

1 **Epidemiological characteristics and transmission dynamics of the outbreak**  
2 **caused by the SARS-CoV-2 Omicron variant in Shanghai, China: a descriptive**  
3 **study**

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32 Abstract: 370 words

33 Main text: 3,992 words

## 34 **Summary**

### 35 **Background**

36 In early March 2022, a major outbreak of the severe acute respiratory syndrome  
37 coronavirus 2 (SARS-CoV-2) Omicron variant spread rapidly throughout Shanghai,  
38 China. Here we aimed to provide a description of the epidemiological characteristics  
39 and spatiotemporal transmission dynamics of the Omicron outbreak under the  
40 population-based screening and lockdown policies implemented in Shanghai.

41

### 42 **Methods**

43 We extracted individual information on SARS-CoV-2 infections reported between  
44 January 1 and May 31, 2022, and on the timeline of the adopted non-pharmacological  
45 interventions. The epidemic was divided into three phases: i) sporadic infections  
46 (January 1–February 28), ii) local transmission (March 1–March 31), and iii) city-wide  
47 lockdown (April 1 to May 31). We described the epidemic spread during these three  
48 phases and the subdistrict-level spatiotemporal distribution of the infections. To  
49 evaluate the impact on the transmission of SARS-CoV-2 of the adopted targeted  
50 interventions in Phase 2 and city-wide lockdown in Phase 3, we estimated the dynamics  
51 of the net reproduction number ( $R_t$ ).

52

### 53 **Findings**

54 A surge in imported infections in Phase 1 triggered cryptic local transmission of the  
55 Omicron variant in early March, resulting in the largest coronavirus disease 2019  
56 (COVID-19) outbreak in mainland China since the original wave. A total of 626,000  
57 SARS-CoV-2 infections were reported in 99.5% (215/216) of the subdistricts of  
58 Shanghai. The spatial distribution of the infections was highly heterogeneous, with 40%  
59 of the subdistricts accounting for 80% of all infections. A clear trend from the city  
60 center towards adjacent suburban and rural areas was observed, with a progressive  
61 slowdown of the epidemic spread (from 544 to 325 meters/day) prior to the citywide  
62 lockdown. During Phase 2,  $R_t$  remained well above 1 despite the implementation of  
63 multiple targeted interventions. The citywide lockdown imposed on April 1 led to a  
64 marked decrease in transmission, bringing  $R_t$  below the epidemic threshold in the entire  
65 city on April 14 and ultimately leading to containment of the outbreak.

66

### 67 **Interpretation**

68 Our results highlight the risk of widespread outbreaks in mainland China, particularly  
69 under the heightened pressure of imported infections. The targeted interventions  
70 adopted in March 2022 were not capable of halting transmission, and the  
71 implementation of a strict, prolonged city-wide lockdown was needed to successfully

72 contain the outbreak, highlighting the challenges for successfully containing Omicron  
73 outbreaks.

74

#### 75 **Funding**

76 Key Program of the National Natural Science Foundation of China (82130093).

77

#### 78 **Research in context**

##### 79 **Evidence before this study**

80 On May 24, 2022, we searched PubMed and Europe PMC for papers published or  
81 posted on preprint servers after January 1, 2022, using the following query:  
82 (“SARS-CoV-2” OR “Omicron” OR “BA.2”) AND (“epidemiology” OR  
83 “epidemiological” OR “transmission dynamics”) AND (“Shanghai”). A total of 26  
84 studies were identified; among them, two aimed to describe or project the spread of the  
85 2022 Omicron outbreak in Shanghai. One preprint described the epidemiological and  
86 clinical characteristics of 376 pediatric SARS-CoV-2 infections in March 2022, and the  
87 other preprint projected the epidemic progress in Shanghai, without providing an  
88 analysis of field data. In sum, none of these studies provided a comprehensive  
89 description of the epidemiological characteristics and spatiotemporal transmission  
90 dynamics of the outbreak.

91

##### 92 **Added value of this study**

93 We collected individual information on SARS-CoV-2 infection and the timeline of the  
94 public health response. Population-based screenings were repeatedly implemented  
95 during the outbreak, which allowed us to investigate the spatiotemporal spread of the  
96 Omicron BA.2 variant as well as the impact of the implemented interventions, all  
97 without enduring significant amounts of underreporting from surveillance systems, as  
98 experienced in other areas. This study provides the first comprehensive assessment of  
99 the Omicron outbreak in Shanghai, China.

100

##### 101 **Implications of all the available evidence**

102 This descriptive study provides a comprehensive understanding of the epidemiological  
103 features and transmission dynamics of the Omicron outbreak in Shanghai, China. The  
104 empirical evidence from Shanghai, which was ultimately able to curtail the outbreak,  
105 provides invaluable information to policymakers on the impact of the containment  
106 strategies adopted by the Shanghai public health officials to prepare for potential  
107 outbreaks caused by Omicron or novel variants.

## 108 **Background**

109 The first wave of coronavirus disease 2019 (COVID-19) in China subsided quickly  
110 with strict containment measures in March 2020.<sup>1</sup> As numerous variants have emerged  
111 across the globe, China successfully contained multiple COVID-19 outbreaks by  
112 adhering to a containment policy. This policy aimed to curb flare-ups of local  
113 transmissions in the shortest possible time, which is done by relying on a set of  
114 non-pharmacological interventions (NPIs) with adjusted intensities according to the  
115 situation on the ground.<sup>2,3</sup>

116

117 In November 2021, the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2)  
118 Omicron variant was first reported in South Africa.<sup>4</sup> Its high transmissibility and  
119 immune escape properties enabled it to rapidly replace previous strains and become  
120 dominant globally.<sup>5,6</sup> Although the containment policy implemented in China had been  
121 effective in containing pre-Omicron outbreaks, its effectiveness against Omicron was  
122 unclear. In fact, a total of 750,000 SARS-CoV-2 infections were reported in mainland  
123 China in the first five months of 2022,<sup>7</sup> the majority in Shanghai.

124

125 Shanghai is one of the most populous and economically advanced metropolises in  
126 China, with a population of nearly 25 million<sup>8</sup> and unique features compared to other  
127 areas in mainland China and abroad. The population in Shanghai was mostly  
128 vaccinated with domestically-developed inactivated vaccines, which had relatively  
129 lower effectiveness in preventing SARS-CoV-2 infections due to their lower  
130 antibody-neutralizing responses compared to mRNA vaccines<sup>9,10</sup>. In addition, the  
131 vaccination coverage was highest in the young population (100.0% in individuals aged  
132 between 12 and 17 years had completed the primary vaccine schedule as of March  
133 2022)<sup>11</sup> and lowest in the elderly (only 62.0% in individuals aged 60 years or older had  
134 completed the primary schedule as of April 2022).<sup>12</sup> Compared to the rest of the cities  
135 in mainland China, Shanghai had the highest risk of SARS-CoV-2 infection  
136 importation due to its global connectivity: about 30-40% of all international flights  
137 arriving in China since 2020 landed in Shanghai.<sup>13</sup> Although a quarantine period of two  
138 weeks was mandatory for all incoming travelers, lapses in these measures may have led  
139 to the repeated seeding of SARS-CoV-2 into the community.<sup>14,15</sup> Consequently, in  
140 early March 2022, a major outbreak of the Omicron BA.2 variant started to spread in  
141 Shanghai.<sup>16</sup>

142

143 In this study, we provide a detailed description of Shanghai's outbreak response  
144 timeline, the spatiotemporal distribution of COVID-19 cases, and Omicron  
145 transmission dynamics since 2022. The analysis of the Shanghai 2022 Omicron

146 outbreak could provide invaluable information to policymakers on the impact of the  
147 containment strategies adopted by Shanghai to prepare for potential outbreaks caused  
148 by Omicron or future novel variants.  
149

## 150 **Methods**

### 151 **Definition of infection and case**

152 An asymptomatic infection is defined as a PCR-confirmed individual who i) does not  
153 meet any of the following clinical criteria: fever, cough, sore throat, and other  
154 self-perceived and clinical-identifiable symptoms or signs; and ii) has no radiographic  
155 evidence of pneumonia. Laboratory-confirmed cases are categorized into four types  
156 based on clinical severity: mild, moderate, severe, and critical cases, with detailed  
157 definitions in the **Supplementary Information**. Detailed surveillance, detection, and  
158 management of infections and cases are also presented in the **Supplementary**  
159 **Information**.

### 161 **Public health response**

162 We summarize the NPIs implemented in Shanghai before and during the Omicron  
163 waves. Prior to the initial surge in Omicron, Shanghai maintained a baseline level of  
164 intensive NPIs against potential outbreaks, including stringent border control policies,  
165 symptom-based surveillance, case isolation, tracing of close contacts (requires  
166 quarantine in separate facilities) and contacts of contacts, occupation-based screening,  
167 targeted screening of individuals at high risk of infection (e.g., contacts of contacts),  
168 and a set of other social distancing measures such as travel restrictions and community  
169 confinement (**Figure 1**). After the Omicron variant was introduced in Shanghai, a set of  
170 additional NPIs were implemented to halt its transmission. In terms of border control  
171 measures, the point of entry for international flights to Shanghai was diverted to other  
172 airports. At the local scale, the closure of in-person educational activities at all levels  
173 was implemented from March 12 to 15, 2022 (starting from primary and secondary  
174 school, then followed by universities). From March 16 to March 27, Shanghai adopted  
175 grid management at the subdistrict level, fractioning subdistricts into high-risk areas.  
176 The lists of high-risk areas changed dynamically over time based on the following  
177 principles: the epidemiological situation measured in terms of number of cases and  
178 infections, gathering of crowds, population density, social characteristics, and  
179 economic activity.<sup>17</sup> During this period, several rounds of nucleic acid screening were  
180 performed within high-risk and non-high-risk areas. In high-risk areas, either one or  
181 two rounds of population-based nucleic acid screening were performed within 48 hours,  
182 together with a lockdown. In non-high-risk areas, a single round of mass PCR screening

183 was performed between March 18 and 20. Afterwards, the entire city entered a phased  
184 stage of lockdown, where eastern Shanghai (which comprises areas east of the  
185 Huangpu River, **Supplementary Figure 1**) entered a population-wide lockdown on  
186 March 28, and then the rest of Shanghai entered the lockdown phase on April 1. The  
187 lockdown restriction was lifted on June 1 when the daily number of infections first  
188 declined to 10..

189

## 190 **Data sources and collection**

191 Aggregated data on the number of local and imported infections according to  
192 symptomatic status in Shanghai from December 2019 were gathered from the Shanghai  
193 Municipal Health Commission through a combined approach of manual downloads and  
194 compiled scripts. Individual line lists of all SARS-CoV-2 infections were retrieved  
195 from multiple publicly available official data sources (websites of municipal health  
196 commissions) and integrated with the supplementary information gathered from the  
197 websites of local government media (**Supplementary Table 1**). The resulting line list  
198 contains the following variables: residential location (address, district, and subdistrict),  
199 symptomatic status (including the clinical outcomes of initially asymptomatic  
200 infections), date of official reporting, and means of infection identification (i.e., routine  
201 screening of community groups or general screening of the quarantined population)  
202 (**Supplementary Table 2**). Specifically, community screening groups referred to those  
203 residents living in areas reporting no new infections, people proactively looking for  
204 healthcare or nucleic acid tests, and essential workers. The quarantined population  
205 corresponded to individuals who were quarantined at home or in designated facilities,  
206 lived in areas reporting new infections or other key settings under close management,  
207 and were identified through screening of close contacts<sup>18</sup>. We obtained the timeline of  
208 the adopted NPI strategies from the official announcements of the authorities of  
209 Shanghai through search engine queries; the list of high-risk areas during March 16 and  
210 March 27 was found by searching on local media pages (**Supplementary Table 3**).  
211 More details on data collection are provided in the **Supplementary Information**.

212

## 213 **Statistical analyses**

214 Definition of the three phases of the epidemic

215 Three phases of the epidemic were defined according to the staged NPIs policies  
216 adopted and the epidemic situation. The first phase covers the period from January 1 to  
217 February 28, 2022, during which the main risk was posed by the importation of  
218 infections and only small numbers of sporadic and locally transmitted cases were  
219 recorded. The second phase started on March 1, immediately before the outbreak of the  
220 BA.2 Omicron variant was confirmed<sup>16</sup>. In addition, no epidemiological link between

221 infections reported before and after March 1, 2022, was identified by the authorities.  
222 The third phase covers the period from April 1 to May 31, 2022, when the entire city  
223 went into staged lockdowns and the epidemic started to subside.

224

#### 225 Inference of delay between sampling and reporting

226 The date of symptom onset or infection was needed to describe the transmission  
227 dynamics, whereas we only had the date of official reporting for each infection. We  
228 estimated the delayed days between the sampling dates and reporting dates and then  
229 generated the sampling date for each infection by assuming that the sampling dates  
230 were close to the dates of symptom onset or infection time.

231

232 The mean reporting delay was determined by computing the cross-correlation  
233 coefficients between the cumulative number of reported SARS-CoV-2 infections and  
234 the cumulative number of sampled specimens under different lengths of lag  
235 (**Supplementary Figure 2**), the details of which are presented in the **Supplementary**  
236 **Information**. We applied delays of 2 days, 3 days, and 2 days for the periods before  
237 March 15, between March 16 and May 14, and after May 15, respectively. In the  
238 sensitivity analysis, we considered a four-day delay for the period from March 16 to  
239 May 14.

240

241 In addition to reporting delay, we also estimated individual locations and the  
242 progression of pre-symptomatic infections for data analysis (see **Supplementary**  
243 **Information** for details).

244

#### 245 Spatial trends

246 Trend surface analysis was used to explore the SARS-CoV-2 spread in Shanghai, China.  
247 Thin-plate spline regression was used to interpolate the first date of sampling in each 3  
248 km × 3 km cell of a gridded population of Shanghai to draw a spatial trend surface plot.  
249 The local slope (time/distance) of the trend surface was then measured using a 3 × 3  
250 moving window filter, and the inverse of the slope value was used to estimate the speed  
251 of the spatial spread of SARS-CoV-2 (distance/time).<sup>20,21</sup>

252

#### 253 Estimation of epidemiological parameters

254 The net reproduction number ( $R_t$ ) was estimated using the EpiEstim R package.<sup>22</sup>  
255 Briefly, the estimate is based on a Bayesian approach that relies on the knowledge of  
256 the time series of infections (sampling date of positive tests in our analysis) and the  
257 distribution of the generation time. In the absence of estimates of the generation time  
258 for the analyzed outbreak, we assumed it to be gamma-distributed with a mean of 2.72



259 days (shape = 3.25, scale = 0.84), corresponding to the estimated serial interval during  
260 the Omicron wave in Hong Kong in the presence of strict interventions.<sup>19</sup>

261

262 We estimated the growth rate ( $r$ ) and 95% confidence interval (CI) in the second  
263 epidemic phase by fitting a linear regression to the logarithm of the incidence of  
264 infections. The doubling time in the early stage was then estimated as  $\ln(2)/r$ , where the  
265 95% CI was calculated using the delta method. All analyses and visualization were  
266 performed in ArcGIS 10.7 and R (version 4.0.2).

267

### 268 **Role of the funding source**

269 The funder of the study had no role in study design, data collection, data analysis, data  
270 interpretation, or writing of the report. The corresponding authors had full access to all  
271 the data in the study and had final responsibility for the decision to submit for  
272 publication.

273

274 This study was approved by the institutional review board of the School of Public  
275 Health, Fudan University (IRB# 2022-05-0968). All data were collected from publicly  
276 available sources. Data were de-identified, and the need for informed consent was  
277 waived.

278

### 279 **Results**

280 Following the successful containment of the initial epidemic wave caused by the  
281 Wuhan outbreak in early 2020, Shanghai was under constant pressure of potential  
282 outbreak introductions due to a high influx of international travelers relative to the rest  
283 of the cities in mainland China. Despite ranking first in the cumulative number of  
284 imported confirmed SARS-CoV-2 infections before 2022, Shanghai had sustained  
285 minimum local transmission prior to the emergence of the Omicron variant (**Figure 2a**),  
286 which was attributable to well-managed entry quarantine, screenings, and case isolation  
287 programs. Nonetheless, sporadic local transmissions events occurred, and all  
288 pre-Omicron outbreaks were swiftly suppressed.

289

290 Starting in mid-February 2022, Shanghai faced a significant surge of imported  
291 infections, the majority of which were linked to the Omicron wave in Hong Kong  
292 (**Supplementary Figure 3**). The number of imported infections reached a peak of 81  
293 per day on February 24—more than 19 times the prior daily average in 2020-2021 (4.1  
294 imported infections per day). Overall, from February 15 to March 15, a total of 1,112  
295 imported infections were reported (**Figure 2b**). Compared to any other time point

296 during the rest of the pandemic, the test positive rate among inbound travelers was at its  
297 highest throughout February 2022, reaching a peak of 96.3 infections per 100,000  
298 travelers (**Supplementary Figure 4**).

299

300 The sudden upsurge of infected travelers ultimately led to a breach of the entry control  
301 program, which might have resulted in the cryptic transmission of the Omicron variant  
302 since early March. A cluster of 14 SARS-CoV-2 positive individuals was detected  
303 starting from March 1 among attendees of a square dance event for seniors. Because of  
304 the Omicron variant's high transmissibility and immune evasive properties, as well as  
305 the delayed discovery of the outbreak, the contact tracing program was quickly  
306 overwhelmed, leading to a fast-growing epidemic that resulted in a total of 626,000  
307 SARS-CoV-2 infections as of May 31, 2022 (**Supplementary Figure 5**).

308

309 Most infections (96.0%) were identified in individuals under quarantine  
310 (**Supplementary Figure 6a**). The extent to which the infections were identified by  
311 community screening (e.g., essential workers) gradually declined from 13.6% in Phase  
312 2 to 2.7% in Phase 3. Meanwhile, the proportion of infections presenting with  
313 symptoms varied from 15.5% in Phase 1 to 9.8% in Phase 3 (**Supplementary Figure**  
314 **6b**).

315

316 The Omicron outbreak was widespread throughout the entire area of Shanghai, with  
317 99.5% (215/216) of the subdistricts reporting positive individuals. However, the spatial  
318 distribution of infections was highly heterogeneous, with 40% of the city area  
319 accounting for 80% of all infections; 35.5% of infections were identified in the Pudong  
320 New Area alone (**Supplementary Figure 5**). We divided the Omicron wave into three  
321 phases: Phase 1 (from January 1 to February 28), characterized by a high risk of  
322 importation; Phase 2 (from March 1 to 31), characterized by local community  
323 transmission; Phase 3 (after April 1), characterized by high levels of local transmission  
324 and the implementation of city-scale lockdowns to contain the outbreak. The total  
325 incidence of reported infections increased from 0.02 infections per 1,000 individuals in  
326 Phase 1 to 2.9 infections per 1,000 individuals in Phase 2 and reaching 21.9 infections  
327 per 1,000 individuals in Phase 3 (**Figure 3a-c**). Overall, an incidence of 24.8 infections  
328 per 1,000 individuals was observed (**Figure 3d**).

329

330 The spatial spread of the reported infections shows a clear trend from the city center  
331 towards adjacent suburban and rural areas (**Figure 4a-c**). Despite the implementation  
332 of a citywide lockdown in Phase 3, the outbreak continued to expand towards suburban  
333 and rural areas (**Figure 4c**). We estimated that spread speed of the infection throughout  
334 the city progressively slowed from an average of 544 meters/day in the first week of the

335 outbreak (February 27-March 5) to approximately 325 meters/day at the end of March  
336 (March 25-31), before the city-scale lockdown (**Figure 4d**). Although this decreasing  
337 trend was common to all districts, the central areas (including the districts of Jing'an,  
338 Yangpu, Hongkou, Putuo, Changning, Xuhui, and Huangpu) showed the fastest rate of  
339 spread, especially in the early stage of the outbreak (**Supplementary Figure 7**).

340

341 From March 16 to March 27, prior to the blanket lockdown of the entire city, Shanghai  
342 launched a set of targeted interventions and management strategies on a smaller  
343 geographical scale. The policy was adopted at the subdistrict level, and subdistricts  
344 were divided into high-risk areas (see definition in the **Methods** section), where the  
345 interventions were implemented, and non-high-risk areas. We further divided the  
346 non-high-risk areas into two categories based on their spatial proximity to the high-risk  
347 areas: moderate-risk areas (i.e., areas that were not classified as high-risk but were  
348 adjacent to high-risk areas) and low-risk areas (i.e., areas that were neither classified as  
349 high-risk nor adjacent to high-risk areas) (**Figure 5a**). From March 16 to March 29, the  
350 cumulative number of infections per 1,000 individuals was highest in high-risk areas  
351 (2.20, 95% CI, 2.18-2.22), followed by moderate-risk areas (1.51, 95% CI, 1.49-1.54)  
352 and low-risk areas (0.53, 95% CI, 0.49-0.57) (**Figure 5b-d**). The implemented targeted  
353 intervention strategies prior to the citywide lockdown were not sufficient to prevent  
354 infections to rise (**Figure 5e**), and  $R_t$  remained well above 1 (the epidemic threshold)  
355 across all three area categories for the entire period (**Figure 5f**). Moreover,  $R_t$  in  
356 moderate-risk and low-risk areas increased slightly from March 20 to March 24. The  
357 sensitivity analysis that considered alternative values for the delay between sampling  
358 and reporting showed similar patterns (**Supplementary Figure 8**). Overall, the growth  
359 rate of the epidemic in Phase 2 was estimated to be 0.216 per day (95% CI, 0.210 to  
360 0.222), with infections doubling every 3.21 days (95% CI, 3.12 to 3.29) (**Figure 5g**).

361

362 Three days after the implementation of a citywide lockdown on April 1 (Phase 3),  $R_t$   
363 started to decrease and fell below the epidemic threshold on April 14 (**Figure 6a-b**).  
364 Despite the marked differences in  $R_t$  observed between eastern and western Shanghai  
365 when the targeted interventions were adopted (Phase 2), immediately after the  
366 implementation of the citywide lockdown, the same level of SARS-CoV-2  
367 transmission ( $R_t$ ) was observed in all of Shanghai (**Figure 6b** and **Supplementary**  
368 **Figure 9**, where a sensitivity analysis confirming this finding is presented).

369

## 370 **Discussion**

371 This study provides a comprehensive picture of the spatiotemporal dynamics of the  
372 Omicron outbreak in Shanghai in 2022. Our findings highlight that, given the

373 Shanghai population's susceptibility to the SARS-CoV-2 Omicron BA.2 variant and  
374 its high transmissibility, the heightened importation risk of infected individuals from  
375 areas with widespread viral circulation was capable of triggering a major epidemic  
376 wave. The targeted interventions implemented in March were not enough to halt  
377 transmission. The citywide lockdown imposed on April 1 led to a decrease in  
378 transmission, bringing the reproduction number below the epidemic threshold in the  
379 entire city. Overall, the Omicron outbreak in Shanghai was successfully contained  
380 through a combination of stringent measures.

381

382 Our findings suggest that when facing a high influx volume of travelers from high-risk  
383 areas, breaches in the screening and quarantine of travelers may have the potential to  
384 trigger major outbreaks. It is also possible that the wave was simultaneously triggered  
385 by domestic travelers, as suggested by previous field investigations by local  
386 authorities.<sup>23</sup> We cannot exclude that the outbreak might have originated from multiple  
387 sources that led to simultaneous transmission chains prior to the identification of the  
388 first local transmission event.

389

390 Compared to previous local SARS-CoV-2 outbreaks that emerged in China after the  
391 original COVID-19 wave, the rapid spread of the Omicron outbreak in Shanghai was  
392 likely driven by the high transmissibility and/or immune-escape properties of Omicron  
393 BA.2.<sup>24</sup> The immunological landscape of Shanghai at the onset of this outbreak was  
394 fragile, with essentially no prior natural immunity (less than 0.01% of the population  
395 was infected prior to the outbreak<sup>25</sup>). In addition, although the coverage of the primary  
396 vaccination series exceeded 90% of the total population, the vaccination coverage for  
397 individuals aged  $\geq 60$  years was relatively low (62% for the primary series and 38% for  
398 the booster dose<sup>26</sup>). Moreover, the inactivated vaccines used in Shanghai provided very  
399 low protection against Omicron variant infection (approximately 17.0% after receiving  
400 a booster dose<sup>11</sup>, based on the level of neutralizing titer), which quickly wanes over  
401 time.<sup>27</sup> As a result, the Shanghai population was particularly vulnerable to the Omicron  
402 variant. Nonetheless, inactivated vaccines could provide high protection against severe  
403 outcomes.<sup>9,28</sup> These considerations further emphasize the importance of administering  
404 vaccines, especially to the most vulnerable segments of the population, and deploying  
405 immunization strategies with improved effectiveness against the current circulating  
406 variants through procuring currently available mRNA vaccines, approve other  
407 domestically developed vaccine candidates with broader protection against Omicron  
408 and future variants, and heterologous boosting strategies that have demonstrated  
409 improved antibody response.

410

411 Another factor that led to the major outbreak might have been the large number of  
412 asymptomatic carriers that contributed to widespread silent transmission.<sup>29</sup> Here, we  
413 found a high proportion (more than 90%) of asymptomatic infections, larger than that  
414 estimated for the ancestral lineage (69%<sup>30</sup>), but in line with other estimates for other  
415 SARS-CoV-2 variants in the presence of vaccination (85%<sup>31</sup>). Lower proportions of  
416 asymptomatic infections were reported in previous studies,<sup>32</sup> including for the  
417 Omicron BA.2 outbreak in Hong Kong<sup>19</sup>; however, those estimates were obtained in  
418 the absence of repeated screenings of the population. Moreover, the criteria adopted  
419 for the definition of asymptomatic infection may vary across locations and the study  
420 period.

421  
422 Between March 1 and March 31, a series of public health measures were adopted to  
423 reduce transmission between individuals living in areas at different risk levels, and the  
424 estimated doubling time in this phase of the epidemic was comparable to that of the  
425 Omicron wave in Hong Kong (3.2 days as compared to 3.4 days estimated for Hong  
426 Kong<sup>19</sup>). However, the implemented interventions were insufficient to curtail the  
427 epidemic. In fact, the repeated two-day lockdowns and population screenings  
428 implemented in high-risk areas were not sufficient to halt SARS-CoV-2 transmission,  
429 and these failed to identify all infected individuals (e.g., those in the early course of the  
430 infection, when the viral load was below the detection limit<sup>33</sup>). Once the two-day local  
431 lockdowns were lifted, unidentified infected individuals were allowed to freely move  
432 and transmit the infection to other areas, as shown by the rise in new daily infections  
433 identified in areas adjacent to high-risk areas.

434  
435 The implementation of the citywide lockdown prevented further growth of the  
436 epidemic and successfully contained the outbreak. However, given the number of  
437 infectious individuals at the time of implementation (over 10,000 new infections  
438 reported per day), it took 13 days for  $R_t$  to fall below the threshold. It is also possible  
439 that it was difficult to transport and isolate all positive individuals in a timely manner  
440 into dedicated facilities during the lockdown. Moreover, residents may still have been  
441 exposed to the virus inside their buildings.<sup>34</sup> It is thus important to carry out  
442 quantitative evaluations of the unavoidable risks posed by the adopted strategies.  
443 Empirical evidence from Shanghai shows that it is possible to maintain containment  
444 policies against the highly transmissible Omicron BA.2 variant, although this required  
445 unprecedented efforts to achieve. However, it is important to stress that the interruption  
446 of onward transmission in Shanghai might have prevented the infection from spreading  
447 into other cities, potentially preventing a major public health crisis in mainland China.<sup>11</sup>  
448 Nonetheless, it is important to stress that the measures adopted during the Omicron  
449 outbreak in Shanghai, including a strict and prolonged city-scale lockdown, would be

450 socially costly and impractical in the long term, emphasizing the importance refining  
451 NPI strategies to be less disruptive of daily lives and boosting population immunity  
452 level to significantly reduce the morbidity and mortality burden of SARS-CoV-2.<sup>11</sup>

453

454 A strength of our study is the unique features of the analyzed dataset. In fact,  
455 Shanghai's population-based screening policy, which allowed infections to be  
456 identified, likely resulted in a high infection ascertainment rate and the identification of  
457 the majority of infections throughout the outbreak. This allowed us to investigate the  
458 spatiotemporal spread of the Omicron BA.2 variant as well as the impact of the  
459 implemented interventions, all without significant amounts of underreporting from  
460 surveillance systems, as experienced in other areas. However, our study suffers from  
461 the limitations rooted in the uncertainty and fragmentary nature of publicly available  
462 sources. First, key variables, such as the date of symptom onset and the addresses of  
463 infected individuals, suffered from a high level of missing data. Second, the sparse  
464 information on the demographic characteristics of positive individuals limits the scope  
465 of our study. Third, the final clinical outcome of infected individuals was only partially  
466 available (e.g., we had no information about patients requiring intensive care treatment).  
467 Moreover, given the real-time nature of this study and the censoring of the final  
468 outcome data, an analysis of the COVID-19 burden is not possible at this point. Finally,  
469 this study does not provide quantitative estimates of the impact of specific  
470 interventions on transmission dynamics, but providing an overall assessment of the  
471 synergetic effect of the adopted interventions.

472

473 In conclusion, this study provides a quantitative description of the spatiotemporal  
474 spread of the Omicron BA.2 variant in Shanghai and explores the impact of the  
475 multifaceted implemented response measures. Our findings highlight the risk of  
476 widespread outbreaks in mainland China, particularly under the large pressure of  
477 imported infections. The successful containment of the Shanghai outbreak through the  
478 implementation of a strict and prolonged citywide lockdown shows that it is possible to  
479 successfully contain an Omicron outbreak in China, although it requires the  
480 implementation of a set of very stringent interventions. Disentangling the effects of  
481 each of the performed interventions might provide further insights into key public  
482 health priorities when faced with the emergence of future novel variants.

483 **Contributors**

484 H.Y. conceived and designed the study. H.Y. and JJ.Z. supervised the study. X.D., JJ.Z.,  
485 Z.C., JY.Z., J.C., Y.W., Y.T., N.Z., X.Y., R.S., X.X., X.Z., S.G., Y.L., and L.Y.  
486 collected and checked data. Z.C., JJ.Z., L.F., T.C., YP.W., JY.Z., J.C., H.L., K.S., Y.W.,  
487 T.W., and Y.T. analyzed the data. Z.C., JJ.Z. and H.Y. wrote the first draft of the  
488 manuscript. Z.C. JJ.Z, K.S., M.A., L.F., J.Y., and H.Y. interpreted the results and  
489 revised the content critically. All authors approved the final version for submission and  
490 agreed to be accountable for all aspects of the work.

491

492 **Declaration of interests**

493 H.Y. has received research funding from Sanofi Pasteur, GlaxoSmithKline, Yichang  
494 HEC Changjiang Pharmaceutical Company, Shanghai Roche Pharmaceutical  
495 Company, and SINOVAC Biotech Ltd. M.A. has received research funding from  
496 Seqirus. None of those research funding is related to this work. All other authors report  
497 no competing interests.

498

499 **Data sharing**

500 The data and code that support the findings of this study will be made available in  
501 GitHub upon manuscript acceptance.

502

503 **Acknowledgments**

504 The findings and conclusions in this report are those of the authors and do not  
505 necessarily represent the official position of the NIH. This study was supported by  
506 grants from the Key Program of the National Natural Science Foundation of China  
507 (grant 82130093 to H.Y.). The funders had no role in study design, data collection, data  
508 analysis, data interpretation or writing of the report.

509

510

511 **Figures legends**

512 **Figure 1. Timeline of the public health response in Shanghai by epidemic phase.**

513

514 **Figure 2. Temporal dynamics of local and imported SARS-CoV-2 infections in**

515 **Shanghai since early 2020. (a)** Number of reported SARS-CoV-2 infections in

516 Shanghai between 2020-2022, stratified by local and imported infections. **(b)** The

517 same as in (a), but for the period from January 1 to May 31, 2022.

518

519 **Figure 3. Geographical distribution of SARS-CoV-2 infections. (a-d)** Cumulative

520 number of new SARS-CoV-2 infections per 1,000 individuals in each phase and

521 overall.

522

523 **Figure 4. Spatial trends and speed of spread of the epidemic in the three phases.**

524 **(a)** Spatial location of the reported infections during the first phase of the epidemic.

525 **(b-c)** Estimated arrival time of the epidemic in the different areas of Shanghai.

526 Estimates are based on the thin spline regression of the interval between the time of

527 the detection of the first infection in each  $3 \text{ km} \times 3 \text{ km}$  grid and February 27, 2022.

528 Triangles indicate the potential source of the outbreak. **(d)** Estimated speed of spread

529 of SARS-CoV-2 (left axis) and cumulative fraction of affected areas of Shanghai

530 (right axis). Red dots indicate the speed of spread over time in each cell. The blue line

531 indicates the average speed per day as obtained using a polynomial regression. Central

532 areas contain the districts of Jing'an, Yangpu, Hongkou, Putuo, Changning, Xuhui,

533 and Huangpu.

534

535 **Figure 5. Characterization of the epidemic dynamics between March 16 and**

536 **March 29, 2022. (a)** Location of high-risk, moderate-risk, and low-risk areas. For

537 each area, its highest risk classification was used. **(b-d)** Number of reported infections

538 between March 16 and March 29 by area type. **(e)** Number of new reported infection

539 per 1,000 individuals by area type. **(f)** Estimated  $R_t$  between March 16 and March 29

540 by area type. **(g)** Estimated epidemic growth rate and doubling time (days).

541

542 **Figure 6. Epidemic dynamics under the effect of interventions. (a)** Number of new

543 SARS-CoV-2 infections by date of sample collection for means of identification. **(b)**

544 Estimated  $R_t$  (mean and 50% confidence interval) in eastern, western, and all

545 Shanghai areas.



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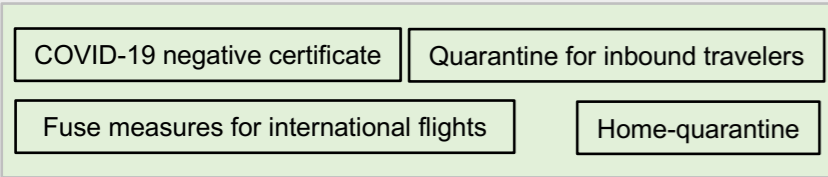
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639

### Phase 1: January 1 to February 28

#### Baseline non-pharmaceutical interventions (NPIs)

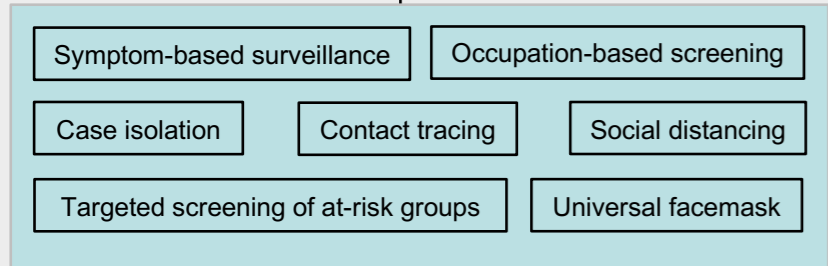
##### Border control policy



January 1, 2022

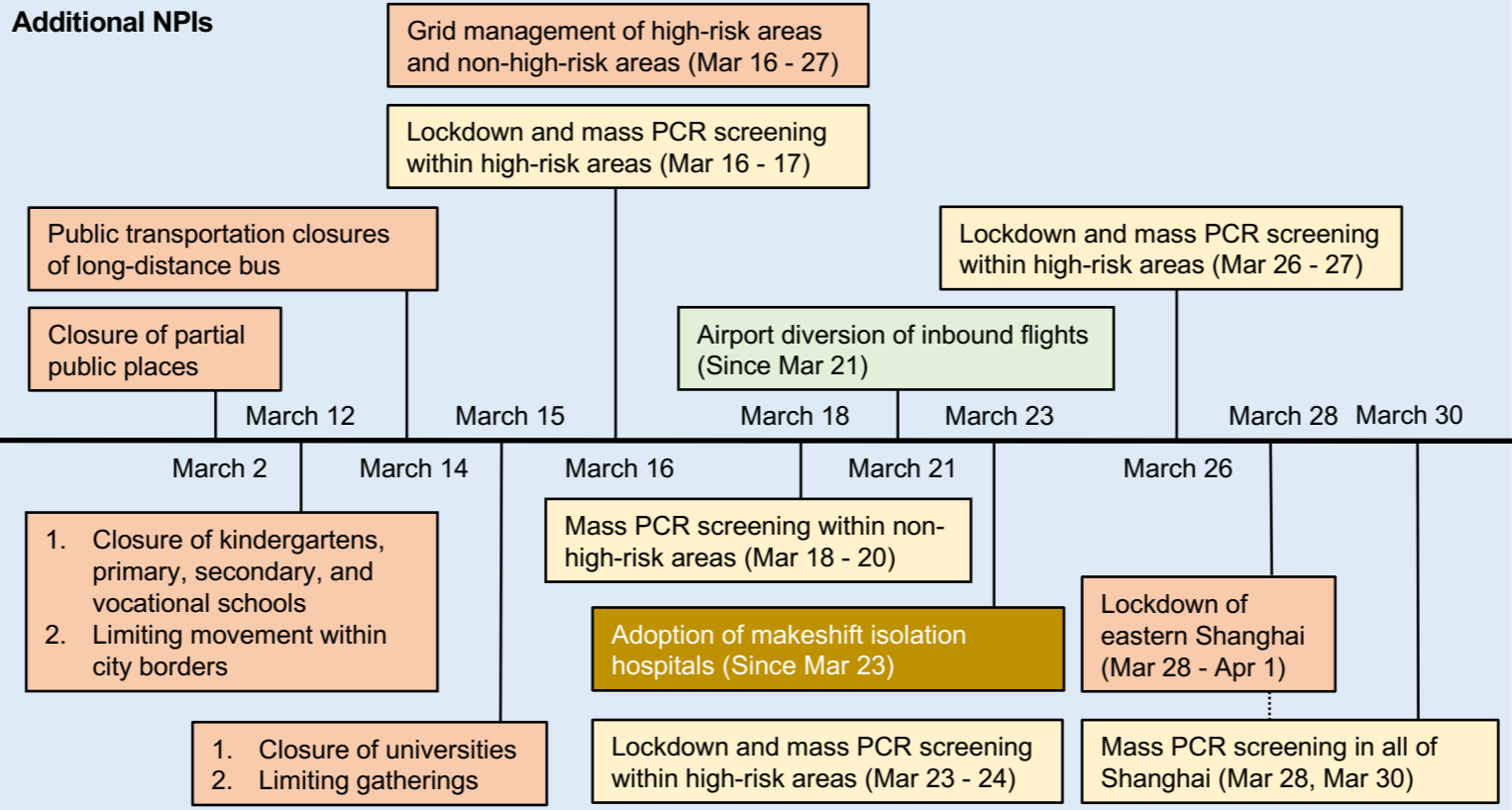
February 28, 2022

##### Local control policy



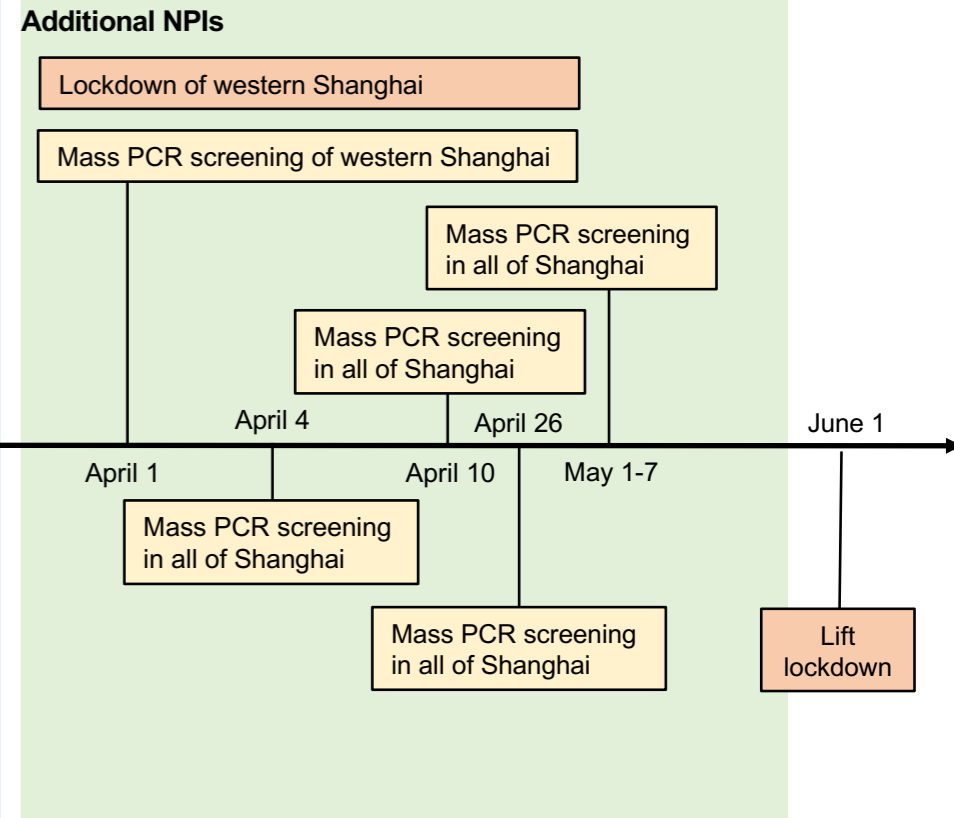
### Phase 2: March 1 to March 31

#### Additional NPIs

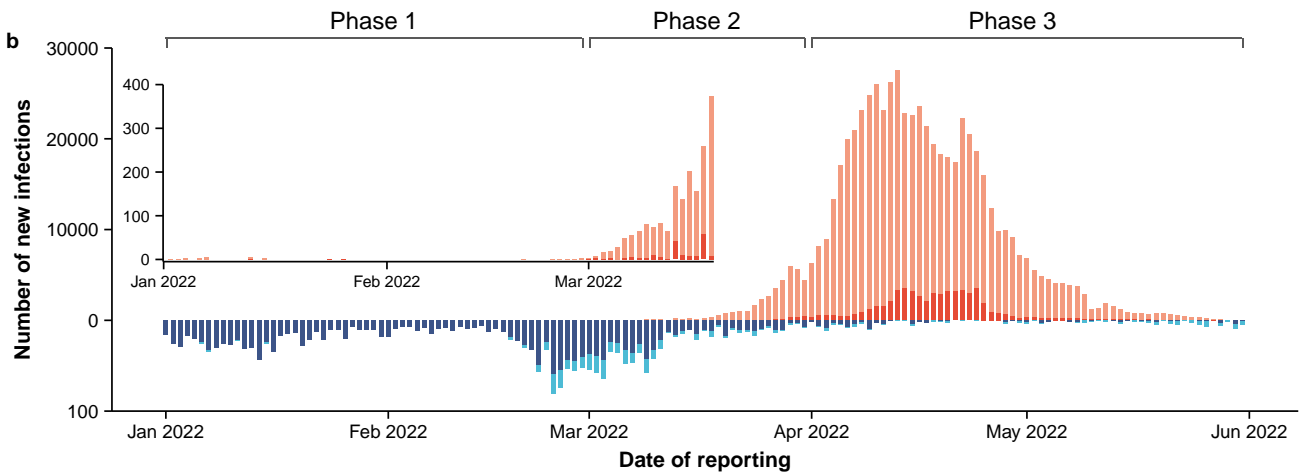
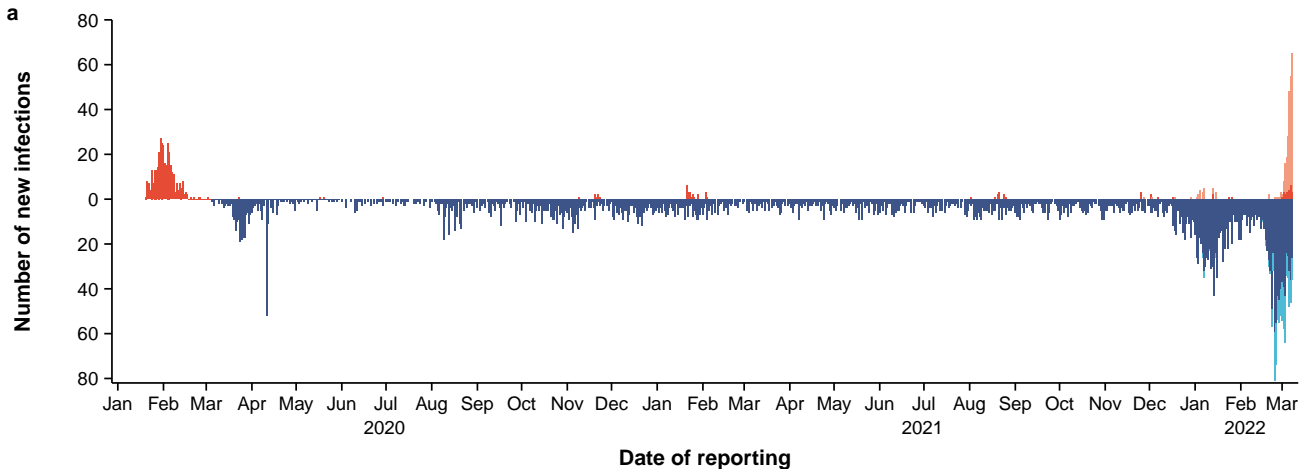


### Phase 3: April 1 to May 31

#### Additional NPIs



Border control policy
  Policy of case isolation
  Mass screening policy
  Social distancing policy



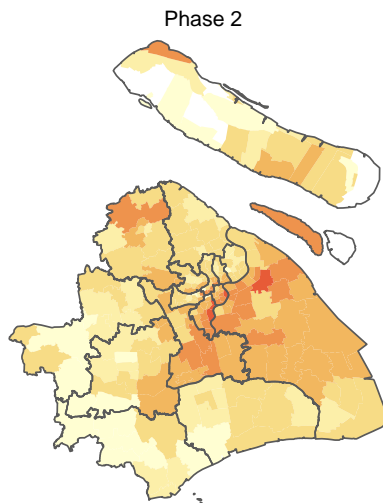
**Symptomatic status and sources**

- Local asymptomatic infections
- Imported asymptomatic infections
- Local symptomatic cases
- Imported symptomatic cases

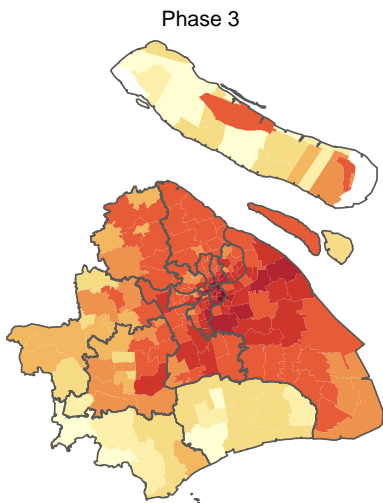
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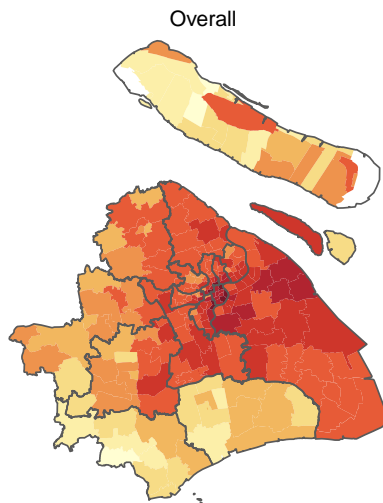
b



c



d



Cumulative number  
of infections per  
1,000 individuals

