1 Version 1

Xiamen University

3	Title: A mathematical model for estimating the age-specific transmissibility of a novel
4	coronavirus
5	Running title: Age-specific transmissibility of SARS-CoV-2
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- 36
- 37 Abstract

38 Background: A novel coronavirus named as "SARS-CoV-2" has spread widely in many countries since

39 December 2019, especially in China. This study aimed to quantify the age-specific transmissibility by

40 using a mathematical model.

41 **Methods:** An age-specific susceptible – exposed – symptomatic – asymptomatic – recovered – 42 seafood market (SEIARW) model was developed based on two suspected transmission routes (from 43 market to person and person to person). The susceptible people from Wuhan City were divided into 44 different age groups. We used the subscript *i* and *j* to represent age group 1 to 4 ($i \neq j$; 1: \leq 14 years; 2:

45 15-44 years; 3: 45-64 years; 4: \geq 65 years) and 1 to 5 ($i \neq j$; 1: \leq 5 years; 2: 6-14 years; 3: 15-24 years; 4: 46 25-59 years; $4 \ge 60$ years), respectively. Data of reported COVID-19 cases were collected from one 47 published literature from 26 November to 22 December, 2019 in Wuhan City, China. The age-specific 48 transmissibility of the virus was estimated accordingly secondary attack rate (SAR). 49 **Results:** The age-specific SEIARW model fitted with the reported data well by dividing the population into four age groups ($\chi^2 = 4.99 \times 10^{-6}$, P > 0.999), and five age groups ($\chi^2 = 4.85 \times 10^{-6}$, P > 0.999). 50 51 Based on the four-age-group SEIARW model, the highest transmissibility occurred from age group 2 to 52 3 (SAR₂₃ = 17.56 per 10 million persons), followed by from age group 3 to 2 (SAR₃₂ = 10.17 per 10 53 million persons). The lowest transmissibility occurred from age group 1 to 2 ($SAR_{12} = 0.002$ per 10 million persons). Based on the five-age-group SEIARW model, the highest transmissibility occurred 54 55 from age group 4 to 5 (SAR₄₅ = 12.40 per 10 million persons), followed by from age group 5 to 4 (SAR₅₄ 56 = 6.61 per 10 million persons). The lowest transmissibility occurred from age group 3 to 4 (SAR_{34} = 57 0.0002 per 10 million persons).

58 Conclusions: SARS-CoV-2 has high transmissibility among adults and elder people but low
 59 transmissibility among children and young people.

60 Key words: Transmissibility; Novel coronavirus; Mathematical model; Age-specific dynamics

62 Introduction

63 In December 2019, a series of cases were identified by a novel coronavirus named severe acute 64 respiratory syndrome coronavirus 2 (SARS-CoV-2) was reported by Wuhan City, Hubei Province, 65 China. The disease was named as coronavirus disease 2019 (COVID-19) by the World Health 66 Organization in 11 February 2020. The patients may be related to have contacted with Huanan 67 Seafood Wholesale Market[1], and the virus can continuously transmit from person-to-person with 68 basic reproduction number (R_0) of 2.2 (95% confidence interval [CI], 1.4 to 3.9)[2]. Another research 69 estimated that the R_0 of SARS-CoV-2 was 2.68 (95% CI, 2.47 to 2.86)[3]. 70 Age-related transmissibility of COVID-19 has become a public health concern. The reported cases 71 at the early stage (before 2 January 2020) of the transmission were higher than 18 years old and 72 mainly were on 25-64 years old[1]. A study with 425 patients indicated that the median age was 59 73 years (range, 15 to 89) and reported different case distributions in four age groups: 0 - 14, 15 - 44, 45 74 -64, and ≥ 65 years[2]. However, there is not enough epidemiological evidence to classify the age 75 groups in transmission. In this study, we added another age classification scenario according to our 76 previous research on influenza that people were divided into five age groups: 0-5, 6-14, 15-24, 2577 -59, and ≥ 60 years[4], to better understand the transmissibility of the disease for different ages.

There were several studies focusing on mathematical modelling on COVID-19[2, 3], calculating the R_0 by using the serial intervals and intrinsic growth rate[2, 5, 6], or using ordinary differential equations and Markov Chain Monte Carlo methods[3]. We also developed a Bats-Hosts-Reservoir-People (BHRP) transmission network model and simplified the BHRP model as Reservoir-People (RP) transmission network model to calculate the transmissibility of SARS-CoV-2[7]. However, there is no age-specific mathematical model for quantifying the age-specific transmissibility SARS-CoV-2.

84	In this study, based on the RP model which we developed, we built an age-specific susceptible –
85	exposed – symptomatic – asymptomatic – recovered – reservoir (SEIARW) model. The age-specific
86	SEIARW model was employed to estimate the age-specific transmissibility of SARS-CoV-2 by fitting
87	the data from 26 November to 23 December, 2019 from published literature[1].
88	
89	Methods
90	Data collection
91	The data of COVID-19 were collected in Wuhan City from 26 November, 2019 to 23 December,
92	2019 from a published literature[1], including 29 COVID-19 cases with the onset date and exposure
93	history. The data of total population which were used in our model was from Wuhan City Statistics
94	Bureau. The data of age group proportions, birth rate and death rate in our model was obtained from
95	Wuhan Statistical Yearbook.
96	Study design
97	In this study, people were divided into 4 age groups based on the published literature[1], and 5 age
98	groups based on our previous study[4]. We used the subscript <i>i</i> and <i>j</i> to represent age group 1 to 4 ($i \neq j$;
99	1: 0 – 14 years; 2: 15 – 44 years; 3: 45 – 64 years; 4: ≥ 65 years) and 1 to 5 ($i \neq j$; 1: 0 – 5 years; 2: 6-14
100	years; 3: 15-24 years; 4: 25-59 years; 4: \geq 60 years), respectively.
101	Age-specific transmission model
102	Our previous study showed that the SEIARW model could be adopted to simulate the infectious
103	diseases transmitted from reservoir (such as water or food) to people and from person to person[8, 9].
104	In this study, the age-specific SEIARW model was developed based on two transmission routes (form

105 market to person and person to person). In the model, people were divided into five compartments:

- 106 susceptible (S), exposed (E), symptomatic (I), asymptomatic (A), recovered (R), and the seafood
- 107 market were defined as reservoir (*W*). The definitions of the six compartments were shown in Table 1.
- 108 The model was based on following assumptions:
- a) Susceptible individuals become infected by contact with infected/asymptomatic people;
- b) SARS-CoV-2 can transmit within each age group. The transmission rate of age group *i* and *j* are
- 111 β_{ii} and β_{jj} respectively.
- 112 c) SARS-CoV-2 can be transmitted between different age groups. The transmission rate from age
- 113 group *i* to *j* is β_{ij} and from *j* to *i* is β_{ji} .
- d) Susceptible people will be infected after contact with seafood market, and the infection rate
- 115 coefficient is β_w ;
- e) All the newborn individuals which were assumed as susceptible were added into age group 1.
- 117 Each age group has a natural mortality rate. We set the natural birth rate is *br*, and the natural mortality
- 118 rate is dr;
- f) The incubation period of exposed person is 1/*a*. The exposed person will become asymptomatic
- 120 people after a latent period of $1/\omega$. We describe $p (0 \le p \le 1)$ as the proportion of asymptomatic
- 121 infection. Exposed individuals will become asymptomatic person A with a daily rate of *pE*, and become
- 122 symptomatic person with a daily rate of (1-*p*)*E*;
- 123 g) The transmissibility of A is κ times ($0 \le \kappa \le 1$) of that of I.
- h) The individuals I and A will become recovered person (R) after an infectious period of $1/\gamma$ and
- 125 $1/\gamma'$. A part of *I* would die due to the infection. We assumed that the case fatality rate was *f*.
- i) *I* and *A* individuals can shed pathogens into *W* with the shedding rate of μI and $\mu'A$, where μ and
- 127 μ' are the shedding coefficients;

- j) SARS-CoV-2 will remove in the market after a given period (the life time of the virus is $1/\varepsilon$), and
- 129 the daily decreasing rate of the pathogen is εW .
- 130 The flowchart of the model was shown in Figure 1. The equations of the age-specific SEIARW
- 131 model were shown as follows:

$$\begin{aligned} \frac{dS_i}{dt} &= brN - \beta_{ii}S_i(l_i + \kappa A_i) - \beta_{ji}S_i(l_j + \kappa A_j) - \beta_w S_i W - drS_i \\ \frac{dE_i}{dt} &= \beta_{ii}S_i(l_i + \kappa A_i) + \beta_{ji}S_i(l_j + \kappa A_j) - (1 - p)\omega E_i - p\omega' E_i - dr E_i \\ \frac{dI_i}{dt} &= (1 - p)\omega E_i - \gamma I_i - (dr + f)I_i \\ \frac{dA_i}{dt} &= p\omega' E_i - \gamma' A_i - dr A_i \\ \frac{dR_i}{dt} &= \gamma I_i + \gamma' A_i - dr R_i \\ \frac{dS_j}{dt} &= brN - \beta_{jj}S_j(l_j + \kappa A_j) - \beta_{ji}S_j(l_i + \kappa A_i) - \beta_w S_j W - dr S_j \\ \frac{dE_j}{dt} &= \beta_{jj}S_j(l_j + \kappa A_j) + \beta_{ji}S_j(l_i + \kappa A_i) + \beta_w S_j W - (1 - p)\omega E_j - p\omega' E_j - dr E_j \\ \frac{dI_j}{dt} &= (1 - p)\omega E_j - \gamma I_j - (dr + f)I_j \\ \frac{dA_j}{dt} &= p\omega' E_j - \gamma' A_j - dr A_j \\ \frac{dR_j}{dt} &= p(I_i + I_j) + \mu' (A_i + A_j) - \varepsilon W \\ N &= S_i + E_i + I_i + A_i + R_i + S_j + E_j + I_j + A_j + R_j \end{aligned}$$

132

133 Since the different dimension of people and the virus, we adopt the following sets to perform the134 normalization:

135
$$s_i = S_i/N$$
, $e_i = E_i/N$, $i_i = I_i/N$, $a_i = A_i/N$, $r_i = R_i/N$, $s_j = S_j/N$, $e_j = E_j/N$, $i_j = I_j/N$, $a_j = A_j/N$,

136
$$r_j = R_j/N, w = \varepsilon W/\mu N, \mu' = c\mu, b_{ii} = \beta_{ii}N, b_{ij} = \beta_{ij}N, b_{jj} = \beta_{jj}N, b_{ji} = \beta_{ji}N, b_w = \mu\beta_w N/\varepsilon.$$

138 In the normalization, parameter *c* refers to relative shedding coefficients of *A* compared to *I*. The

139 normalized model is expressed as follows:

$$\begin{aligned} \frac{as_i}{dt} &= brn - b_{ii}s_i(i_i + \kappa a_i) - b_{ji}s_i(i_j + \kappa a_j) - b_w s_i w - drs_i \\ \frac{de_i}{dt} &= b_{ii}s_i(i_i + \kappa a_i) + b_{ji}s_i(i_j + \kappa a_j) - (1 - p)\omega e_i - p\omega' e_i - dre_i \\ \frac{di_i}{dt} &= (1 - p)\omega e_i - \gamma i_i - (dr + f)i_i \\ \frac{da_i}{dt} &= p\omega' e_i - \gamma' a_i - dra_i \\ \frac{dr_i}{dt} &= pi_i + \gamma' a_i - drr_i \\ \frac{ds_j}{dt} &= brn - b_{jj}s_j(i_j + \kappa a_j) - b_{ji}s_j(i_i + \kappa a_i) - b_w s_j w - drs_j \\ \frac{de_j}{dt} &= b_{jj}s_j(i_j + \kappa a_j) + b_{ji}s_j(i_i + \kappa a_i) + b_w s_j w - (1 - p)\omega e_j - p\omega' e_j - dre_j \\ \frac{di_j}{dt} &= (1 - p)\omega e_j - \gamma i_j - (dr + f)i_j \\ \frac{da_j}{dt} &= p\omega' e_j - \gamma' a_j - dra_j \\ \frac{dr_j}{dt} &= pi_j + \gamma' a_j - drr_j \\ \frac{dw}{dt} &= \varepsilon(i_i + i_j + c(a_i + a_j) - w) \\ n &= s_i + e_i + j_i + a_i + r_i + s_j + e_j + i_j + a_j + r_j \end{aligned}$$

140 The left side of the equation shows the instantaneous rate of change of S, E, I, A, R and W at time

141 *t*. The subscript *i* and j ($i \neq j$) represent age group 1 to 4/5 in the equation, respectively.

142 Parameters estimation

The mean incubation period was 5.2 days (95% confidence interval [*CI*]: 4.1–7.0) [2]. As reported by Xu *et al*[10], the median time from exposure to onset of illness (infected) was 4 days (interquartile range 3-5 days). Another study showed that the mean of incubation period was around 5 days and the period falls within the range of 2-14 days [11]. We set the 5-day as the incubation period and the latent period in this study. Thus, $\omega = \omega' = 0.2$.

Since there was no data on the proportion of asymptomatic infection of the virus, we simulated the baseline value of proportion of 0.5 (p = 0.5). Since there was no evidence about the transmissibility of asymptomatic infection, we assumed that the transmissibility of asymptomatic infection was 0.5 times that of symptomatic infection ($\kappa = 0.5$), which was the similar value as influenza[12]. We assumed that the relative shedding rate of *A* compared to *I* was 0.5. Thus, c = 0.5. Since the SARS-CoV-2 is an RNA virus, we assumed that it could be died in the environment in a short time, but it could stay for a longer time (10 days) in the unknown hosts in the market. We set $\varepsilon = 0.1$.

155 There is a mean 5-day delay from symptom onset to detection/hospitalization of a case (the cases 156 detected in Thailand and Japan were hospitalized from 3-7 days after onset, respectively) [13-15]. 157 Another study showed that the mean time from illness onset to hospital admission (for treatment and/or 158 isolation) was estimated at 3-4 days without truncation and at 5-9 days when right truncated[11]. As 159 reported by Xu et al[10], the median time from onset of illness to first hospital admission was 2 (range 160 from 1-4) days. A study including 45 patients diagnosed before January 1 was estimated to have a 161 mean of 5.8 days (95% CI: 4.3-7.5) from illness onset to first medical visit[2]. In our model, we set the 162 infectious period of the cases as 6 days. Therefore, $\gamma = \gamma' = 0.1667$.

According the official report by National Health Commission of the People's Republic of China, we collected the data of daily fatality from January 24th to 30th, 2020[16]. Therefore, we set *f* as 0.02348 (range: 0.02198-0.03186). In this study, we set the total population as 11,081,000 so that $br = 4.266 \times 10^{-5}$ and $dr = 3.184 \times 10^{-5}$ based on the 2018 Wuhan Statistical Yearbook (Table 2).

167 A two-step curve fitting method was adopted to estimate the parameter b_{W} , b_{ii} , b_{jj} , b_{ij} , and b_{ji} . At the 168 first step, we used the mix-age SEIARW model to fit the data and to estimate the parameter b_{W} and 169 mix-age *b* (named as b_{p}). At the second step, we used the age-specific SEIARW model to fit the data

170 and to estimate the parameter b_{ij} , b_{jj} , b_{ij} , and b_{ji} .

171 Quantification of the age-specific transmissibility of SARS-CoV-2

- 172 In the model the age-specific secondary attack rate (SAR) was employed to quantify the
- 173 transmissibility of SARS-CoV-2. They were calculated as follows:

 $SAR_{ii} = \beta_{ii}/\gamma$

 $SAR_{ij} = \beta_{ij}/\gamma$

 $SAR_{ji} = \beta_{ji}/\gamma$

$$SAR_{jj} = \beta_{jj}/\gamma$$

174 In the equations, SAR_{ii} , SAR_{ij} , SAR_{jj} , and SAR_{jj} refer to the age-specific SAR among age group *i*,

175 from age group *i* to *j*, from age group *j* to *i*, and among age group *j*, respectively.

To quantify the age-specific transmissibility of SARS-CoV-2, we also performed a simulation which was named as "knock-out" simulation in our previous study[17]. In this study, the "knock-out" simulation was defined as cutting off the route of transmission between or within different age groups, and was performed in the following scenarios: A) $\beta_{ii} = 0$; B) $\beta_{ji} = 0$; C) $\beta_{ij} = 0$; D) $\beta_{jj} = 0$; E) control (no cutting off transmission route).

181 Simulation method and statistical analysis

Berkeley Madonna 8.3.18 (developed by Robert Macey and George Oster of the University of California at Berkeley. Copyright ©1993-2001 Robert I. Macey & George F. Oster) was employed to perform the curve fitting and the simulation. The simulation methods (Runge-Kutta method of order four with tolerance set at 0.001) were the same as the previously published researches [18-24]. The data was analyzed and figured by using Microsoft Office Excel 2010 (Microsoft, Redmond, WA, USA) and GraphPad Prism 7.0 (GraphPad Software, La Jolla, CA). The goodness of fit was judged by coefficient

188 of determination (R^2) and Chi-square (χ^2) value calculated by SPSS 21.0 (IBM Corp, Armonk, NY, 189 USA).

190 Sensitivity analysis

- 191 Since nine parameters (p, κ , c, ε , ω , ω' , γ , γ' and f) in the models were collected from literatures,
- 192 there might be some uncertainties in our model. Therefore, we performed sensitivity analysis of
- parameters p, κ , c, ε , ω , ω' , γ , γ' and f by splitting them into 1,000 values ranging from 0 0.9, 0 1, 0 –
- 194 1, 0 1, 0.1428 0.25, 0.1428 0.25, 0.125 0.25, 0.125 0.25, 0.02198 0.03186, respectively
- 195 (Table 2).
- 196
- 197 Results

198 Epidemiological characteristics and curve fitting of SARS-CoV-2

199There were 29 COVID-19 cases data collected in Wuhan City from 26 November, 2019 to 23200December, 2019, among which 10 cases had a history of the seafood market exposure and 19 cases

201 had no history of the exposure (Figure 2).

202 The mix-age SEIARW model including all age groups fitted the reported data well ($\chi^2 = 4.77 \times 10^{-6}$,

203 P > 0.999). The results of the curve fitting were shown in Figure 3. The results also showed that the b_p

- 204 = 1.1329 and b_W = 0.5255. The age-specific SEIARW model fitted with the reported data well by
- dividing the population into four age groups ($\chi^2 = 4.99 \times 10^{-6}$, P > 0.999), and five age groups ($\chi^2 = 4.85$)
- $\times 10^{-6}$, *P* > 0.999). Thus, the prevalence of COVID-19 in each age group was simulated in Figure 4.

207 Transmissibility of SARS-CoV-2

Based on the four-age-group SEIARW model, the highest transmissibility occurred from age group 209 2 to 3 ($SAR_{23} = 17.56$ per 10 million persons), followed by from age group 3 to 2 ($SAR_{32} = 10.17$ per 10 million persons), from age group 4 to 4 ($SAR_{44} = 6.99$ per 10 million persons), and from age group 3 to 4 ($SAR_{34} = 5.69$ per 10 million persons). The lowest transmissibility occurred from age group 1 to 2 ($SAR_{12} = 0.002$ per 10 million persons), followed by from age group 3 to 1 ($SAR_{31} = 0.003$ per 10 million persons), from age group 4 to 2 ($SAR_{42} = 0.52$ per 10 million persons), and from age group 1 to 3 ($SAR_{13} = 1.08$ per 10 million persons) (Figure 5-A).

Based on the five-age-group SEIARW model, the highest transmissibility occurred from age group 4 to 5 ($SAR_{45} = 12.40$ per 10 million persons), followed by from age group 5 to 4 ($SAR_{54} = 6.61$ per 10 million persons), from age group 4 to 4 ($SAR_{44} = 5.08$ per 10 million persons), and from age group 2 to 1 ($SAR_{21} = 4.90$ per 10 million persons). The lowest transmissibility occurred from age group 3 to 4 ($SAR_{34} = 0.0002$ per 10 million persons), followed by from age group 1 to 1 ($SAR_{11} = 0.22$ per 10 million persons), from age group 1 to 3 ($SAR_{13} = 1.54$ per 10 million persons), and from age group 1 to 2 ($SAR_{12} = 1.59$ per 10 million persons) (Figure 5-B).

The results of the "knock-out" simulation showed that, based on the four-age-group SEIARW model, the scenarios $b_{23} = 0$, $b_{32} = 0$, and $b_{22} = 0$ led to the highest decrease for the number of cases. However, the scenarios $b_{12} = 0$, $b_{31} = 0$, and $b_{11} = 0$ led to the lowest decrease for the number of cases (Figure 6-A). Based on the five-age-group SEIARW model, the scenarios $b_{44} = 0$, $b_{45} = 0$, and $b_{54} = 0$ led to the highest decrease for the number of cases. However, the scenarios $b_{12} = 0$, $b_{31} = 0$, and $b_{11} = 0$ led to the highest decrease for the number of cases (Figure 6-B).

228 Sensitivity analysis

- 229 The results of sensitivity analysis showed that the models were slightly sensitive to parameters p,
- 230 ω , and γ but not sensitive to parameters κ , c, ε , f, ω ', and γ ' (Figures 7 8).
- 231

232 Discussion

233	This is the first study to develop an age-specific SEIARW model to quantify the transmissibility of
234	COVID-19 among different age groups. The results showed that our model fitted the reported data well,
235	thus has the capability of estimating or predicting? the age-specific transmissibility of the virus.
236	Based on the four-age-group SEIARW model, the highest transmissibility occurred between the
237	age groups 15 – 44 years and 45 – 64 years, among those \geq 65 years, or from 45 – 64 years to \geq 65
238	years. The lowest transmissibility occurred from age group 0-14 years to 15 – 44 years, or from 45 – 64
239	years to \leq 14 years. Based on the five-age-group SEIARW model, the highest transmissibility occurred
240	between age group 25 – 59 years and \geq 60 years, or among 25 – 59 years. The lowest transmissibility
241	occurred from age group 15 – 24 years to 25 – 59 years, or from age group 0-5 years to 6-14 years, or
242	to 15-24 years. The "knock-out" simulation had the similar results. We concluded that the virus was
243	more likely to transmit among older population.
244	These results revealed that SARS-CoV-2 has high transmissibility among adult older than 25 years
245	old or elder people, but has low transmissibility among children or people younger than 14 years.
246	These results were similar to the age-specific transmissibility of influenza A (H1N1)[4]. The age-specific
247	control and prevention interventions are needed.

The reasons for the difference of age-specific transmissibility remain unclear. It might be related to the different contact characteristics among different age groups. Adults were more likely to work outside and came into contact with different individuals in work places, buses, subways, or even airplanes. However, children or younger people could stay at home during the outbreak, and they were less likely to get infected, or if they were infected, it was mostly from adults or elder people in the same family.

253 Our findings were based on the parameter estimation and the data from literatures. It is known that

254 the asymptomatic infection of COVID-19 exists. As reported by Bai et al[25], 1 of 6 cases was 255 asymptomatic infected, with the proportion of asymptomatic of 0.17. As reported by Pan et al[26], 2 of 3 256 cases was asymptomatic infected, with the proportion of asymptomatic of 0.67. There was a research 257 showed that the asymptomatic infection would shed SARS-CoV-2 for 5 days[27]. However, there was 258 not enough evidence to provide the clear epidemiological estimates of the parameters p, κ , ω' , and γ' 259 which were related to asymptomatic characteristics. Although there was clinical evidence to calculate 260 the parameters related to symptomatic cases such as incubation period, fatality rate, and duration from 261 symptoms onset to diagnosis[1, 2], more epidemiological data are needed to explore the parameters. 262 However, we performed a sensitivity analysis of all the nine parameters from the literatures, and we 263 found that the models was slightly sensitive to parameters p, ω , and γ but not sensitive to parameters κ , 264 c, ε , f, ω' , and γ' . Therefore, our results might be affected slightly by the parameter estimation. 265 Since we could not obtain the first-hand data, the results of our findings might have some 266 uncertainties. However, our study aimed to develop and provide an age-specific transmission model for

267 the public health department who has the big data to investigate the age-specific transmissibility in 268 real world scenarios.

269 Conclusions

By calculating the published data, our model showed that SARS-CoV-2 has a high transmissibility among adults and elder people but low transmissibility among children and young people. Our results provide a mathematical model for investigating the age-specific transmissibility of SARS-CoV-2. More data and studies are needed to estimate the age-specific transmissibility in real world scenarios to better understand the characteristics of the widely spread virus.

276 Abbreviations

- 277 SARS-CoV-2: severe acute respiratory syndrome coronavirus 2; COVID-19: coronavirus disease 2019;
- 278 *R*₀: basic reproduction number; CI: confidence interval; BHRP: Bats-Hosts-Reservoir-People; RP:
- 279 Reservoir-People; SEIARW: susceptible exposed symptomatic asymptomatic recovered -
- 280 reservoir; SAR: secondary attack rate; R^2 : coefficient of determination.

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288 Availability of data and materials

- Not applicable.
- 290 Authors' contributions
- 291 TMC and ZYZ designed research; TMC, YHS, BHZ, and ZYZ conceived the experiments, TMC, ZYZ,
- 292 YZZ, JWX, ZL, JR, XCL, YW, MY, LL, SSY, JL, RYL, FX, YYS, and YCC conducted the experiments
- and analyzed the results; TMC, ZYZ, and QQH wrote the manuscript. All authors read and approved
- the final manuscript.

295 Competing interests

- 296 The authors declare that they have no competing interests.
- 297 Consent for publication

- 298 Not applicable.
- 299 Ethics approval and consent to participate 300 Not applicable. 301 302 303 304 References 305 Huang C, Wang Y, Li X, Ren L, Zhao J, Hu Y, et al. Clinical features of patients infected with 1. 306 2019 novel coronavirus in Wuhan, China. Lancet. 2020;395:497-506. 307 2. Li Q, Guan X, Wu P, Wang X, Zhou L, Tong Y, et al. Early Transmission Dynamics in Wuhan, 308 China, of Novel Coronavirus-Infected Pneumonia. N Engl J Med. 2020; doi: 309 10.1056/NEJMoa2001316. 310 3. Wu JT, Leung K, Leung GM. Nowcasting and forecasting the potential domestic and 311 international spread of the 2019-nCoV outbreak originating in Wuhan, China: a 312 modelling study. Lancet. 2020; doi: 10.1016/S0140-6736(20)30260-9. 313 4. Liu R, Leung RK, Chen T, Zhang X, Chen F, Chen S, et al. The Effectiveness of Age-Specific 314 Isolation Policies on Epidemics of Influenza A (H1N1) in a Large City in Central South 315 China. PloS one. 2015; 10:e0132588. 316 5. Zhao S, Lin Q, Ran J, Musa SS, Yang G, Wang W, et al. Preliminary estimation of the basic 317 reproduction number of novel coronavirus (2019-nCoV) in China, from 2019 to 2020: A 318 data-driven analysis in the early phase of the outbreak. Int J Infect Dis. 2020;92:214-7. 319 6. Zhao S, Musa SS, Lin Q, Ran J, Yang G, Wang W, et al. Estimating the Unreported Number

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381 Figures

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384 Figure 1. Flowchart of the age-specific SEIARW model



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387 Figure 2. Epidemic curve of collected data of reported COVID-19 cases in Wuhan City from 26

388 November, 2019 to 23 December, 2019



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391 Figure 3. Results of curve fitting of the mix age SEIARW model to the reported data



394 Figure 4. Results of curve fitting of the age-specific SEIARW model to the reported data. A-a:

393

curve fitting based on four age groups; A-b: the simulated prevalence by the age-specific SEIARW
model based on the four age groups; B-a: curve fitting based on five age groups; B-b: the simulated
prevalence by the age-specific SEIARW model based on the five age groups.





400 Figure 5. SAR simulated from the age-specific SEIARW model. A: SAR based on four age groups;

401 B: SAR based on five age groups.



404 Figure 6. Results of the "knock-out" simulation from the age-specific SEIARW model. A: results

405 based on four age groups; B: results based on five age groups.

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408 Figure 7. Results of sensitivity analysis of *p*, *κ*, *c*, and *ε*. A: sensitivity analysis of *p*; B: sensitivity

409 analysis of κ ; C: sensitivity analysis of c; D: sensitivity analysis of ϵ .

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411

412 Figure 8. Results of sensitivity analysis of ω , ω' , γ , γ' and f. A: sensitivity analysis of ω ; B:

413 sensitivity analysis of ω '; C: sensitivity analysis of γ ; D: sensitivity analysis of γ '; F: sensitivity analysis of

414 f.

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417 Tables

418 Table 1. Variables in the age-specific SEIARW model

Variables	/ariables Description	
Si	Susceptible individuals density of age group <i>i</i>	Individuals.km ⁻²
S_j	Susceptible individuals density of age group <i>j</i>	Individuals · km ⁻²
E _i	Exposed individuals density of age group <i>i</i>	Individuals · km ⁻²
E_j	Exposed individuals density of age group <i>j</i>	Individuals · km ⁻²
l _i	Infectious individuals density age group <i>i</i>	Individuals · km ⁻²
l _j	Infectious individuals density age group <i>j</i>	Individuals · km ⁻²
A _i	Asymptomatic individuals density age group <i>i</i>	Individuals · km ⁻²
A_j	Asymptomatic individuals density age group <i>j</i>	Individuals.km ⁻²
R _i	Recovered individuals density age group <i>i</i>	Individuals.km ⁻²
R_{j}	Recovered individuals density age group <i>j</i>	Individuals · km ⁻²
W	Pathogen concentration in the seafood market	Viruses⋅mL ⁻³
Ν	Total number of population density	Individuals.km ⁻²

419

Parameter	Description	Unit	Value	Range	Method
β _{<i>ii</i>} *	Transmission relative rate among age group <i>i</i>	Individuals ⁻¹ .days ⁻¹	-	-	-
<i>b</i> ;; [*]	Scaled <i>i</i> -to- <i>i</i> contact rate	days ⁻¹	-	≥ 0	Curve fitting
${oldsymbol{eta}_{ij}}^{*}$	Transmission relative rate from age group <i>i</i> to <i>j</i>	Individuals ⁻¹ .days ⁻¹	-	-	-
b _{ij} *	Scaled <i>i</i> -to- <i>j</i> contact rate	days ⁻¹	-	≥ 0	Curve fitting
β_{ji} *	Transmission relative rate from age group <i>j</i> to <i>i</i>	Individuals ⁻¹ .days ⁻¹	-	-	-
<i>b_{ji}*</i>	Scaled <i>j</i> -to- <i>i</i> contact rate	days ⁻¹	-	≥ 0	Curve fitting
${oldsymbol{eta}_{jj}}^{*}$	Transmission relative rate among age group <i>i</i>	Individuals ⁻¹ .days ⁻¹	-	-	-
b _{jj} *	Scaled <i>j</i> -to- <i>j</i> contact rate	days ⁻¹	-	≥ 0	Curve fitting
β_w	Seafood market contact rate	mL ⁻³ ·cells ⁻¹ ·days ⁻¹	-	-	-
b _w	Scaled market-to-person contact rate	days ⁻¹	0.9122	≥ 0	Curve fitting
	Relative transmissibility rate of asymptomatic to	A	0.5	0.4	Deference [40]
К	symptomatic individuals	1	0.5	U-1	Reference [12]

421 Table 2. Description and values of parameters in the age-specific SEIARW model

p	Proportion of the asymptomatic	1	0.5	0-1	Reference [12]
(4)	Incubation relative rate	days ⁻¹	0.2	0.1428-0.25	References [2, 10,
ω					11]
ω'	Latent relative rate	days ⁻¹	0.2	0.1428-0.25	References [2, 10,
					11]
V	Removed rate of the infectious	days ⁻¹	0.1667	0.125-0.25	References [2, 10,
Ŷ					11, 13-15]
<i></i>	Removed rate of the asymptomatic	days ⁻¹	0.1667	0.125-0.25	References [2, 10,
Ŷ					11, 13-15]
f	Fatality of the disease	1	0.02348	0.02198-0.03186	References [12]
3	Pathogen lifetime relative rate	days ⁻¹	0.1	-	Assumption
	Person-to-reservoir contact rate ("shedding"	cells·mL ⁻³ ·day ⁻¹ ·km ² ·individuals ⁻¹	-		
μ	by Infectious)			-	-
μ'	Person-to-reservoir contact rate ("shedding" by	cells·mL ⁻³ ·day ⁻¹ ·km ² ·individuals ⁻¹	-	-	-

	Asymptomatic)				
0	Relative shedding rate of asymptomatic	1	0.5	0.1	Poforonco [12]
C	compared to infectious	0.5		0-1	
br	Birth rate of the population	1	0.00004266		Wuhan Statistical
			0.00004200		Yearbook
dr	Death rate of the population	1	0 00003184	-	Wuhan Statistical
			0.00003184		Yearbook

422 *: *i* and *j* represent age group 1 to 4/5, respectively, and $i \neq j$