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Epidemiological characteristics of sporadic nosocomial COVID-19 infections from June 2020 to June 2021 in China: An overview of vaccine breakthrough infection events

Zhigang He¹, Hong-bing Xiang¹, Anne Manyande², Weiguo Xu¹, Li Fan³, Boqi Xiang^{4*}

¹Tongji Hospital, Tongji Medical College, Huazhong University of Science and Technology, China, ²University of West London, United Kingdom, ³Department of Orthopedics, Union Hospital of Tongji Medical College, Huazhong University of Science and Technology, China, ⁴Rutgers University, United States

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Author contribution statement

All the authors contributed to the paper presented methodology and conceptualization. They all contributed to data analysis and paper writing.

Keywords

COVID-19 disease, Asymptomatic carrier, Nosocomial infection, Scientific protective strategy, vaccine breakthrough infection

Abstract

Word count: 108

The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) pandemic has wreaked havoc to human beings around the world. Although China quickly brought the Coronavirus disease (COVID-19) pandemic under control, there have been several sporadic outbreaks in different regions of China since then. This article describes the chronological nosocomial COVID-19 infection events that related to several sporadic outbreaks of SARS-CoV-2 in different regions of China. We reported epidemiological characteristics and management measures of sporadic nosocomial COVID-19 infections from June 2020 to June 2021 in China, specially focused on domestic COVID-19 breakthrough infection in China — a vaccinated healthcare professional working in the isolation ward of a designated COVID-19 hospital.

Contribution to the field

The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) pandemic has wreaked havoc to human beings around the world. Although China quickly brought the Coronavirus disease (COVID-19) pandemic under control, there have been several sporadic outbreaks in different regions of China since then. This article describes the chronological nosocomial COVID-19 infection events that related to several sporadic outbreaks of SARS-CoV-2 in different regions of China. We reported epidemiological characteristics and management measures of sporadic nosocomial COVID-19 infections from June 2020 to June 2021 in China, specially focused on domestic COVID-19 breakthrough infection in China — a vaccinated healthcare professional working in the isolation ward of a designated COVID-19 hospital.

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Inclusion of identifiable human data

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| 19 | Abstract |
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| 20 | The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) pandemic has wreaked |
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| 22 | disease (COVID-19) under control, there have been several sporadic outbreaks in different regions |
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| 24 | infection events related to several sporadic outbreaks of SARS-CoV-2 in different regions of |
| 25 | China. We report epidemiological characteristics and management measures of sporadic |
| 26 | nosocomial COVID-19 infections from June 2020 to June 2021, and specially focused on the |
| 27 | domestic COVID-19 breakthrough infection in China including domestic COVID-19 |
| 28 | breakthrough infection - a vaccinated healthcare professional working in the isolation ward of a |
| 29 | designated COVID-19 hospital. |
| 30 | Keywords: COVID-19 disease; asymptomatic carrier; nosocomial infection; Scientific protective |
| 31 | strategy; vaccine breakthrough infection |
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Introduction

The outbreak of the COVID-19 virus was first reported in Wuhan, China, in December 2019. The COVID-19 pandemic led China to quarantine the population in order to protect them ¹. Chinese authorities decided to adopt extraordinary measures to contain and limit the spread of the SARS-CoV-2 virus. From 1 January to 8 April 2020, >8,000 patients with COVID-19 were hospitalized and the Chinese government imposed the Wuhan lockdown on January 23, which ended on April 8, 2020.

With the COVID-19 epidemic quickly under control in the early stages of 2020, importing the SARS-CoV-2 virus could pose great challenges to the control and prevention of nosocomial COVID-19 infection in healthcare settings. Our previous report showed the impact of the novel coronavirus SARS-CoV-2 among healthcare workers in hospitals during the early phase of the COVID-19 epidemic², and suggested that local authorities need to be extremely cautious and implement stringent protective measures to safeguard healthcare workers in order to counteract the threats brought by the pandemic³⁻⁵. Though medical staff belong to the susceptible population to a certain extent, hospitals have practiced cohorting in accordance with recommendations from COVID-19 infection prevention and control professional societies, which has reduced the risk of hospital-acquired COVID-19. Because healthcare workers are at the interface between hospitals and the community, where there is significant COVID-19 transmission, they may be infected by asymptomatic carriers with COVID-19 or COVID-19 patients. Furthermore, in the designated hospital admitting COVID-19 patients or asymptomatic carriers, medical staff are highly exposed to nosocomial COVID-19 acquisition and SARS-CoV-2 transmission. Therefore, healthcare staff may play a key role in initiating or amplifying sporadic COVID-19 outbreaks in healthcare settings including hospitals and other care facilities.

The importing of the SARS-CoV-2 virus from overseas, induced sporadic outbreaks with COVID-19 from June 2020 to June 2021 in China^{6, 7}. This article describes the chronological events that led to several sporadic nosocomial COVID-19 infections in different regions of China from June 2020 to June 2021, including domestic COVID-19 breakthrough infection — a vaccinated healthcare professional working in the isolation ward of a designated COVID-19 hospital⁸. Lastly, we provide an overview of local COVID-19 outbreak induced by the B.1.617.2 (Delta) variant of the COVID-19 virus in China.

Methods

From the beginning of June 1, 2020, every day, we prospectively focused on the COVID-19 epidemic data from the Chinese Center for Disease Control and Prevention. Once we had the new report of nosocomial COVID-19 infection in China, we tracked this epidemic and collected its epidemiological characteristics from announcements by the local Health Commission and presented a narrative research for geographical and epidemiological characteristics of nosocomial COVID-19 infection from June 2020 to June 2021.

We searched epidemiologic data published on the website of WHO, the China Center for Disease Control and Prevention, National Health Commission, the Health Commission of Qingdao, Shenyang, Shijiazhuang, and Dalian city, Jilin, Shaanxi, and Guangdong Province from June 2020

- 77 to January 2021. Using the keywords "nosocomial infection", "COVID-19 variant", "SARS-CoV-
- 78 2", "B1.617.2 (Delta) lineage", and Boolean operator 'AND', we periodically searched the
- published medical literature using the PubMed service maintained by the U.S. National Library of
- 80 Medicine of NIH. Confirmed COVID-19 cases are defined as persons who tested positive for SARS-
- 81 CoV-2 and had clinical symptoms. Asymptomatic carriers refer to persons without clinical
- symptoms who tested positive for SARS-CoV-2.

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Results

Regional distribution of nosocomial COVID-19 infections from June 2020 to June 2021

- From June 2020 to June 2021, regional distribution of the sporadic nosocomial COVID-19
- infections is shown in Figure 1, 2 and Table 1. Most of these cities are located in coastal or airline
- 88 hub areas, for example, Dalian, Qingdao, Shanghai are coastal cities, and Xi'an, Shenyang,
- 89 Shijiazhuang, and Guangzhou are airline hub cities.

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Occupational distribution of nosocomial COVID-19 infections from June 2020 to June 2021

- 92 From June 2020 to June 2021, six cities in China had reported over 45 cases with nosocomial
- 93 COVID-19 infections, including 39 confirmed cases and 6 asymptomatic cases (Table 2). Among
- 94 them, nursing staff (3), doctors (6), patients (12), accompanying staff (22), and laboratorian (2) were
- 95 diagnosed. The route of these nosocomial infections mainly included workplace accidental exposure
- 96 to COVID-19, cross infection among healthcare workers, patients, and accompanying staff.

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SARS-CoV-2 gene sequencing results of nosocomial COVID-19 infections from June 2020 to

- 99 **June 2021**
- 100 SARS-CoV-2 gene sequencing is crucial work for nosocomial COVID-19 infections in new
- 101 sporadic outbreak regions. The virus strain of nosocomial COVID-19 infection found in Qingdao
- 102 city is the virus strain lineage B.1.1 imported from overseas (Table 2). Gene sequencing results
- showed that the coronavirus found in Dalian, Shenyang, and Shijiazhuang cities is similar to the
- strain imported from Europe; The virus strain found in Xi'an city is a COVID-19 variant B.1.1.7
- lineage; SARS-CoV-2 strain found in Guangzhou city is the imported virus strain B1.617.2 (Delta)
- lineage (Table 2).
- A report from Fang et al⁹ showed that the strains associated with specific outbreak in Dalian City
- 108 were as follows: LNDL-BHQ-0722-Y S12 L001 R1 001, LNDL-SFL-0722-
- 109 Y S9 L001 R1 001, LNDL-WY-0722-Y S11 L001 R1 001, and LNDL-XY-0722-
- 110 Y S10 L001 R1 001; The parent strain from Wuhan was
- 111 NC 045512.2 Severe acute respiratory syndrome coronavirus 2 isolate Wuhan-Hu-
- 112 1 complete genome. After accessing the public database GISAID and GenBank, 3 Russian strains
- detected in July were found to share the 10 variation sites with the 2 Hebei strains (GISAID IDs:
- EPI ISL 596266, EPI ISL 569792, and EPI ISL 569793)¹⁰. Evidence indicates that these the
- Shijiazhuang strains may have originated from these Russian strains¹⁰.

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Discussion

- 118 This paper provides us with an inspiring vision regarding the current COVID-19 pandemic. The
- main findings of sporadic nosocomial COVID-19 infections from June 2020 to June 2021 are listed
- below: (1) the importance of rolling out an overall nucleic acid test campaign to all staff in healthcare

settings, is a crucial part of the COVID-19 surveillance; (2) The case of coronavirus transported onto the workplace warns us that the current hospital disinfection concept urgently requires updating, especially in designated hospitals, and there is an urgent demand of the management of staff accompanying patients in non-designated hospitals; (3) The phenomenon about COVID-19 vaccine breakthrough infection deserves our great attention.

For healthcare workers, patients/accompanying staff, and cleaner, their COVID-19 infection may be from hospital (hospital-acquired infection, also nosocomial infection) or community (community-acquired infection). Distinguishing nosocomial infection and community-acquired infection is an important base to control the epidemic in nosocomial COVID-19 infections. Because healthcare workers are at the interface between hospitals and the community, where there is significant COVID-19 transmission, they may be infected by asymptomatic carriers from the community with COVID-19 or COVID-19 patients. These healthcare workers (infected in the community) do not belong to nosocomial infection (also called hospital-acquired infection). For example, in Figure 1, the infected cases (including healthcare workers, cleaner, etc.) in Beijing, Dalian (27 Dec. 2020) and Shanghai were community-acquired infection, not nosocomial infection. For nosocomial infection of healthcare workers, patients/accompanying staff, and cleaner, as the effective control measures taken by the local government must include: (1) these hospitals imposed lockdown, especially in the designated COVID-19 hospital; (2) rolling out an overall nucleic acid test campaign to all staff is key in healthcare settings.

Among 9 sporadic nosocomial COVID-19 infections, our results showed that 8 series cases were found by rolling out overall a nucleic acid test campaign to all hospital staff, suggesting the importance of regular screening for all staff in healthcare settings. As part of efforts to control the COVID-19 infections, one of the effective prevention measures taken by local authorities is that all hospital personnel labeled a key population are closely monitored. Whether the epidemic in nosocomial COVID-19 infections can be brought under control depends on how many new infections emerge of their close contacts and secondary close contacts in the next two weeks. If these hospitals imposed lockdown as the effective control measures taken by the local government, then ideally, new confirmed cases with COVID-19 infections will see a downward trend within two weeks. Cases such as hospital-acquired infection in the designated hospital of Qingdao city are a warning that workplace storage and transport could be a hotbed for the coronavirus, or other pathogens. There is an urgent demand for workplace disinfection to protect the health safety of medical staff, patients, and accompanying staff in the current pandemic or into another worse outbreak.

Our data also showed that a vaccinated healthcare professional who received inactivated vaccine was infected with COVID-19 while working in the isolation ward of a designated COVID-19 hospital, and the coronavirus strain was determined to be the imported COVID-19 variant strain B.1.1.78, suggesting that there exists the domestic COVID-19 vaccine breakthrough infection in China, and this phenomenon deserves our serious attention. Vaccination is well-known to be key to stopping the virus from circulating and more variants from popping up¹¹⁻¹³. The vaccine breakthrough infection case was defined as an individual with positive SARS-CoV-2 nucleic acid amplification tested after receiving at least one dose of a SARS-CoV-2 vaccine¹⁴⁻¹⁶. Jacobson et al¹⁷ addressed post-vaccination SARS-CoV-2 infections and the incidence of the B.1.427/B.1.429

variant among healthcare personnel at a northern California academic medical center. Hacisuleyman et al¹⁸ reported two women with vaccine breakthrough infection in a cohort of 417 persons who had received the second dose of BNT162b2 (Pfizer–BioNTech) or mRNA-1273 (Moderna) vaccine at least two weeks previously, and the viral sequencing showed that they were infected with the new variant virus including E484K in 1 woman and three mutations (T95I, del142–144, and D614G) in both. These observations revealed a potential risk of COVID infection with the variant virus after successful vaccination.

Vaccine breakthrough cases with SARS-CoV-2 were reported in many countries¹⁹⁻²¹. An analysis from Israel's vaccination campaign showed that COVID-19-related hospitalizations, severe disease, and death were reduced in infected cases with SARS-CoV-2 after vaccination, including symptomatic and asymptomatic infections²¹. Antonelli et al²² identified risk factors for post-vaccination SARS-CoV-2 infection and describe the characteristics of post-vaccination illness, and found that Almost all symptoms were reported less frequently in infected vaccinated individuals than in infected unvaccinated individuals, and vaccinated participants were more likely to be completely asymptomatic, especially if they were 60 years or older. Our vaccine breakthrough case from Xi'an supported vaccine effectiveness and cautioned around relaxing physical distancing and other personal protective measures in the post-vaccination era.

SARS-CoV-2 virus delta variants have drawn worldwide attention. The WHO proposed three labels for global SARS-CoV-2 variants, including variant of concern (VOC), variant of interest (VOI), and variant under monitoring, to be used alongside the scientific nomenclature in communications about variants to the public²³. The B.1.617 variant of the COVID-19 virus has been called a triple mutant variant since it splits into three lineages including the B.1.617.1 (Kappa) variant, the B.1.617.2 (Delta) variant, and the B.1.617.3 variant²⁴. The delta COVID-19 variant, which was first detected in India in October 2020, had been reported in more than 80 countries on June 20, 2021²⁵. The WHO declared the delta variant a "variant of concern" on May 10, 2021. In Mid-June 2021, the US Centers for Disease Control and Prevention upgraded its classification of the delta from a "variant of interest" to a "variant of concern."²⁶ Our results showed that two medical staff (1 laboratorian and 1 emergency doctor), infected with COVID-19 in the designated hospital, occurred due to workplace accidental exposure to COVID-19, and the coronavirus strain was determined to be the imported B.1.617.2 (Delta) variant.

To conclude, our findings add to the accumulating evidence regarding the importance of regular screening for all staff in healthcare settings. Furthermore, our study highlights the need for an update of the current hospital disinfection procedures in designated hospitals to prevent the nosocomial spread of SARS-CoV-2 infection. Finally, the epidemiological exposure of vaccinated medical staff should draw concern in order to minimize the impact of a new outbreak induced by virus mutants.

Table 1: Epidemiological characteristics of sporadic nosocomial COVID-19 infections

| Date | City | Number of | Hospital type | Profession | Classification of | Route of infection | Database |
|---------|---------|-----------------|---------------|------------|-------------------|--------------------|----------|
| | | hospital/ cases | | | infection | | |
| 19 Jun. | Beijing | 1/1 | Designated | nurse | Community- | family member | Link 1 |
| 2020 | | | hospital | | acquired | exposure with | |
| | | | | | infection | COVID-19 | |

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Designated hospital: a hospital designated for treatment of imported COVID-19 cases

Link 1: https://finance.sina.com.cn/china/gncj/2020-06-19/doc-iircuyvi9385332.shtml?r=9&tj=none&tr=9

Link 2: https://www.zhihu.com/question/425220159

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208 Link 3: https://new.qq.com/omn/20201030/20201030A05GXU00.html

209 Link 4: http://www.xinhuanet.com/politics/2021-01/06/c_1126953634.htm

| 210 | Link 5: https://new.gg.com/g | omn/20201229/20201229A0259M00.html |
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Link 6: https://new.qq.com/omn/20210111/20210111A0DFD500.html

Link 7: https://news.sina.com.cn/c/2021-01-20/doc-ikftpnnx9631142.shtml

 $213 \qquad \text{Link 8: https://new.qq.com/omn/20210131/20210131A0071200.html} \\$

214 Link 9: https://news.sina.com.cn/c/2021-03-18/doc-ikkntiam4831560.shtml

Link 10: https://news.sina.com.cn/c/2021-06-14/doc-ikqcfnca0979774.shtml

Link 11: http://med.china.com.cn/content/pid/281087/tid/1026

Link 12: http://news.hexun.com/2021-08-18/204179459.html

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Table 2: Epidemiological characteristics of sporadic nosocomial COVID-19 infections

| Date | City | Confirmed/ asymptomatic cases | How to discover the source of infection | Virus type | Database |
|-----------------|--------------|-------------------------------------|--|--|----------|
| 19 Jun. 2020 | Beijing | 1/0 | Nucleic acid testing for close contacts of people with infection cases | One ancestral virus strain (XFDM strain) ²⁷ | Link 1 |
| 11 Oct. 2020 | Qingdao | 6/6 | Regular screening for medical staff | Imported virus strain Lineage B.1.1 ²⁸ | Link 2 |
| 29 Oct. 2020 | Qingdao | 1/0 | Regular screening for medical staff | Imported virus strain Lineage B.1.1 ²⁸ | Link 3 |
| 18 Dec. 2020 | Dalian | 3/0 | Regular screening for medical staff | European family branch 1 of the L genotype | Link 4 |
| 27 Dec. 2020 | Dalian | 1/0 | Regular screening for medical staff | European family branch 1 of the L genotype | Link 5 |
| 3 Jan. 2021 | Shenyang | 12/0 | Regular screening for medical staff | Imported virus strain | Link 6 |
| 14 Jan. 2021 | Shijiazhuang | 14/0 | Regular screening for medical staff | the strain imported from Europe ²⁹ | Link 7 |
| 20 Jan. 2021 | Shanghai | 2/0 | Nucleic acid testing for close contacts of people with infection cases | Imported virus strain | Link 8 |
| 17 Mar. 2021 | Xi'an | 1/0 | Regular screening for medical staff | Imported virus strain B.1.1.78 | Link 9 |
| 14 Jun. 2021 | Guangzhou | 2/0 | Regular screening for medical staff | Imported virus strain B1.617.2 (Delta) Lineage | Link 10 |
| 27 Jul. 2021 | Nanjing | 4/0 | Regular screening for medical staff | Imported virus strain B1.617.2 (Delta) Lineage | Link 11 |
| 10Aug. 2021 | Yangzhou | 3/0 | Regular screening for medical staff | Imported virus strain B1.617.2 (Delta) Lineage | Link 12 |

Delta Lineage: the highly contagious Covid-19 variant first identified in India

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Link 1: http://weekly.chinacdc.cn/en/article/doi/10.46234/ccdcw2020.246

 $224 \qquad \text{Link 2: http://weekly.chinacdc.cn/en/article/doi/10.46234/ccdcw2020.224}$

- 225 Link 3: https://new.qq.com/omn/20201030/20201030A05GXU00.html
- 226 Link 4: http://www.xinhuanet.com/politics/2021-01/06/c 1126953634.htm
- 227 Link 5: https://new.qq.com/omn/20201229/20201229A0259M00.html
- 228 Link 6: https://new.qq.com/omn/20210111/20210111A0DFD500.html
- Link 7: http://weekly.chinacdc.cn/en/article/doi/10.46234/ccdcw2021.006
- 230 Link 8: https://new.qq.com/omn/20210131/20210131A0071200.html
- Link 9: http://weekly.chinacdc.cn/en/article/doi/10.46234/ccdcw2021.094
- 232 Link 10: https://news.sina.com.cn/c/2021-06-14/doc-ikqcfnca0979774.shtml
- Link 11: http://med.china.com.cn/content/pid/281087/tid/1026
- 234 Link 12: http://news.hexun.com/2021-08-18/204179459.html

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Authors contribution

All the authors contributed to the paper presented methodology and conceptualization. They all contributed to data analysis and paper writing.

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Declaration of competing interest

- 245 The authors declare that they have no known competing financial interests or personal relationships
- 246 that could have appeared to influence the work reported in this paper

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Figure 1: Geographical distribution of sporadic nosocomial COVID-19 infections from June 2020 to June 2021 in China. Seven cities (Dalian, Beijing, Qingdao, Shijiazhuang, Xi'an, Guangzhou, and Shanghai) reported the nosocomial COVID-19 infections. Medical staff, patients and accompanying staff (12) in Shenyang (Jan 3, 2021), patients (3) in Dalian (Dec 18, 2020), patients and accompanying staff (6) in Qingdao (Oct 11, 2020), laboratory personnel (1) in Xi'an (Mar 17, 2021), medical staff, patients and accompanying staff (14) in Shijiazhuang (Jan 14, 2021), and Medical staff (2) in Guangzhou (Jun 14, 2021) were diagnosed as infected cases.

Figure 2: The graph's left-right axis is used as a timeline of the key events and dynamic profile of sporadic nosocomial COVID-19 infections from June 2020 to June 2021 in China.

- From June 2020 to June 2021, five cities in China had reported 10 series cases with nosocomial COVID-19 infections, including 39 confirmed cases and 6 asymptomatic cases.
- The delta COVID-19 variant was first detected in India in October 2020. The World Health Organization designated Delta as a variant of interest in April and a variant of concern on 11 May
- 2021. In Mid-June 2021, the US Centers for Disease Control and Prevention upgraded its classification of delta from a "variant of interest" to a "variant of concern."



