 0.9829 m/s and 1.996 m/s, respectively, which is the maximum velocity. For  
145 fully developed condition, maximum velocity is double of the average velocity.

146 **SARS CoV-2 particle are smaller in size and it's approximately around 120 nm.**

147 ([https://www.pptaglobal.org/media-and-information/ppta-statements/1055-2019-novel-  
148 coronavirus-2019-ncov-and-plasma-protein-therapies](https://www.pptaglobal.org/media-and-information/ppta-statements/1055-2019-novel-<br/>148 coronavirus-2019-ncov-and-plasma-protein-therapies)).

149 Therefore, nano-particle transport equations are solved (Inthavong, Tu and Ahmadi 2009).

$$\frac{du_i^p}{dt} = F_D + F_{Brownian} + F_{Lift} + \frac{\rho_p - \rho_g}{\rho_p} g_i$$

$$150 \quad F_D = \frac{1}{C_c} C_D A_p \frac{\rho_g |u_i^g - u_i^p| (u_i^g - u_i^p)}{2m_p} = \frac{18\mu_g}{\rho_p d_p^2 C_c} (u_i^g - u_i^p) \quad (5)$$

$$C_c = 1 + \frac{2\lambda}{d_p} (1.257 + 0.4e^{\frac{-0.11d_p}{2\lambda}})$$

151 where  $F_D$  is the drag force per unit particle mass  $m_p$ ,  $C_D$  is the drag coefficient,  $A_p$  is the cross  
152 sectional area of the particle, and  $C_c$  is the Cunningham correction factor.  $\lambda$  is the mean free path of  
153 the gas molecules.  $u_i^p$  is the i-th component of the time-averaged particle velocity while  $u_i^g$  is the i-th  
154 component of the time-averaged gas (air) velocity.  $\rho_p$  and  $\rho_g$  are the density of particle material and  
155 gas (air), respectively.  $g_i$  is the gravitational component.  $\mu_g$  denotes the gas (air) viscosity and  $d_p$  is  
156 defined as particle diameter. For the low particle Reynolds number ( $Re_p < 0.5$ ), the drag coefficient  
157  $C_D$  can be defined as (Haider and Levenspiel 1989);

$$158 \quad C_D = \frac{24}{Re_p}, \quad Re_p < 0.5 \quad (6)$$

159 The particle Reynolds number can be calculated from,

$$160 \quad Re_p = \rho_g \frac{d_p |u_r|}{\mu_g} \quad (7)$$

161 where,  $u_r$  is the relative velocity. The particle Re for 120nm particle for this study is 0.0168.

162 Amplitude for Brownian force is applied as

$$163 \quad F_{Brownian} = \zeta \sqrt{\frac{\pi S_0}{\Delta t}} \quad (8)$$

164 where  $\zeta$  is the unit variance for independent Gaussian random number, time-step integration of  
165 the particle  $\Delta t$ . The spectral intensity ( $S_0$ ) is defined as

$$S_o = \frac{216\mu k_B T}{\pi^2 \rho_p d_p^5 \left(\frac{\rho_p}{\rho_g}\right)^2 C_c} \quad (9)$$

167  $T$  is the fluid in absolute temperature,  $k_B$  is the Boltzmann constant,  $\rho_g$  is the gas density.

168 SIMPLE coupling scheme, and second order pressure discretisation technique are employed.

169 Second order upwind technique is utilised for energy and momentum equations. Hybrid

170 initialisation and pressure-based solver are used. The present model has considered the particle

171 with a density of 1.0 g/cm<sup>3</sup>; all particles were initiated from the one inlet area that was the

172 trachea. Steady injection method is used. In reality, SARS-CoV-2 particles are spherical in

173 shape like other viruses ([https://www.nih.gov/news-events/nih-research-matters/novel-](https://www.nih.gov/news-events/nih-research-matters/novel-coronavirus-structure-reveals-targets-vaccines-treatments)

174 [coronavirus-structure-reveals-targets-vaccines-treatments](https://www.nih.gov/news-events/nih-research-matters/novel-coronavirus-structure-reveals-targets-vaccines-treatments)). Goldsmith et al. (2020) reports that

175 SARS CoV-2 viruses are spherical in shape, and the structure is surrounded by dark dots, which

176 might have been interpreted as spikes on coronavirus. The transmission electron microscope

177 image also showed the spherical shape of the SARS CoV-2 virus. SARS CoV-2 aerosols are

178 injected beginning from the inlet surface in the normal direction. The mass flow rate at the inlet

179 is 0.5003 kg/s. The particles are injected from the inlet surface area, and each face of the surface

180 inject a single particle. The particle distribution at the inlet surface is uniform, and all particles

181 are injected at once. A total 14800 particles were injected. The outlet boundary condition is

182 used as pressure outlet, and open pressure condition is used at the terminal outlets.

183 The SARS CoV-2 aerosols were considered as secondary phase, and the air was the continuous

184 phase. The interaction between discrete phase and continuous phase is considered. The

185 maximum number of steps of tracking parameters is  $5 \times 10^8$  and the step length factor is 5.

186 Crowe et al. (2011) calculated the particle momentum response and collision time ratio, which

187 eventually indicated whether the air was dilute or dense.

188 The momentum response time of the particle can be explained as;



189 
$$\tau_V = \frac{\rho_p d_p^2}{18\mu} \quad (10)$$

190 where particle density is  $\rho_p$ , particle diameter is  $d_p$ , and  $\mu$  is the viscosity of air.

191 The time between the collisions can be defined as;

192 
$$\tau_c = \frac{1}{n\pi d_p^2 v_r} \quad (11)$$

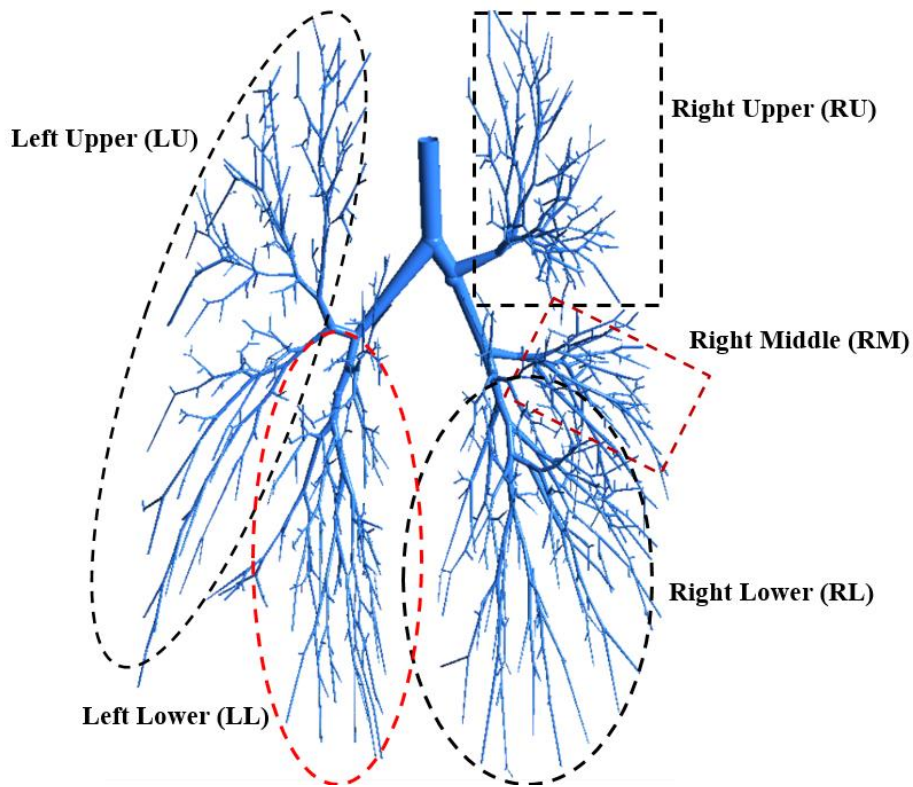
193 where  $n$  is the particle number,  $v_r$  is the particle relative velocity. If the ratio value is less than  
194 one, the fluid is dilute and one-way coupling can be considered. The value obtained for the  
195 ratio in the present study was 0.00041, which meant that this study is a one-way.

196 This study used ‘trap’ boundary condition for the wall. The physical meaning of the ‘trap’  
197 condition’ is particle will stick on the wall once it touches the wall. Once the particle touches  
198 the airway wall, the trajectory calculations will be terminated, and the fate of the particle will  
199 be recorded as ‘trapped’. The deposition fraction is defined as;

200 
$$DF = \frac{\text{Number of deposited particles in the wall}}{\text{Number of virus particles entering the inlet}}$$

## 201 **Grid Refinement and Model Validation**

202 A 17-generation airway model (Gemci et al. 2008; Schmidt et al. 2004) is employed for the  
203 SARS CoV-2 aerosol transport and deposition to the lower part of the lung. Figure 1 shows the  
204 airway model with five different lobes. The study performed a proper grid refinement, the  
205 details of the mesh at different section of the airway as well as the details of the grid refinement  
206 can be found in author's previous study (Islam et al. 2018).



207

208

Figure 1: Highly asymmetric 5-lobes 17-generation human lung model

209

210

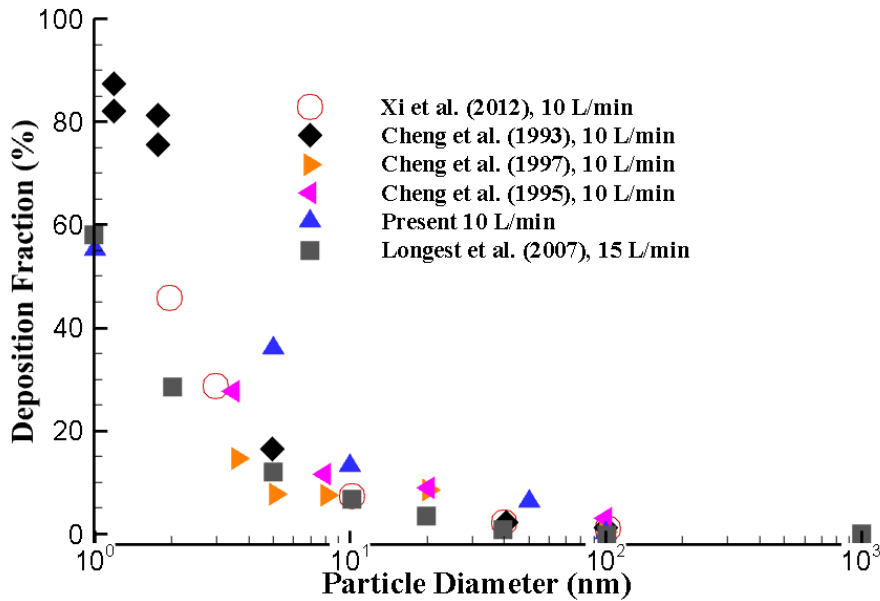
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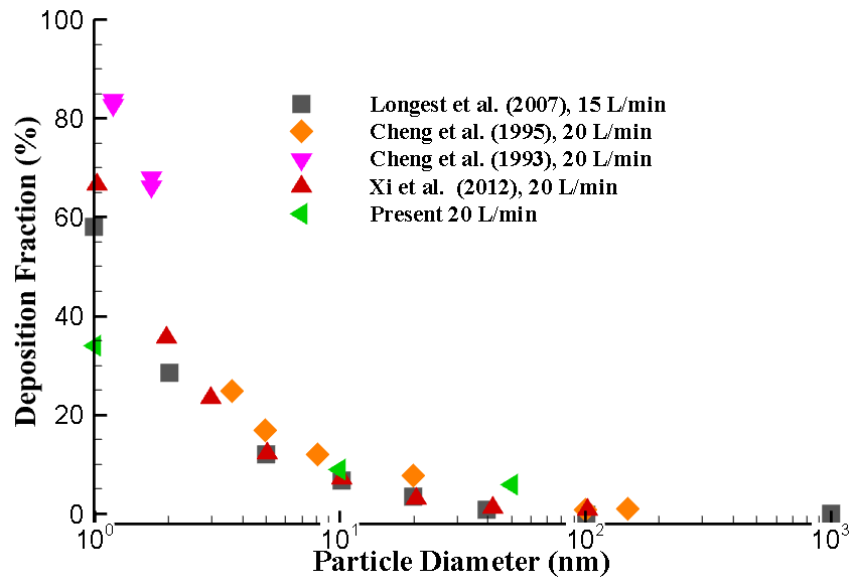
The numerical approach is validated with the available experimental and computations measurement in literature. A range of ultrafine particles are used to validate the deposition fraction (DF) at the upper airways. The ultrafine particle transport and DF is calculated for different airflow rates. Figure 2 illustrates the DF of the present calculation at 10 L/min inlet conditions with available literature data. The overall DF of the present study shows a close match with the available experimental and numerical measurements.



215

216 Figure 2: Numerical results validation with available literature at 10 L/min inlet airflow .

217 The DF of the present approach at higher flow rate is also comparing with the existing literature  
 218 (Cheng et al. 1996; Cheng et al. 1995; Cheng et al. 1993; Longest, Xi and Technology 2007;  
 219 Xi et al. 2012). Figure 3 demonstrates the comparison of DF at 20 L/min flow rate at the upper  
 220 airways. The DF for the smaller diameter particle is found to be higher when comparing to the  
 221 larger diameter nano-particle, which also support the hypothesis of the Brownian motion. The  
 222 DF of this present calculation indicates an good agreement with the available results for larger  
 223 nano-size particles .



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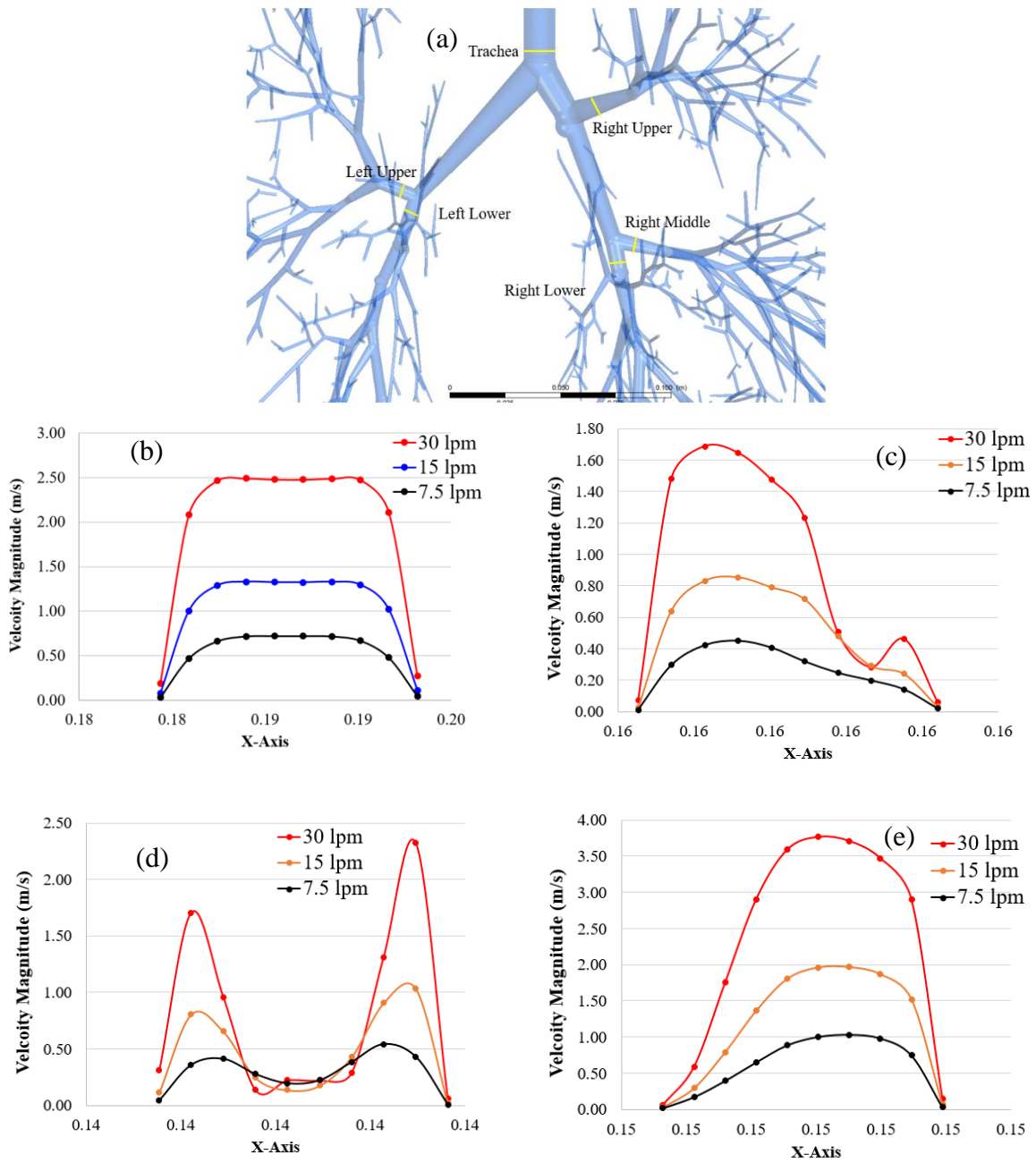
225 Figure 3: Numerical results validation with available literature at 20 L/min inlet flow (Cheng  
 226 et al. 1995; Cheng et al. 1993; Longest, Xi and Technology 2007; Xi et al. 2012).

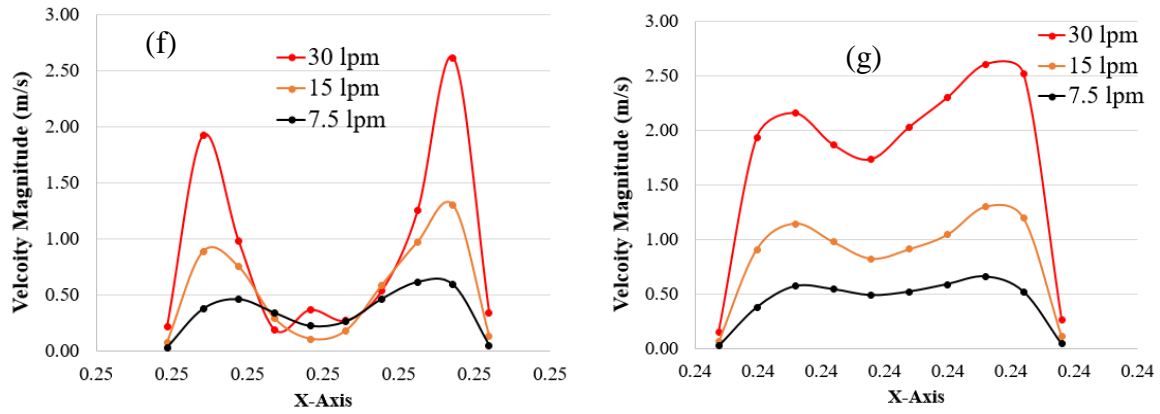
227 **Results and Discussion**

228 The airflow and SARS CoV-2 aerosol transport to the lower part of lung are simulated for  
 229 different inlet conditions. A highly asymmetric 17-generation bifurcating model is utilised to  
 230 analyse the SARS CoV-2 aerosol transport and deposition to the lower airways. The overall  
 231 investigation is performed for three different airflow rates, which consist of 7.5 L/min, 15  
 232 L/min, and 30 L/min.

233 The velocity profiles are plotted at various positions of the upper airways, and the lung at the  
 234 left side and right side for different breathing conditions. SARS CoV-2 aerosol usually follows  
 235 the air streamline inside the respiratory tube. An accurate understanding of the upper and lower  
 236 airways flow pattern is important to analyse the SARS CoV-2 transport and lung deposition.  
 237 Figure 4 presents the velocity profiles for three different flow rates at selected cross-sections  
 238 in different areas for the 17-generation lung model, from the trachea to terminal part of the  
 239 lung. Figure 4a presents the velocity profiles at the trachea. Figure 4b-d presents the velocity

240 profiles at three lobes (upper, lower and middle) of the right lung. Figure 4e-f presents the  
 241 velocity profiles at two lobes (lower and upper) of left lung.

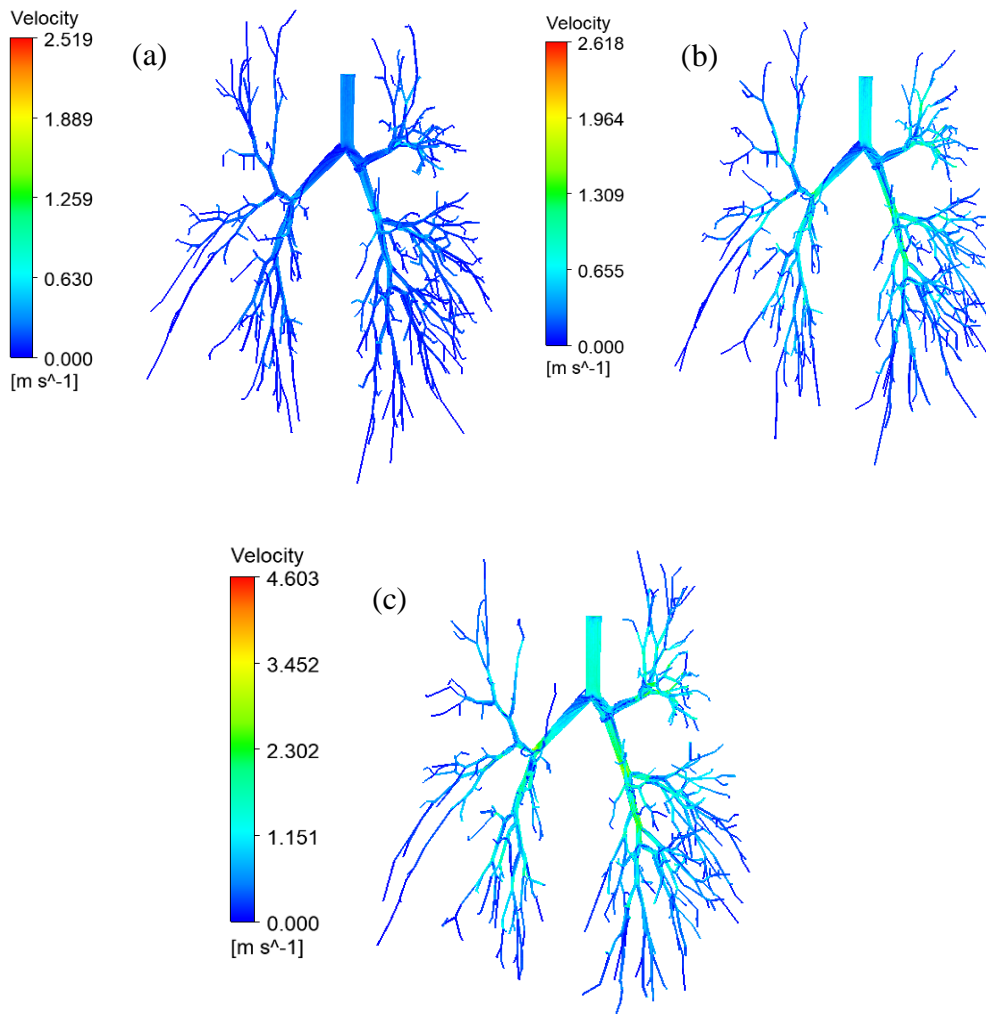




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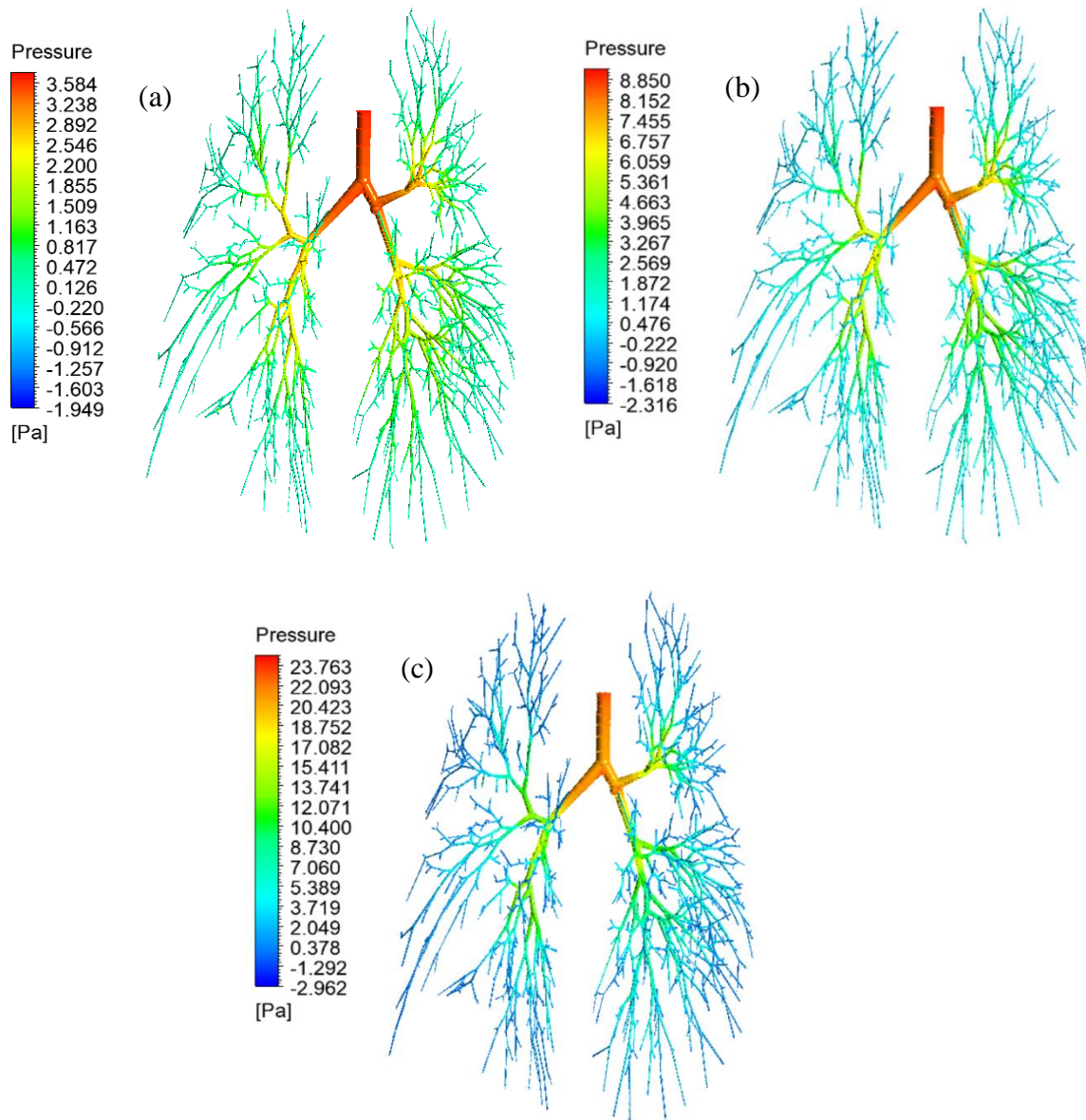
243 Figure 4: Air velocity profiles at randomly chosen location of the airway, (a) random location  
 244 definition, (b) trachea, (c) right upper (RU) lobe, (d) right middle (RM) lobe, (e) right lower  
 245 (RL) lobe, (f) left upper (LU) lobe and (g) left lower (LL) lobe.

246 Figure 4(a) shows the randomly selected locations at trachea and various lobes of the airway  
 247 model. The velocity profiles for all three air flow rates at the tracheal area show a fully  
 248 developed behaviour (Figure 4b) and the velocity magnitude is maximum at the centre of the  
 249 airways for all cases. However, the velocity field for all lobes tends to be locally transitional,  
 250 especially at RM lobe (Figure 4d) and LU lobe (Figure 4f), which have similar air velocity  
 251 magnitudes and nearly reach the 0.1 m/s at the middle point of the cross-section for all three  
 252 air flow rates. The velocity magnitude is higher close to the airway wall for RM and LU lobes,  
 253 which potentially increases the SARS CoV-2 aerosol deposition at the airways of the RM and  
 254 LU lobes. At the RU lobe, the velocity profile for 30 L/min shows a different trend than other  
 255 flow rates (Figure 4c). AT 30 L/min condition, the flow becomes locally unstable. The velocity  
 256 magnitude at the RL lobe (Figure 4e) is found maximum, whereas the RU lobe (Figure 4c) is  
 257 found to have the lowest velocity magnitude.



258 Figure 5: Velocity streamline at different airflow rates, (a) 7.5 L/min, (b) 15 L/min and (c) 30  
 259 L/min.

260 Figure 5 presents the velocity streamlines throughout the bifurcating model for various airflow  
 261 rates. Figure 5a shows the air velocity streamlines at 7.5 L/min, whereas Figure 5b and 5c  
 262 presents the air velocity streamline at 15 L/min and 30 L/min, respectively. The overall velocity  
 263 streamline shows a higher velocity magnitude at the upper area of the bifurcating model. The  
 264 velocity streamlines figure for low inlet velocity conditions (Figure 5a) indicates relatively  
 265 low-velocity magnitude to the terminal bronchioles than the high-velocity conditions (figure  
 266 5c). For low inlet flow condition (Figure 5a), the highest velocity magnitude is reported at the  
 267 upper bronchioles for all three right lobes. On the contrary, 15 L/min (Figure 5b) and 30 L/min  
 268 (Figure 5c) have the highest air velocity at the initial area for all five main lobes.



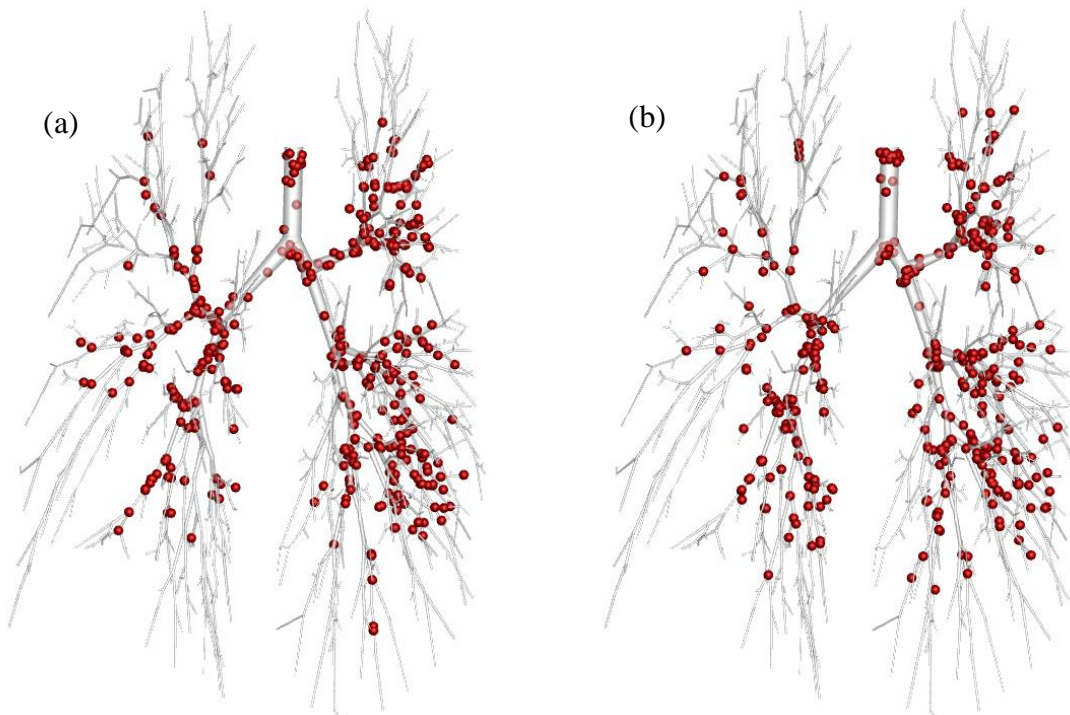
269 Figure 6: Pressure contour throughout the lung airways, (a) 7.5 L/min, (b) 15 L/min and (c)  
 270 30 L/min

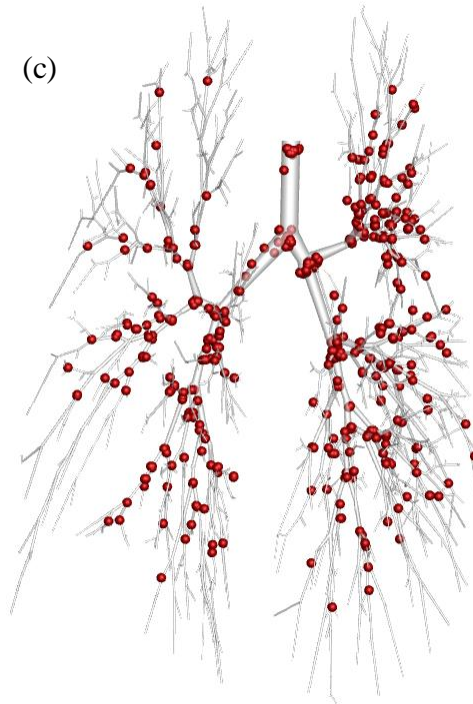
271 The precise knowledge of the airway pressure and pressure-drop to the terminal airways is  
 272 important for airway health risk analysis. Figure 6 presents the pressure contours for all three  
 273 different airflow rates that include 7.5 L/min (Figure 6a), 15 L/min (Figure 6b), and 30 L/min  
 274 (Figure 6c). Figure 6 reports that the pressure generally decreases from the initial area (trachea)  
 275 to the lower generation (17<sup>th</sup> generation). The pressure at the tracheal wall and the upper  
 276 airways is found maximum for all cases, and a significant pressure drop is obviously found in  
 277 the lower airways. The maximum pressure of 23.763 Pa is found at the highest airflow rate at



278 30 L/min (Figure 6c), while the maximum pressure of 3.584 Pa is found at the lowest airflow  
279 rate at 7.5 L/min (Figure 6a). Figure 6a shows, at low inlet condition (7.5 L/min), the pressure  
280 drop from the upper airways to the lower airways is insignificant. At high inlet velocity  
281 condition (30 L/min), the pressure drop is found significant from the upper airways to the lower  
282 airways. At high flow conditions, the terminal airways velocity magnitude is found relatively  
283 higher than the low inlet condition, which is reported in figure 5c. The higher velocity  
284 magnitude at the terminal airways eventually generates low pressure at the lower airways.

285 The SARS CoV-2 Aerosol deposition scenario is presented in Figure 7 under various airflow  
286 rate conditions. Figure 7a illustrates the deposition for airflow rate at 7.5 L/min, while Figure  
287 7b, c shows the deposition for the airflow rate at 15 L/min and 30 L/min, respectively.

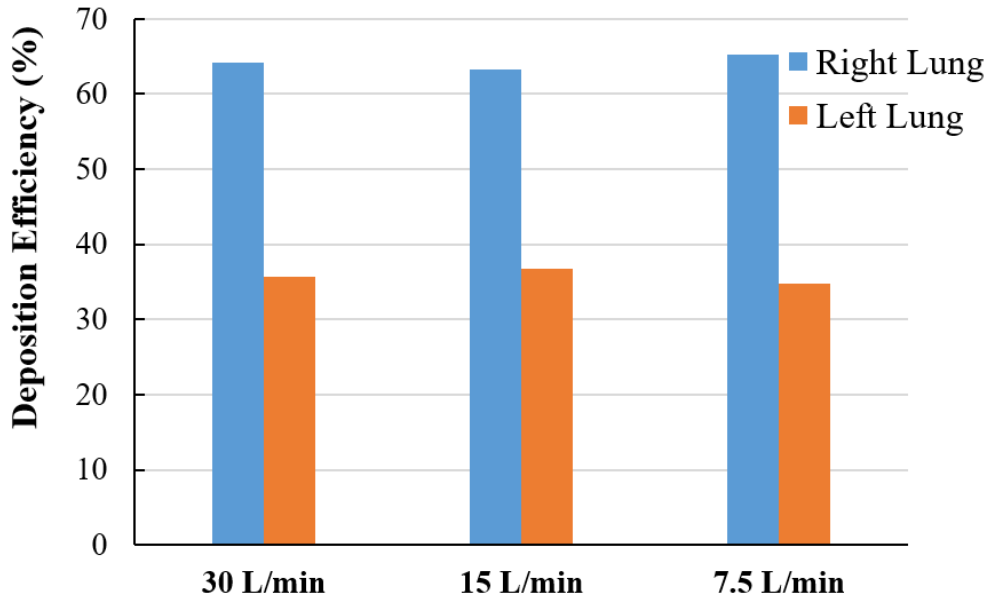




288 Figure 7: SARS CoV-2 Aerosol deposition (120nm) at different physical conditions, (a) 7.5  
289 L/min, (b) 15 L/min and (c) 30 L/min flow rate. (Sphere size is increased during post-  
290 processing for visualisation purpose)

291 Figure 7 shows that the SARS CoV-2 Aerosols are more trapped in the tracheal area and the  
292 bifurcation of all generations in the lung lobes for both lung sides. Figure 7 shows the higher  
293 SARS CoV-2 aerosol which is trapped at the tracheal inlet, and upper airways at 7.5 L/min  
294 (Figure 7a) and 15 L/min (Figure 7b) compared to the airflow rate at 30 L/min (Figure 7c).  
295 Brownian motion effect is dominant for smaller aerosol like SARS CoV-2 transport and  
296 deposition. SARS CoV-2 aerosol can spontaneously transport through the airways at a low  
297 flow rate, and the random movement of the SARS CoV-2 aerosol increases the overall  
298 deposition rate at the upper airways. Simultaneously, the Brownian motion effect becomes less  
299 effective with the increase of the velocity magnitude. The SARS CoV-2 aerosol deposition at  
300 the right side is higher than the left side for all airflow rates. At 15 L/min (Figure 7b) and 7.5  
301 L/min (Figure 7a) shows a cluster of SARS CoV-2 aerosol deposition at the three right lobes,

302 whereas the high airflow rate at 30 L/min (Figure 7c) shows a cluster SARS CoV-2 aerosol  
303 deposition at the upper area of the right side only.

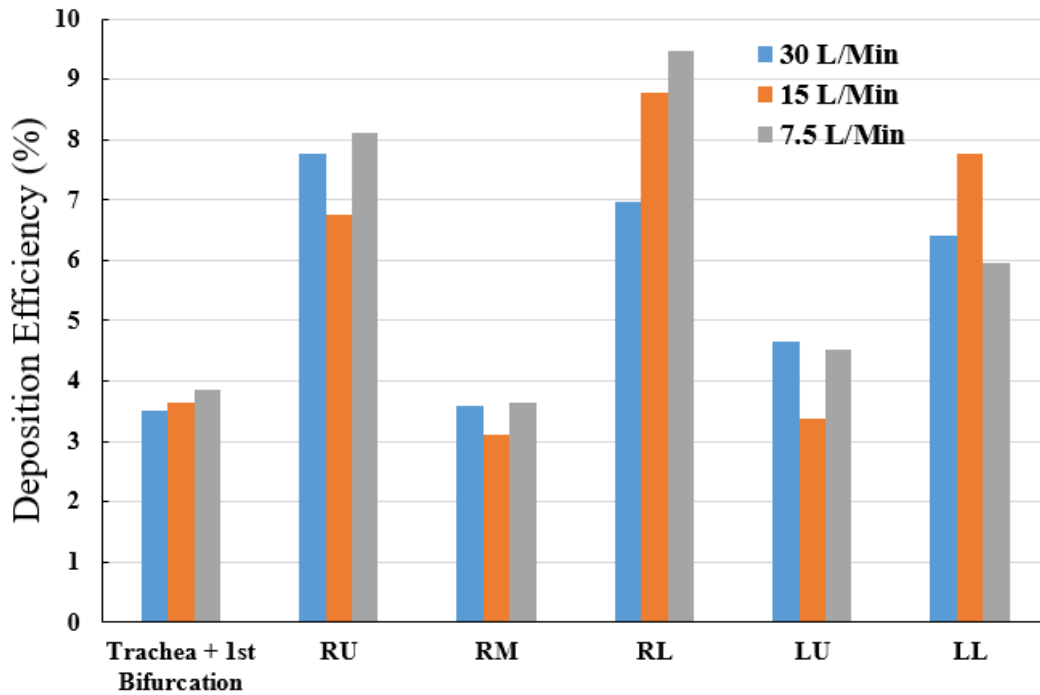


304

305 Figure 8: DE comparison at right and left lung for different airflow rate conditions.

306 Comparing SARS CoV-2 aerosol deposition efficiency (DE) between the right and the left lung  
307 is calculated and figure 8 reports the DE under the various airflow rate conditions. Figure 8  
308 shows that the DE of the SARS CoV-2 aerosol at the left side is found to be lower when  
309 comparing to the right side for all airflow rates. At 7.5 L/min, the highest DE 65.22% is  
310 reported at the right lung, and the highest DE at the right lung causes the lowest DE at the left  
311 lung. At 30 L/min inlet case, the DE of the SARS CoV-2 aerosol in the lung at the right side is  
312 64.25% and the left side is 35.75%. The anatomical structure of the right lung and left lung are  
313 different, and right lung airway diameter is higher than the left lung. A number of studies  
314 analysed the total flow distribution (%) in the right lung and left lung and found higher flow  
315 distribution to the right lung than the left lung. Cohen, Sussman and Lippmann (1990) found  
316 60% of the total flow goes through the right lung, Horsfield et al. (1971) reports 54.6% of the  
317 total flow goes through the right lung, and Islam et al. (2018) reports 54.93% total flow goes

318 through the right lung. Higher flow distribution to the right lung indicates a higher amount of  
 319 particle will go through the right bronchioles, which increases the deposition efficiency at the  
 320 right lung. The SARS CoV-2 aerosol follows the air pathlines and more SARS CoV-2 aerosol  
 321 enters into the right bifurcations due to high flow distribution to the right lung. The higher  
 322 SARS CoV-2 aerosol in the right bifurcations increase the overall DE at the right lung.



323

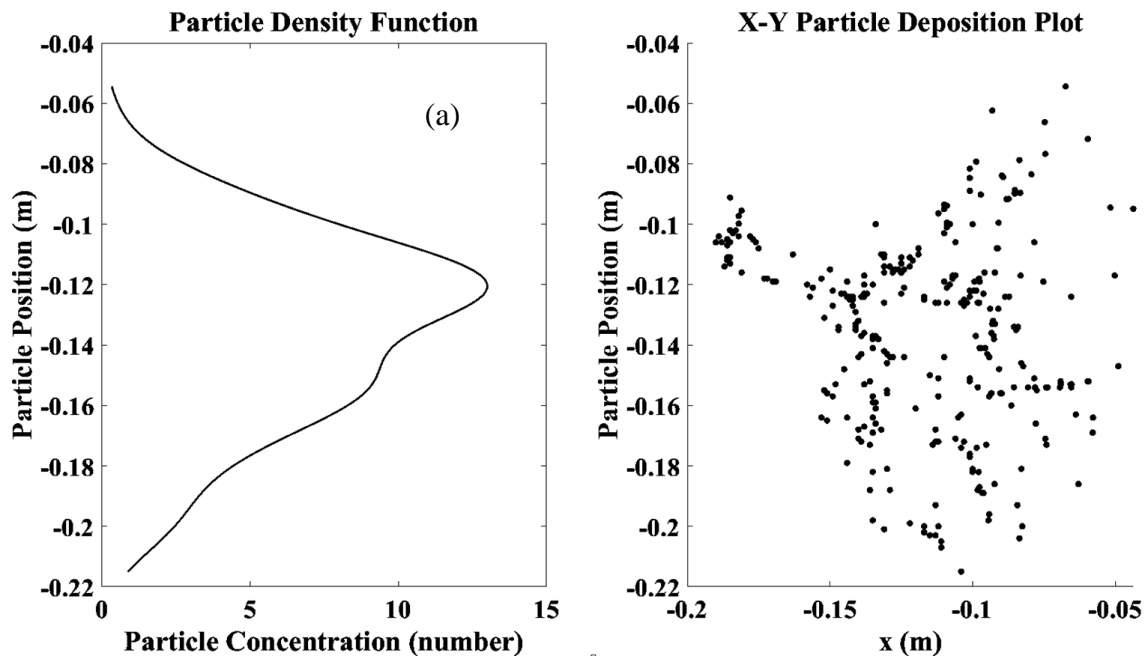
324 Figure 9: Local deposition of SARS CoV-2 Aerosol at different inlet conditions. LU, Left  
 325 upper lobe; LL, left lower lobe; RU, right upper lobe; RM, right middle lobe; and RL, right  
 326 lower lobe.

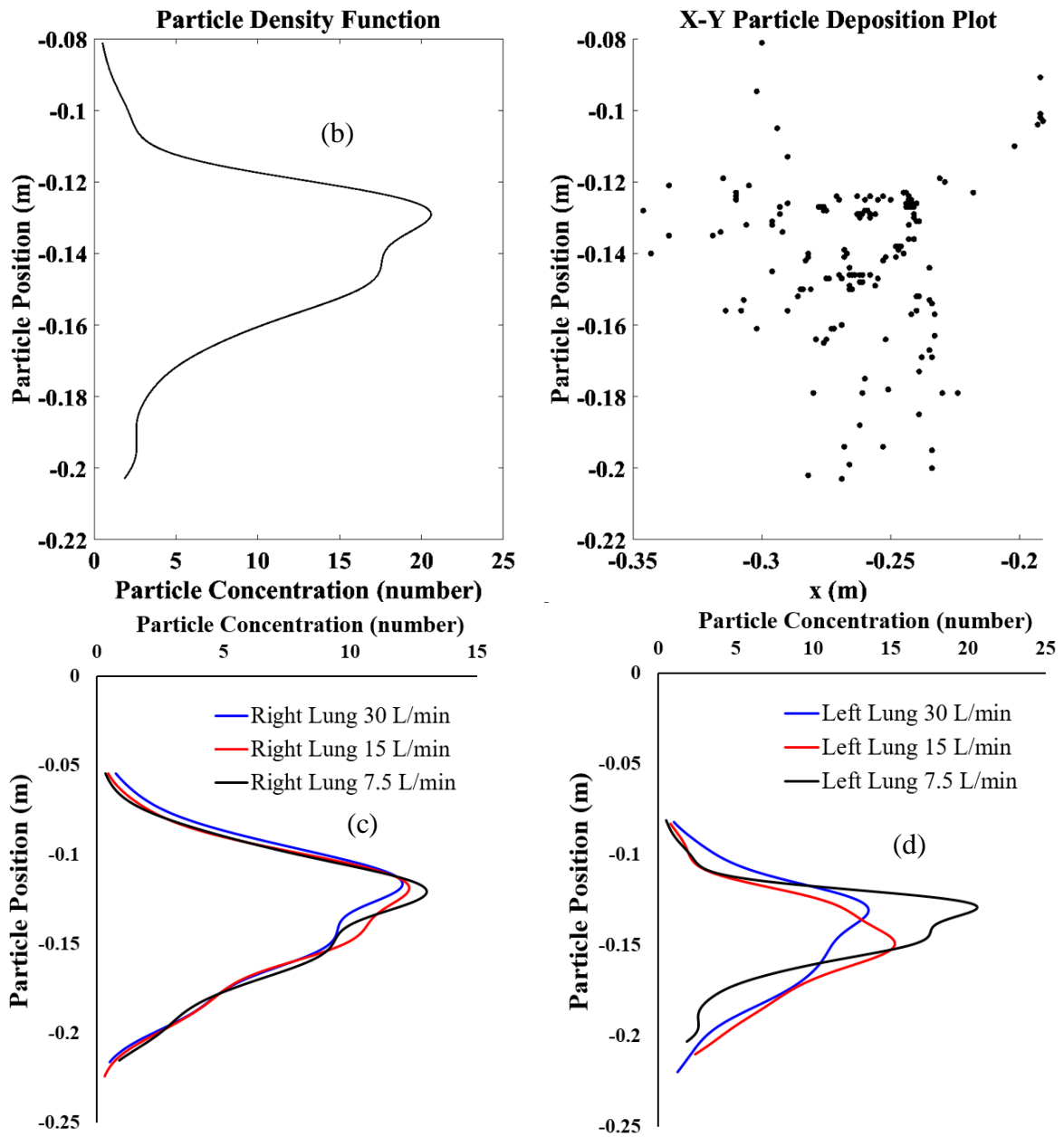
327 Figure 9 presents the SARS CoV-2 aerosol deposition at local areas involving the trachea & 1<sup>st</sup>  
 328 bifurcation, the three lung lobes of the right side, and the two lung lobes of the left side. The  
 329 local DE of the SARS CoV-2 aerosol is calculated for different airflow rates. The local DE of  
 330 the SARS CoV-2 aerosol illustrates higher deposition at the trachea and first bifurcation region  
 331 at low inlet condition (7.5 L/min). At 7.5 L/min condition, 3.85% of the total SARS CoV-2  
 332 aerosols are deposited at trachea and first bifurcation area, whereas 3.51% SARS CoV-2

333 aerosols are deposited for 30 L/min case. The local SARS CoV-2 aerosol DE at the bronchioles  
334 of the RL lobe is found higher than other lobes. At 7.5 L/min inlet condition, 9.46% of the total  
335 SARS CoV-2 aerosols are deposited at the RL lobe, whereas 6.96% for 30 L/min airflow rate.  
336 At 15 L/min inlet case, the SARS CoV-2 aerosol deposition DE is found maximum at the LL  
337 lower lobe which is 7.77%. The overall SARS CoV-2 aerosol DE curve reports that the DE for  
338 30 L/min inlet case is lower at all lobes, including the trachea and first bifurcation of the 17  
339 generation model. For the comparison between the two lung sides, the overall DE at the right  
340 side is found to be higher than the left side. The DE at RM lobe and LU lobe is lower than other  
341 areas. To be concluded, the SARS CoV-2 aerosols are mostly trapped at RL lobe and RU lobe  
342 and rarely trapped at RM lobe and LU lobe for 7.5 L/min and 30 L/min. In contrast, for 15  
343 L/min, the majority of the SARS CoV-2 aerosol deposition generally locates at RL lobe and  
344 LL lobe, while the minority of this aerosol deposition is in the RM lobe and LU lobe. The first-  
345 ever lobe-specific SARS CoV-2, aerosol DE analysis for the 17-generation model, would  
346 improve the knowledge of the SARS CoV-2 transport to the lower part of the lung airways of  
347 a large-scale model. A Recent study have investigated the (Kwee and Kwee 2020b) CT-images  
348 of SARS CoV-2 positive patient from a RT-PCR test. The CT-Scan imaged reports the SARS  
349 CoV-2 presence at the RU lobe (figure 10a), RM and LL lobe (figure 10b), RU lobe (figure  
350 10c) and both lower lobes (figure 10d), which necessarily indicate the significance of the  
351 present study. A comprehensive lobe-specific analysis is presented in figure 9, which would  
352 potentially improve the knowledge of the field.

353 Figure 10 demonstrates the SARS CoV-2 aerosol deposition concentration in the right lung and  
354 left lung in the different airflow rate conditions. This aerosol deposition concentration is plotted  
355 based on the SARS CoV-2 aerosols x, y, z in the airway wall. Figure 10a presents the SARS  
356 CoV-2 aerosol concentration for 7.5 L/min flow condition at the right lung, while figure 10b  
357 shows the left lung concentration. Both figure's right panel shows the deposited SARS CoV-2

358 aerosol and the left panel shows the deposition concentration curve. The concentration curve  
359 on the left panel demonstrates the deposition hot spot at the bifurcating airways, that is  
360 presented at the right panel of the figure. The concentration curve shows the SARS CoV-2  
361 aerosol deposition concentration is found to be higher at the upper as well as middle  
362 bifurcations of the 17-generation model. A comprehensive SARS CoV-2 aerosol deposition  
363 concentration for all cases are presented in figures 10 (c, d). At right lung, the SARS CoV-2  
364 aerosol concentration shows a similar trend for all cases whereas a different trend is observed  
365 at the left lung. The asymmetric brunching pattern of the left and the right lung influences the  
366 overall SARS CoV-2 aerosol transport to the lower airways. This analysis would provide an  
367 understanding of the SARS CoV-2 aerosol deposition hot spot at the lower airways of a large  
368 model.





369

370 Figure 10: SARS CoV-2 aerosol deposition concentration from trachea to the terminal  
 371 airways for various airflow rate conditions, (a) 7.5 L/min at right lung, (b) 7.5 L/min at left  
 372 lung, (c) right lung, and (d) left lung.

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375

376 **Conclusions**

377 In this paper, the SARS CoV-2 aerosol transport to the lower part of the lung airways of a 17-  
378 generations lung airway model is investigated numerically for the first time. The SARS  
379 CoV-2 aerosol transportation to the lower airways is investigated for different inlet  
380 conditions. The key findings of the study are listed as following;

- 381 • The cluster of SARS CoV-2 aerosols is found at the right lung, which is more than one  
382 time of the left lung for all airflow rates. A total of 35.55%, 33.45% and 32.90% of SARS  
383 CoV-2 injected aerosols are deposited to the airway wall for 7.5 L/min, 15 L/min and 30  
384 L/min, respectively. The remaining aerosols escape and transport to lower generations and  
385 alveolar region.
- 386 • The highest deposition efficiency is located at the RL lobe with the low airflow rate of  
387 7.5 L/min, whereas the lowest DE is found at the RM and LU lobes with the airflow rate of  
388 15 L/min.
- 389 • The majority of SARS CoV-2 aerosols is trapped at RL and RU lobes, and the minority  
390 is trapped at RM and LU lobes for 7.5 L/min and 30 L/min airflow rates. For 15 L/min, the  
391 minority of aerosol deposition is in the RM and LU lobes which are similar to other airflow  
392 rates but the majority of this aerosol deposition is located at RL and LL lobes instead.
- 393 • The SARS CoV-2 deposition concentration curves show a similar trend for the right  
394 lung, while the left lung is different. The deposition hot spot (DHS) of the right lung is  
395 found at the first bifurcation of the RU lobe. For the left lung, the DHS is found at LU and  
396 LL lobes for 7.5 L/min and 30 L/min, while the 15 L/min has the DHS point at LU lobe  
397 only.

398 The numerical study demonstrates the SARS CoV-2 aerosol transport, and deposition  
399 concentration at different lobes of the large airway model. The numerical study investigated



400 the SARS CoV-2 aerosol transport to the lower area of the lung airways of a 17-generation  
401 model for the first time and a comprehensive lobe-specific analysis is performed, which would  
402 improve the SARS CoV-2 aerosol transport knowledge to the lower airways and help the health  
403 risk assessment of the covid patients. The numerical study also analysed the deposition hotspot  
404 of the SARS CoV-2 aerosol to the right and the left lung. The present study along with more  
405 patient-specific study would improve the knowledge of the field. The future study will  
406 investigate the age and patient-specific whole lung model for better understanding of the SARS  
407 CoV-2 aerosol to the lower airways.

#### 408 **Assumptions of the Study**

409 In reality, the aerosol emitted during exhalation exhibits a wide size distribution. The smaller  
410 droplet could evaporate during transportation and become more smaller. During exhalation, the  
411 aerosol could contain a single SARS CoV-2 virus or more than one SARS CoV-2 virus. If the  
412 aerosol contains more than one SARS CoV-2 viruses, then the size and shape of the virus-laden  
413 particle could be different. This study assumed a single isolated virus and did not consider the  
414 aggregation of the SARS CoV-2 viruses. The future study will perform a comprehensive  
415 analysis on virus-laden particles, and the aggregation of the viruses on aged people lung as the  
416 virus is found deadly for older people. The study assumed that virus particles have no electrical  
417 charges for the intermolecular forces and van der Waals interactions are neglected as the study  
418 did not investigate the particle and lung surfactant interaction.

419

#### 420 **Conflicts of the Interest**

421 No conflicts of interested are associated with this study.

#### 422 **Acknowledgement:**

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425 **Data Availability:**

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

428  
429 **References:**

- 430 Anderson, P.J., 2005, History of Aerosol Therapy: Liquid Nebulization to MDIs to DPIs,  
431 Respiratory Care, vol. 50, no. 9, pp. 1139-1150.
- 432 Wells, W.F., 1955, Airborne contagion and air hygiene. Cambridge, MA: Harvard  
433 University Press.
- 434 Mason, R.J., 2020. Pathogenesis of COVID-19 from a cell biology perspective. Eur Respir  
435 Journal. vol. 55, pp. 1-3. doi: 10.1183/13993003.00607-2020
- 436 Atkinson, J.; Chartier, Y.; Pessoa-Silva, C. L.; Jensen, P.; Li, Y., Seto, W. (2009). Annex  
437 C: Respiratory droplets. *Natural Ventilation for Infection Control in Health-Care*  
438 *Settings*. World Health Organization. ISBN: 978-92-4-154785-7
- 439 Weibel, E.R., 1963, Morphometry of the Human Lung. Heidelberg: Springer Verlag.
- 440 Phillips, C.G., Kaye, S.R., 1995, Diameter-based analysis of the branching geometry of  
441 four mammalian bronchial trees. *Respiration Physiology*. vol. 102, pp. 303–316.
- 442 Kitaoka, H., Takaki, R., Suki, B. 1999, A three-dimensional model of the human airway  
443 tree. *Journal of Applied Physiology*. vol. 87, pp. 2207–2217.
- 444 Martonen, T.B., Yang, Y., Hwang, D., Fleming, J.S. 1995. Computer simulations of  
445 human lung structures for medical applications. *Computer Biology and Medicine*. vol.  
446 25, pp. 431–446.
- 447 Lizal, F., Elcner, J., Hopke, P.K., Jedelsky, J., Jicha, M., 2012. Development of a realistic  
448 human airway model. Proc Inst Mech Eng H. vol. 226, no. 3, pp. 197-207. doi:  
449 10.1177/0954411911430188.
- 450 Srivastav, V.K., Paul, A.R., Jain, A., 2013. Effects of cartilaginous rings on airflow and  
451 particle transport through simplified and realistic models of human upper respiratory  
452 tracts, Acta Mechanica Sinica, vol. 29, no. 6, pp. 883–892. doi: 10.1007/s10409-013-  
453 0086-2
- 454 Burrowes, K.S., Swan, A.J., Warren, N.J., Tawhai, M.H., 2008. Towards a virtual lung:  
455 multiscale, multiphysics modelling of the pulmonary system. *Philos Trans A Math Phys*  
456 *Eng Sci*. vol. 28, no. 366(1879), pp. 3247-63. doi: 10.1098/rsta.2008.0073.

- 457 Islam, M.S., Paul, G., Ong, H.X., Young, P.M., Gu, Y.T., Saha, S.C., 2020. A review of  
458 respiratory anatomical development, air flow characterisation and particle deposition,  
459 International Journal of Environmental Research and Public Health. vol. 17, no. 380,  
460 pp. 1-28.
- 461 Xi, J., Longest, P.W. 2007, Transport and deposition of micro-aerosols in realistic and  
462 simplified models of the oral airway. *Ann Biomed Eng.* vol. 35, no. 4, pp. 560-581. doi:  
463 10.1007/s10439-006-9245-y
- 464 Islam, M.S., Saha, S.C., Sauret, E., Gemci, T., Gu, Y.T., 2017. Pulmonary aerosol transport  
465 and deposition analysis in upper 17 generations of the human respiratory tract, *Journal*  
466 *of Aerosol Science*, vol. 108, pp. 29-43.
- 467 Lizal, F., Elcner, J., Jedelsky, J., Maly, M., Jicha, M., Farkas, Á., Belka, M., Rehak, Z.,  
468 Adam, J., Brinek, A., Laznovsky, J., Zikmund, T., Kaiser, J., 2020. The effect of oral  
469 and nasal breathing on the deposition of inhaled particles in upper and tracheobronchial  
470 airways. *J Aerosol Sci.* vol. 150:105649. doi: 10.1016/j.jaerosci.2020.10564
- 471 Kleinstreuer, C., Zhang, Z., Donohue, J.F., 2008. Targeted drug-aerosol delivery in the  
472 human respiratory system. *Annual Revision Biomedical Engineering*, vol. 10, pp. 195–  
473 220.
- 474 Martonen, T., Fleming, J., Schroeter, J., Conway, J., Hwang, D. 2003. In silico modeling  
475 of asthma. *Advanced Drug Delivery Reviews.* vol. 55, pp. 829-49, 2003.
- 476 Srivastav, V.K., Kumar, A., Shukla, S.K., Paul, A.R., Bhatt, A.D., Jain, A., 2014. Airflow  
477 and aerosol-drug delivery in a CT scan based human respiratory tract with tumor using  
478 CFD, *Journal of Applied Fluid Mechanics.* vol. 7, no. 2, pp. 345-356.
- 479 Kumar, S., Lee, H.P., 2020. The perspective of fluid flow behavior of respiratory droplets  
480 and aerosols through the facemasks in context of SARS-CoV-2. *Phys. Fluids.* vol. 32,  
481 111301 (2020); doi: 10.1063/5.0029767
- 482 Prather, K. A., Wang, C. C., Schooley, R. T. 2020. Reducing transmission of SARSCoV-  
483 2, *Science* 368, 1422.
- 484 Vuorinen, V., Aarnio, M., Alava, M., Alopaeus, V., Atanasova, N., Auvinen, M.,  
485 Balasubramanian, N., Bordbar, H., Erästö, P., Grande, R., Hayward, N., Hellsten, A.,  
486 Hostikka, S., Hokkanen, J., Kaario, O., Karvinen, A., Kivistö, I., Korhonen, M.,  
487 Kosonen, R., Kuusela, J., Lestinen, S., Laurila, E., Nieminen, H.J., Peltonen, P., Pokki,  
488 J., Puisto, A., Råback, P., Salmenjoki, H., Sironen, T., Österberg, M., 2020. Modelling  
489 aerosol transport and virus exposure with numerical simulations in relation to SARS-  
490 CoV-2 transmission by inhalation indoors. *Saf. Sci.* vol. 130, 104866.
- 491 Xie, X., Li, Y., Sun, H., Liu, L., 2009, Exhaled droplets due to talking and coughing. *J. R.*  
492 *Soc., Interface.* vol. 6, S703.
- 493 Feng, Y., Marchal, T., Sperry, T., Yi, H., 2020. Influence of wind and relative humidity  
494 on the social distancing effectiveness to prevent COVID-19 airborne transmission: A  
495 numerical study," *J. Aerosol Sci.* vol. 147, 105585.
- 496 Chaudhuri, S., Basu, S., Kabi, P., Unni, V.R., Saha, A., 2020. Modeling the role of  
497 respiratory droplets in COVID-19 type pandemics," *Phys. Fluids.* vol. 32, 063309.

- 498 Bhardwaj, R., Agrawal, A., 2020a. Likelihood of survival of coronavirus in a respiratory  
499 droplet deposited on a solid surface. *Phys. Fluids*. vol. 32, 061704 (2020)  
500 <https://doi.org/10.1063/5.0012009>
- 501 Bhardwaj, R., Agrawal, A., 2020b. How coronavirus survives for days on surfaces. *Phys.*  
502 *Fluids*, vol. 32, 111706. doi: 10.1063/5.0033306
- 503 Balázs G. Madas, Péter Fűri, Árpád Farkas, Attila Nagy, Aladár Czitrovszky, Imre  
504 Balásházy, Gusztáv G. Schay, Alpár Horváth, 2020. Deposition distribution of the new  
505 coronavirus (SARS-CoV-2) in the human airways upon exposure to cough-generated  
506 aerosol. *medRxiv* 2020.05.13.2010005 doi: 10.1101/2020.05.13.20100057
- 507 Koblinger, L., Hofmann, W., 1990. Monte Carlo modeling of aerosol deposition in human  
508 lungs. Part I: Simulation of particle transport in a stochastic lung structure. *Journal of*  
509 *Aerosol Science*. vol. 21, pp. 661–674. doi:10.1016/0021-8502(90)90121-D
- 510 Fitzpatrick, M.F., McLean, H., Urton, A.M., Tan, O'Donnell, D., Driver, H.S., 2003.  
511 Effect of nasal or oral breathing route on upper airway resistance during sleep.  
512 *European Respiratory Journal*. vol. 22, pp. 827-832; DOI:  
513 10.1183/09031936.03.00047903
- 514 Fernández Tena A, Casan Clarà P., 2012. Deposition of Inhaled Particles in the  
515 Lungs. *Arch Bronconeumol*. 2012;48(7):240–246
- 516 Mortazavi, H., Mortazavy Beni, H., Aghaei, F., Sajadian, S.H., 2020. SARS-CoV-2  
517 droplet deposition path and its effects on the human upper airway in the oral inhalation.  
518 *Computer Methods and Programs in Biomedicine*.  
519 <https://doi.org/10.1016/j.cmpb.2020.105843>
- 520 Heyder, J.. 2004. Deposition of Inhaled Particles in the Human Respiratory Tract and  
521 Consequences for Regional Targeting in Respiratory Drug Delivery. *Proc Am Thorac*  
522 *Soc*. vol. 1. pp 315–320. DOI: 10.1513/pats.200409-046TA
- 523 Frederix, E.M.A., Kuczaj, A.K., Nordlund, M., Bělka, M., Lizal, F., Jedelský, J., Elcner,  
524 J., Jícha, M., Geurts, B.J., Simulation of size-dependent aerosol deposition in a realistic  
525 model of the upper human airways. *Journal of Aerosol Science* 115 (2018) 29–  
526 45 <https://doi.org/10.1016/j.jaerosci.2017.10.007>
- 527 Pozin, N. 2017. Multiscale lung ventilation modeling in health and disease. *Modeling and*  
528 *Simulation*. Université Pierre et Marie Curie - Paris VI, 2017. English. ffNNT:  
529 2017PA066581ff.
- 530 [https://www.nih.gov/news-events/nih-research-matters/novel-coronavirus-structure-](https://www.nih.gov/news-events/nih-research-matters/novel-coronavirus-structure-reveals-targets-vaccines-treatments)  
531 [reveals-targets-vaccines-treatments](https://www.nih.gov/news-events/nih-research-matters/novel-coronavirus-structure-reveals-targets-vaccines-treatments) (access on 5<sup>th</sup> May 2021)
- 532 Ciloglu, D., A numerical study of the aerosol behavior in intra-acinar region of a human  
533 lung. 2020. *Phys. Fluids* 32, 103305 (2020); <https://doi.org/10.1063/5.0024200>
- 534 Munir, B., Xu, Y., 2020. Effects of gravity and surface tension on steady microbubble  
535 propagation in asymmetric bifurcating airways. *Phys. Fluids* 32, 072105 (2020);  
536 <https://doi.org/10.1063/5.0012796>
- 537 Xi, J., Si, X.A., Nagarajan, R., 2020. Effects of mask-wearing on the inhalability and  
538 deposition of airborne SARS-CoV-2 aerosols in human upper airway. *Phys. Fluids* 32,  
539 123312 (2020); <https://doi.org/10.1063/5.0034580>

540 Mallik, A.K., Mukherjee, S., Panchagnula, M.V., 2020. An experimental study of  
541 respiratory aerosol transport in phantom lung bronchioles. *Phys. Fluids* 32, 111903  
542 (2020); <https://doi.org/10.1063/5.0029899>

543 Zhou, M., Zou, J., 2021. A dynamical overview of droplets in the transmission of  
544 respiratory infectious diseases. *Phys. Fluids* 33, 031301 (2021);  
545 <https://doi.org/10.1063/5.0039487>

546 Si, X.A., Talaat, M., Xi, J., 2021. SARS COV-2 virus-laden droplets coughed from deep  
547 lungs: Numerical quantification in a single-path whole respiratory tract geometry. *Phys.*  
548 *Fluids* 33, 023306 (2021); <https://doi.org/10.1063/5.0040914>

549 Chaudhuri, S., S. Basu, P. Kabi, V. R. Unni, A. J. P. o. F. Saha. 2020. Modeling the role of respiratory  
550 droplets in covid-19 type pandemics 32:063309. doi.

551 Cheng, K.-H., Y.-S. Cheng, H.-C. Yeh, R. A. Guilmette, S. Q. Simpson, Y.-H. Yang, D. L. J. A. S. Swift.  
552 1996. In vivo measurements of nasal airway dimensions and ultrafine aerosol deposition in  
553 the human nasal and oral airways. *J Journal of Aerosol Science* 27:785-801. doi.

554 Cheng, K.-H., Y.-S. Cheng, H.-C. Yeh, D. L. J. A. S. Swift, Technology. 1995. Deposition of ultrafine  
555 aerosols in the head airways during natural breathing and during simulated breath holding  
556 using replicate human upper airway casts. *J Aerosol Science* 23:465-474. doi.

557 Cheng, Y.-S., Y.-F. Su, H.-C. Yeh, D. L. J. A. S. Swift, Technology. 1993. Deposition of thoron progeny in  
558 human head airways. *J Aerosol Science* 18:359-375. doi.

559 Cohen, B. S., R. G. Sussman, M. Lippmann. 1990. Ultrafine particle deposition in a human  
560 tracheobronchial cast. *Aerosol Science and Technology* 12:1082-1091. doi.

561 Crowe, C. T., J. D. Schwarzkopf, M. Sommerfeld, Y. Tsuji. 2011. *Multiphase flows with droplets and*  
562 *particles*: CRC press.

563 Gemci, T., V. Ponyavin, Y. Chen, H. Chen, R. Collins. 2008. Computational model of airflow in upper  
564 17 generations of human respiratory tract. *Journal of Biomechanics* 41:2047-2054. doi.

565 Goldsmith, C. S., S. E. Miller, R. B. Martines, H. A. Bullock, S. R. Zaki. 2020. Electron microscopy of  
566 sars-cov-2: A challenging task. *J The Lancet* 395:e99. doi.

567 Haider, A. and O. J. P. t. Levenspiel. 1989. Drag coefficient and terminal velocity of spherical and  
568 nonspherical particles. *J Powder technology* 58:63-70. doi.

569 Horsfield, K., G. Dart, D. E. Olson, G. F. Filley, G. Cumming. 1971. Models of the human bronchial  
570 tree. *Journal of applied physiology* 31:207-217. doi.

571 [https://www.pptaglobal.org/media-and-information/ppta-statements/1055-2019-novel-](https://www.pptaglobal.org/media-and-information/ppta-statements/1055-2019-novel-coronavirus-2019-ncov-and-plasma-protein-therapies)  
572 [coronavirus-2019-ncov-and-plasma-protein-therapies](https://www.pptaglobal.org/media-and-information/ppta-statements/1055-2019-novel-coronavirus-2019-ncov-and-plasma-protein-therapies). doi.

573 Inthavong, K., J. Tu, G. Ahmadi. 2009. Computational modelling of gas-particle flows with different  
574 particle morphology in the human nasal cavity. *The Journal of Computational Multiphase*  
575 *Flows* 1:57-82. doi.

576 Islam, M. S., S. C. Saha, T. Gemci, I. A. Yang, E. Sauret, Y. T. Gu. 2018. Polydisperse microparticle  
577 transport and deposition to the terminal bronchioles in a heterogeneous vasculature tree.  
578 *Scientific Reports* 8:16387. doi: 10.1038/s41598-018-34804-x.

579 Kwee, T. C. and R. M. Kwee. 2020a. Chest ct in covid-19: What the radiologist needs to know. *J*  
580 *RadioGraphics* 40:1848-1865. doi.

581 Kwee, T. C. and R. M. Kwee. 2020b. Chest ct in covid-19: What the radiologist needs to know. *J*  
582 *RadioGraphics* 40:1848-1865. doi.

583 Longest, P. W., J. Xi, Technology. 2007. Effectiveness of direct lagrangian tracking models for  
584 simulating nanoparticle deposition in the upper airways. *J Aerosol Science* 41:380-397. doi.

585 Schmidt, A., S. Zidowitz, A. Kriete, T. Denhard, S. Krass, H.-O. Peitgen. 2004. A digital reference  
586 model of the human bronchial tree. *Computerized Medical Imaging and Graphics* 28:203-  
587 211. doi: <http://dx.doi.org/10.1016/j.compmedimag.2004.01.001>.

588 White, F. M. 2003. Fluid mechanics. 5th. *Boston: McGraw-Hill Book Company*. doi.

589 Xi, J., A. Berlinski, Y. Zhou, B. Greenberg, X. J. A. o. b. e. Ou. 2012. Breathing resistance and ultrafine  
590 particle deposition in nasal–laryngeal airways of a newborn, an infant, a child, and an adult. *J*  
591 *Annals of biomedical engineering* 40:2579-2595. doi.

592

593