INTRODUCTION

Coronavirus disease 2019 (COVID-19) is an infectious disease caused by the new coronavirus (2019-nCoV) [1]. It began spreading in Wuhan, China, in December 2019, and on January 14, 2020, the World Health Organization officially stated that there was a potential for inter-personal infection of COVID-19 in a limited range between families, and on January 22, 2020, the officials in Wuhan, China, reported that there was evidence of COVID-19 infection among people [2]. In Korea, the first patient was confirmed with COVID-19 on January 20, 2020. Thereafter, 30 confirmed cases appeared by February 16, 2020, and the rate of increase in confirmed cases was limited to one or two cases per day. However, the situation rapidly changed after the #31 confirmed case on February 18, 2020 in Daegu city. As of March 26, 2020, of the 9,241 confirmed cases in Korea, 6,482 confirmed cases occurred in Daegu, accounting for 70.1% of the total cases. Among them, 4,391 (67.7%) confirmed cases in Daegu were related to Shincheonji. Assuming there are 2.5 million citizens in Daegu, and 9,000 members are part of Daegu Shincheonji, the 2019-nCoV infection rate of Daegu citizens was 0.08%, while the infection rate of Daegu Shincheonji members was 48.78%, which is 583 times higher. This shows that the COVID-19 epidemic in

METHODS

Indirect-based simulation model for COVID-19 transmission in Daegu, Korea

Woo-Sik Son¹, RISEWIDs Team*
¹National Institute for Mathematical Sciences, Daejeon, Korea

OBJECTIVES: The aims of this study were to obtain insights into the current coronavirus disease 2019 (COVID-19) epidemic in the city of Daegu, which accounted for 6,482 of the 9,241 confirmed cases in Korea as of March 26, 2020, to predict the future spread, and to analyze the impact of school opening.

METHODS: Using an individual-based model, we simulated the spread of COVID-19 in Daegu. An individual can be infected through close contact with infected people in a household, at work/school, and at religious and social gatherings. We created a synthetic population from census sample data. Then, 9,000 people were randomly selected from the entire population of Daegu and set as members of the Shincheonji Church. We did not take into account population movements to and from other regions in Korea.

RESULTS: Using the individual-based model, the cumulative confirmed cases in Daegu through March 26, 2020, were reproduced, and it was confirmed that the hotspot, i.e., the Shincheonji Church had a different probability of infection than non-hotspot, i.e., the Daegu community. For 3 scenarios (I: school closing, II: school opening after April 6, III: school opening after April 6 and the mean period from symptom onset to hospitalization increasing to 4.3 days), we predicted future changes in the pattern of COVID-19 spread in Daegu.

CONCLUSIONS: Compared to scenario I, it was found that in scenario III, the cumulative number of patients would increase by 107 and the date of occurrence of the last patient would be delayed by 92 days.

KEY WORDS: COVID-19, Mathematical model, Model prediction, Infections, Korea
Daegu was because the Shincheonji Church in Daegu became a hotspot, leading to the spread of COVID-19 in the Daegu community.

A compartmental model [3], which is widely used as a mathematical study model for the spread of infectious diseases, generally divides the entire population into several groups depending on the state of infection, but it is not an appropriate model to reproduce the hotspot and non-hotspot (communities that are not members of Shincheonji) groups that appear in Daegu. In addition, although a two-patch model [4], that divides the entire population into two groups, was proposed to include hotspots and non-hotspots in the compartmental model, there are difficulties in reproducing infections caused by close contact between the two groups.

To overcome the above problems, we simulated the COVID-19 spread in Daegu using an individual-based model suggested by Ferguson et al. [5]. In the individual-based model, each individual can be infected through contact with infected persons in households, workplaces/schools, and communities (religious and social gatherings), and for this, a virtual population of the same size as the population in Daegu was created. The purpose of this study was to use the individual-based model to understand the current COVID-19 epidemic in Daegu, predict the future spread, and analyze how the reopening of schools scheduled for April 6, 2020 will affect the spread of COVID-19.

MATERIALS AND METHODS

The individual-based model [5] simulates the spread of infectious diseases through close contact between people in households, workplaces/schools, and communities. Through the individual-based model, a virtual population group of the size of Daegu was created to understand the spread of COVID-19 in Daegu and predict future spread. Each individual in the virtual population group has information regarding the household, work/school, and community. We generated a virtual population of 2,171,000 people from the 2015 census 2% sample data of the MicroData Integrated Service (as of 2015) [6]. Household information and the status of students and workers were set using the age and commuting status of each individual listed in the census data. Assuming that close contact at work or school occurs in the same office or classroom, a student's classroom ID and employee's office ID were virtually generated as follows: the classroom ID was set by randomly selecting students of the same age, city, and district so that an average of 30 students were assigned to the same classroom. Figure 1A shows the histogram according to the number of students in each classroom in Daegu. The office ID was set by randomly selecting the workers in Daegu without the distinction of municipalities and assigning an average of 20 people to the same office. Figure 1B shows the histogram according to the number of workers in each office in Daegu.

Table 1 shows some of the virtual population groups created in this way. Each row represents one individual, and each column represents the attributes of the individual used to simulate the spread of the infectious disease. Individuals with the same household, classroom, and office IDs belong to the same household, classroom, and office, respectively. If the classroom ID and the office ID are marked as NA, it means that the individual is neither a student nor worker. Items on hotspot indicates whether the individual is a member of Shincheonji or not, and 9,000 people were randomly selected from the population of Daegu and assigned to the hotspot. The infection status in the last column of Table 1 represents the infection status of each individual on a certain date, and the possible infection status were as follows: those susceptible
Table 1. Each row of the synthetic population data represents one individual, and each column represents the attributes of the individual used to simulate the spread of coronavirus disease 2019 (COVID-19).

| Individual | Household ID | Age (yr) | Class- | Office | Hotspot | Infectious status |
|------------|--------------|----------|room ID| ID     |         |                 |
| 1          | 1            | 48       | NA    | 3      | False   | S                |
| 2          | 1            | 44       | NA    | NA     | True    | I                |
| 3          | 1            | 15       | 2 NA  | False  | S        |                 |
| 4          | 2            | 45       | NA    | 3      | False   | S                |
| 5          | 2            | 43       | NA    | NA     | False   | S                |
| 6          | 2            | 17       | 25 NA | False  | S        |                 |
| 7          | 3            | 51       | NA    | 5      | False   | S                |
| 8          | 3            | 50       | NA    | NA     | False   | S                |
| 9          | 3            | 17       | 25 NA | False  | S        |                 |

NA, not available; S, susceptible; I, infectious.

To COVID-19 - susceptible (S); those in the latent stage after infection - latent (L); those that can infect the susceptible - infectious (I); those isolated after being confirmed with COVID-19 - hospitalization (H); and those that recovered or died from COVID-19 - recovered (R). Figure 2 is a diagram showing the change in each infection status. λ is the infection probability of the susceptible and was calculated as follows [5]:

\[
\lambda = \frac{\beta_h/N_h}{N_0} + \frac{\beta_w/N_w}{N_0} + \frac{\beta_s/N_s}{N_0} + \frac{\beta_{hot}/3/N_{hot}}{N_0}
\]

β_h (β_w, β_s, β_{hot}) is the probability of encountering an infected individual in the household (work, school, hotspot) and getting infected. We set this as β_h = β_w = β_s = 2β, same as that set by Ferguson et al. [5]. N_h (N_w, N_s, N_{hot}) represents the total number of households (work, school, hotspot), and N_h (N_w, N_s, N_{hot}) represents the total number of those infected in households (work, school, hotspot). 1/κ = 5.2 [7], 1/α = 4.3 [8], and 1/η = 14 are the average latent period (days), average period between symptom onset to confirmation (days), and average period from being confirmed to recovery (days), respectively.

In February 1, 2020, 10 infected individuals in the hotspot were set as the initial confirmed patients. The #31 patient was confirmed with COVID-19 on February 18, 2020, but it was assumed that the symptoms started on February 7, 2020, and considering the latent period of COVID-19, the #31 patient was assumed to be infected from the initial patients of the hotspot on February, 2020. The individual-based model was simulated on a daily basis, and it was assumed that the population inflow into Daegu using public transportation was minimal after the COVID-19 epidemic in Daegu. That is, it was assumed that there was no influx of new COVID-19 infections from other regions in Korea and from abroad.

Ethics statement

This research is based on data which is open to public. Neither ethical approval of an institutional review board nor written informed consent was required.

RESULTS

The individual-based model and all parameter values except for β_{hot} and β are described in the Materials and Methods section. Data on the date of onset of symptoms in confirmed patients with COVID-19 are not currently available. We set β_{hot} and β to realize the confirmed COVID-19 patients in Daegu until March 26, 2020 (4,391 cases in the hotspot, 2,091 in the non-hotspot). In the individual-based model, changes in infection status are statistically simulated; that is, the changes in which a susceptible person becomes a latent person is realized as follows. Every day, a uniform random number between 0 and 1 is generated for all susceptible people, and if this value is less than lambda, which is the probability of becoming latent after infection, that susceptible person becomes a latent patient. Since the random number changes every time it is generated, the simulation results can be different even with the same parameters and initial patient settings. Therefore, it is necessary to check the distribution of simulation results using different random seeds rather than a single simulation. We performed 100 simulations using different random seeds and confirmed whether the median of this result reproduces the statistics of cumulative confirmed cases as of March 26, 2020. Among β_{hot} and β, the parameter with a higher determinant power for reproducing the cumulative confirmed cases in Daegu was β_{hot}. When β_{hot} was determined, the results of the cumulative confirmed cases were realized, and while adjusting for β, a parameter value for reproducing the non-hotspot cumulative cases was found. The parameter results were β_{hot} = 3.06 and β = 0.33. This shows that the probability of infection between the hotspot and household and workplace differed by more than nine times. Figure 3 shows the cumulative confirmed cases and simu-
The reopening of elementary/middle/high schools on April 6, 2020, mentioned in scenarios II and III, is the current plan as of March 26, 2020. The assumption of scenario III that the average period from symptom onset to confirmation would increase again to 4.3 days considers the students' relatively passive expression of symptoms after the reopening of schools. The individual-based model simulation results for the above three scenarios are shown in Table 2, Figures 4 and 5. In scenario I, the number of cumulative confirmed cases in Daegu was 6,677 (4,394 in hotspot, 2,322 in non-hotspot), and the last newly confirmed case occurred on April 26, 2020. In scenario II, the number of cumulative confirmed cases in Daegu was 6,716 (4,394 in hotspot, 2,322 in the non-hotspot) and compared with scenario I (based on the median), 39 Daegu citizens were additionally infected (non-hotspot, not a member of Shincheonji). The last newly confirmed cases occurred on May 3, 2020, 7 days later than in scenario I. In scenario III, the number of cumulative confirmed cases were 6,784 (4,394 in hotspot, 2,390 in the non-hotspot), and 107 additional cases occurred on May 3, 2020, 7 days later than in scenario I. The median value of 1/α was set to 2.7 instead of 4.3 after February 29, 2020 to reflect the effect of the shortened 1/α, the average period from symptom onset to confirmation, after mass screening tests for the Shincheonji Church members.

The statistics of the confirmed cases as of March 26, 2020 were reproduced, and to predict the spread of COVID-19 in Daegu thereafter, we considered the three following scenarios.

- Scenario I: Maintaining vacations in elementary/middle/high schools
- Scenario II: Reopening of elementary/middle/high schools on April 6, 2020
- Scenario III: Reopening of elementary/middle/high schools on April 6, 2020 & after April 6, 2020, 1/α, the average period from symptom onset to confirmation, increases again to 4.3 days

| Parameters | Scenario I | Scenario II | Scenario III |
|------------|------------|-------------|--------------|
| $1/\alpha$ | $\beta_s = 0, \frac{1}{\alpha} = \frac{4.3}{2.7}$ (Feb 28) | $\beta_s = 2\beta_h, \frac{1}{\alpha} = \frac{4.3}{2.7}$ (Feb 28) | $\beta_s = 2\beta_h, \frac{1}{\alpha} = \frac{4.3}{2.7}$ (Feb 29-Apr 5) |
| Cumulative no. of hospitalization cases | 6,677 (median) | 6,716 (compared with scenario I + 39 cases) | 6,784 (compared with scenario I + 107 cases) |
| Hospitalization date of the last patient | Apr 26 | May 3 (compared with scenario I + 7 d) | Jul 27 (compared with scenario I + 92 d) |
Figure 4. Cumulative number of hospitalization cases in the simulation. Here, we show the median and 5th to 95th percentile range for (A) scenario I, (B) scenario II, and (C) scenario III.

Figure 5. Cumulative and daily hospitalization cases in the simulation (A, B) scenario I, (C, D) scenario II, and (E, F) scenario III. Here, we show the median of 100 simulation results for different random seeds.

The individual-based model was selected to explain the characteristics demonstrated by the spread of COVID-19 in Daegu, specifically, that the cumulative infection rate of Shincheonji members was about 583 times higher than that of non-Shincheonji members. Compared with the compartmental model which is widely used as a mathematical model for the spread of infectious diseases [3], the individual-based model simulates the transmission of infectious diseases through close contact among people using the socio-demographic information of each individual (household, workplace/school, community such as religious and social gatherings). Therefore, it has the advantage of being able to analyze the effect of quarantine policies, such as closing schools and implementing shifts at work, in preventing infection spread in more detail.

Using the individual-based model, we reproduced the cumulative COVID-19 confirmed cases of Daegu until March 26, 2020. The number of newly confirmed cases per day in Daegu sharply
increased from February 21, 2020, after the #31 confirmed case, and the largest number of confirmed cases was reported on February 29, 2020, with 656 cases. Since then, and after March 11, 2020 with 131 confirmed cases, a decreasing trend was maintained with less than 100 confirmed cases per day. However, it cannot be said that the rapid increase in the number of confirmed cases during this period reflects the actual rate of COVID-19 spread in Daegu. Because of the intensive large-scale screening of the members of Shincheonji, it is likely that the rate increased more steeply than the actual rate of spread because more confirmed cases arose in a short period of time compared with the previous order and rate of infection. Since data on the date of symptom onset in patients in Daegu are not available, the results of this study using the cumulative data may be different from the actual COVID-19 transmission patterns in Daegu. If the data on the date of symptom onset are collected in the future, a follow-up study using this information should be conducted. In addition, because we did not assume any other additional group infections, such as in nursing homes, other than Shincheonji, this study may show a different pattern from the actual transmission of COVID-19.

The above results assume that newly infected COVID-19 cases in Daegu did not come from abroad or other regions in Korea. For a more accurate prediction and analysis of the effect of quarantine policies, such simulations should be expanded to reflect the whole country and consider the entry of latent patients from other regions and abroad. Studies on this are currently under way.

SUPPLEMENTARY MATERIALS

Korean version is available at http://www.e-epih.org/.

CONFLICT OF INTEREST

The authors have no conflicts of interest to declare for this study.

FUNDING

This project was supported by the Government-wide R&D Fund Project for Infectious Disease Research (GFID), Republic of Korea (Grant No. HG18C0088).

ACKNOWLEDGEMENTS

We thank all individuals who are struggling in healthcare fields to overcome the COVID-19 outbreak. This study was performed under the research project named ‘Research and Development on Integrated Surveillance System for Early Warning of Infectious Diseases (RISEWIDs).’ The investigators of this project were Jaeyong Kim (Yonsei University), Sunwool Kim (National Medical Center), Eui Jung Kwon (National Health Insurance Review and Assessment Service), Dong Wook Kim (National Health Insurance Service), Moran Ki (National Cancer Center), Hyunjin Son (Pusan National University Hospital), Jong-Hun Kim (Sungkyunkwan University), Jin Yong Kim (Incheon Medical Center), Heeyoung Lee (Seoul National University Bundang Hospital), Boyoung Park (Hanyang University), Woo-Sik Son (National Institute for Mathematical Sciences), Jungsoo Lee (University), Sunhwa Choi (National Cancer Center), Okyu Kwon (National Institute for Mathematical Sciences), Hyojung Lee (National Institute for Mathematical Sciences), Jong-Hoon Kim (International Vaccine Institute), Heecheon Kim (MISO Info Tech), and Bo Youl Choi (Hanyang University).

AUTHOR CONTRIBUTIONS

All work was done by WSS and RISEWIDs Team.

ORCID

Woo-Sik Son: https://orcid.org/0000-0002-0421-4546

REFERENCES

1. World Health Organization. Naming the coronavirus disease (COVID-19) and the virus that causes it [cited 2020 Apr 1]. Available from: https://www.who.int/emergencies/diseases/novel-coronavirus-2019/technical-guidance/naming-the-coronavirus-disease-(covid-2019)-and-the-virus-that-causes-it.
2. World Health Organization. WHO timeline-COVID-19; 2020 Apr 27 [cited 2020 Jun 20]. Available from: https://www.who.int/news-room/detail/27-04-2020-who-timeline---covid-19.
3. Brauer F, van den Driessche P, Wu J. Mathematical epidemiology. Berlin: Springer; 2008, p. 19-79.
4. Dowdy DW, Golub JE, Chaixson RE, Saraceni V. Heterogeneity in tuberculosis transmission and the role of geographic hotspots in propagating epidemics. Proc Natl Acad Sci U S A 2012;109:9557-9562.
5. Ferguson NM, Cummings DA, Cauchemez S, Fraser C, Riley S, Meeyai A, et al. Strategies for containing an emerging influenza pandemic in Southeast Asia. Nature 2005;437:209-214.
6. MicroData Integrated Service. Data in service [cited 2020 Jul 20]. Available from: https://mdis.kostat.go.kr/infoData/detailData.do?statsConfmNo=101001%20(2020) (Korean).
7. Li Q, Guan X, Wu P, Wang X, Zhou L, Tong Y, et al. Early transmission dynamics in Wuhan, China, of novel coronavirus-infected pneumonia. N Engl J Med 2020;382:1199-1207.
8. Ki M; Task Force for 2019-nCoV. Epidemiologic characteristics of early cases with 2019 novel coronavirus (2019-nCoV) disease in Korea. Epidemiol Health 2020;42:e2020007.