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## Title page

## Original Article

**Title:** Epidemiologic trends and distributions of imported infectious diseases among travelers to Japan before and during the COVID-19 pandemic, 2016 to 2021: a descriptive study

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Accepted Version

## 1 **Abstract**

2 **Background:** Little is known about the trends of imported infectious diseases among  
3 travelers to non-endemic countries during the COVID-19 pandemic. This article aimed to  
4 describe those among travelers to Japan.

5 **Methods:** This is a descriptive study based on national surveillance data. Imported infectious  
6 disease cases were defined as those with a reported overseas source of infection among 15  
7 diseases pre-selected based on the probability and impact of importation. The number of  
8 notified cases from April 2016 to March 2021 were described by disease and time of  
9 diagnosis. The relative ratio and absolute difference in case counts—both by number and per  
10 arrival—were calculated by disease comparing those from the pandemic period (April 2020–  
11 March 2021) to the pre-pandemic period (April 2016–March 2020).

12 **Results:** A total of 3524 imported infectious disease cases were diagnosed during the study  
13 period, including 3439 cases before and 85 cases during the pandemic. The proportionate  
14 distribution of diseases changed but notification counts of all 15 diseases decreased during the  
15 pandemic. Accounting for arrivals, however, seven diseases showed a two-fold or greater  
16 increase, with a notable absolute increase per million arrivals for amebiasis (60.1; 95%CI,  
17 41.5–78.7), malaria (21.7; 10.5–33.0), and typhoid fever (9.3; 1.9–16.8).

18 **Conclusion:** The epidemiology of imported infectious diseases changed during the pandemic.  
19 While the number of imported infectious disease cases decreased, the number of cases per  
20 arrivals increased considerably both in relative and absolute terms for several diseases of  
21 public health and clinical importance.

22

23 **Key words:** COVID-19; epidemiology; imported infectious disease; surveillance; travel

24

## 25 **Background**

26 In response to COVID-19, which was declared a pandemic by the World Health  
27 Organization on March 11, 2020,<sup>1</sup> governments globally have implemented international  
28 travel restrictions.<sup>2</sup> These measures have been followed by a substantial decline in the number  
29 of international travelers worldwide.<sup>2</sup> Japan has been no exception. Since February 2020,  
30 Japan had taken measures to enhance border control measures, including entry restrictions  
31 depending on the country of departure for foreign nationals.<sup>3</sup> Subsequently, the total number  
32 of travelers entering Japan in 2020 fell sharply from the hitherto upward trend.<sup>4</sup> On the other  
33 hand, some exceptions were allowed, such as permitting foreign nationals with “special  
34 exceptional circumstances” (e.g., spouses or children of Japanese nationals/permanent  
35 residents, or foreign nationals with residency status of “Diplomat” or “Official”) to enter  
36 Japan,<sup>3</sup> and establishing special quota pertaining to cross-border business travelers between  
37 Japan and certain countries and regions.<sup>5</sup> In this context, not only the number of travelers to  
38 Japan but also traveler characteristics and countries of travel origin could have changed  
39 during the pandemic.

40 Several studies have shown that, since before the pandemic, air travel data such as  
41 the number and demographic characteristics of travelers as well as their countries of origin are  
42 associated with patterns of infectious disease importations.<sup>6-9</sup> Given the situation in Japan  
43 under the COVID-19 pandemic, it was possible that the number and distributions of infectious  
44 disease cases among travelers to non-endemic countries had changed. The aim of this study  
45 was to describe the important changes in trends and distributions of imported infectious  
46 diseases among travelers entering Japan before and during the pandemic.

47

## 48 **Methods**

49 Study design

50 This is a retrospective, mostly descriptive analysis of national surveillance data combined  
51 with publicly available national migration statistics.

52

### 53 Data sources

54 The National Epidemiological Surveillance of Infectious Diseases (NESID) system has been  
55 operating under the Infectious Diseases Control Law since 1999. Physicians are required to  
56 notify all notifiable diseases to the public health centers (the reporting criteria and the  
57 notification form for each disease are publicly available),<sup>10</sup> who coordinate with prefectural  
58 and municipal public health institutes (e.g. laboratory testing). The notification form includes  
59 demographic, clinical, laboratory, and exposure information. The data collected are then  
60 reported by the public health centers and institutes via the electronic NESID system. We  
61 extracted data on cases diagnosed between April 1, 2016 and March 31, 2021 on December  
62 28, 2021. The variables used in the analysis were patient name (to differentiate between  
63 Japanese and non-Japanese), sex, symptoms/signs, diagnostic methods, date of diagnosis,  
64 presumed date of infection, and suspected country of infection.

65 The monthly number of arrivals into Japan and their nationality was obtained in April  
66 2022 from the Immigration Services Agency of Japan website.<sup>4</sup> Demographic and travel-  
67 related information of arrivals, including length of stay and status of residence, were obtained  
68 from the annual report on the same website.

69

### 70 Case definition

71 In 2018, 15 notifiable diseases with a historically sizable number and proportion of imported  
72 cases were systematically selected as priority imported infectious diseases by the National  
73 Institute of Infectious Diseases; these 15 have been monitored continuously since then. These  
74 are amebiasis, chikungunya, cryptosporidiosis, dengue, giardiasis, hepatitis A, hepatitis E,

75 leptospirosis, malaria, measles, paratyphoid fever, rubella, shigellosis, typhoid fever, and Zika  
76 virus disease. In accordance with the definition, an “imported case” was defined as a reported  
77 case whose source of infection was determined to be overseas by the physician who  
78 diagnosed and reported the case.

79

#### 80 Data analysis

81 We described the number of arrivals in Japan by month and by fiscal year (FY) running from  
82 April to March. Demographic and travel-related information were presented by calendar year  
83 owing to the limitations of the available data.

84 We described the monthly trends in the number of 15 imported infectious disease  
85 cases and the annual trends in the proportionate distribution of the 15 diseases. We then  
86 assessed the importation data by first describing the case counts before (April 1, 2016 to  
87 March 31, 2020, corresponding to FY2016–2019) and during the pandemic period (April 1,  
88 2020 to March 31, 2021, corresponding to FY2020), based on the timing of the sharp decline  
89 in the number of travelers due to travel restrictions imposed by the government.<sup>4</sup> Next, we  
90 compared the case counts by calculating the ratio and the difference between the two periods;  
91 the ratio provides a relative comparison while the difference indicates an absolute change. To  
92 account for the denominator of number of travelers, we similarly calculated the ratios and  
93 differences, per million arrivals, for the respective periods using the aggregate number of  
94 arrivals of 184,042,453 in FY2016–2019 and 697,618 in FY2020.<sup>4</sup> The 95% confidence  
95 interval (CI) for the ratio and difference per million arrivals before and during the pandemic  
96 was estimated to indicate the level of precision.

97 Amebiasis and malaria, with the largest absolute increase in the number of cases per  
98 arrivals, were further analyzed to explore the potential reasons for this increase. We described  
99 amebiasis cases based on sex, symptoms, and time of FY infection. Amebiasis cases with

100 specific colonic mucosal lesions or positive fecal occult blood but that were otherwise  
101 asymptomatic were categorized as asymptomatic cases.<sup>11</sup> The ratio and the difference of non-  
102 Japanese malaria cases per million foreign national arrivals, before and during the pandemic,  
103 were described by region, based on the travel origin of the cases and nationality of the  
104 arrivals. Japanese nationals were excluded from the analysis because of a lack of regional  
105 denominator data. Cases whose suspected country of infection were unknown or included  
106 more than one country were excluded from the analysis by region. Statistical analysis was  
107 performed using Stata/MP version 16.0.

108

#### 109 Ethical consideration

110 Information on notified cases was collected under the Infectious Diseases Control Law. The  
111 use of national surveillance data for public health purposes does not require informed consent  
112 from the patient or ethical approval from the relevant authorities. For those diseases that  
113 included identifiable data, strict data management practices were implemented per standard  
114 protocol.

115

## 116 **Results**

### 117 Arrivals to Japan

118 The traveler volume to Japan dropped remarkably during February–April 2020 and remained  
119 low from April 2020 onwards (Figure 1). The total annual number of arrivals in FY2020 was  
120 697,618, a 98% decrease from the annual average of 46,641,690 in FY2016–2019 (Table 1).  
121 Whilst the proportion of foreign national arrivals exceeded that of Japanese nationals in  
122 FY2016–2019, this was not the case in FY2020. Asian nationals consistently constituted the  
123 majority of foreign national arrivals across all five years, and the distribution of nationalities  
124 from other regions remained largely stable. The proportion of African nationals remained low,

125 averaging 0.2% in FY 2016–2019 and 1.2% in FY2020.

126 The demographic and travel-related statistics of arrivals by nationality are presented  
127 in eFigures 1 and 2. The proportion of Japanese national arrivals who stayed at their travel  
128 origin for more than a month increased noticeably during the pandemic, compared to the pre-  
129 pandemic period (eFigure 1). During the pandemic, the proportion of temporary foreign  
130 visitors decreased, while the proportion of arrivals with employment qualifications or  
131 residency status increased (eFigure 2).

132

133 Notification trends and distributions of 15 imported infectious diseases

134 A total of 3524 imported infectious disease cases were diagnosed during FY2016–2020,  
135 including 3439 cases during FY2016–2019 and 85 cases in FY2020. During the pandemic  
136 FY2020 period, 8/15 diseases were reported but chikungunya, cryptosporidiosis,  
137 leptospirosis, measles, paratyphoid fever, rubella and zika virus infection were not reported.  
138 The monthly number of reported cases declined over February–April 2020 and remained low  
139 thereafter (Figure 2). The decline in case counts coincided with the drop in the number of  
140 travelers.<sup>4</sup>

141 Among the 15 imported infectious diseases, dengue accounted for the highest  
142 proportion in FY2016–2019, averaging 34.7% (308/896 (34.4%), 219/749 (29.2%), 245/818  
143 (30.0%) and 423/976 (43.3%), respectively) (Figure 3). However, the proportion of dengue  
144 declined to 10.6% (9/85) in FY2020. Meanwhile, the proportion of amebiasis cases among the  
145 15 diseases increased substantially from its average of 15.9% (186/896 (20.8%), 156/749  
146 (20.8%), 107/818 (13.1%), and 99/976 (10.1%)) in FY2016–2019 to 51.8% (44/85) in  
147 FY2020. Similarly, the proportion of malaria increased almost 3-fold from 6.5% (56/896  
148 (6.3%), 63/749 (8.4%), 48/818 (5.9%), and 56/976 (5.7%)) to 18.8% (16/85).

149 In all 15 diseases, the number of cases declined (Table 2). The largest decrease was



150 observed in dengue, followed by amebiasis and shigellosis. However, none of the 15 diseases  
151 showed a substantial decline in the number of cases when accounting for the number of  
152 travelers; in fact, the number of dengue cases per million arrivals increased (Table 2). Relative  
153 to the pre-pandemic period, a two-fold or greater increase in the number of cases per arrival  
154 was observed for dengue, malaria, amebiasis, giardiasis, hepatitis A, hepatitis E, and typhoid  
155 fever. The highest absolute increase was observed in amebiasis, malaria, and typhoid fever,  
156 with an increase of 60.1 (95%CI, 41.5–78.7), 21.7 (10.5–33.0), and 9.3 (1.9–16.8) per million  
157 arrivals, respectively. Despite having the same modes of transmission,<sup>12</sup> different trends were  
158 observed in the change in the number of cases per arrival for vector-borne (e.g. malaria vs.  
159 dengue) and food-borne infectious diseases (e.g. typhoid fever vs. paratyphoid fever).

160

#### 161 Amebiasis case analysis

162 The proportion of male cases averaged 88% (163/186 (88%), 142/156 (91%), 92/107 (86%),  
163 and 85/99 (86%)) for amebiasis cases in FY2016–2019 and 86% (38/44) in FY2020, showing  
164 no major changes before and during the pandemic. The proportion of asymptomatic cases  
165 increased from an average of 25.9% (42/186 (22.6%), 39/156 (25.0%), 32/107 (29.9%), and  
166 26/99 (26.3%)) during FY2016–2019 to 34.1% (15/44) in FY2020. The annual proportion of  
167 cases with unknown time of infection also increased from an average of 61.1%  
168 (120/186(64.5%), 104/156 (66.7%), 65/107 (60.7%), and 52/99 (52.5%)) to 70.5% (31/44), as  
169 did the proportion with presumed infection for more than one year before diagnosis, from  
170 7.5% (10/186 (5.4%), 11/156 (7.1%), 10/107 (9.3%), and 8/99 (8.1%)) to 15.9% (7/44).

171

#### 172 Malaria case analysis

173 The proportion of non-Japanese malaria cases increased to 68.8% (11/16) during the  
174 pandemic, compared to an average annual proportion of 55.3% (28/56 (50.0%), 44/63

175 (69.8%), 23/48 (47.9%), and 30/56 (53.6%)) in FY2016–2019 before the pandemic. The  
176 absolute increase in non-Japanese notifications per million foreign national arrivals was 31.1  
177 (12.0–50.1) (Table 3), higher than 21.7 (10.5–33.0) when including all arrivals. When  
178 stratified by region, excluding five cases (pre-pandemic, 4/125; pandemic, 1/11) with  
179 unknown or multiple suspected countries of infection, the majority of non-Japanese cases  
180 consistently originated from Africa (38/56 (67.9%), 47/63 (74.6%), 38/48 (79.2%), 45/56  
181 (80.4%), and 12/16 (75.0%)). Following the pandemic, the absolute increase in non-Japanese  
182 cases per million foreign national arrivals from Africa was high, at 1537.7 (88.9–2986.6). In  
183 terms of the relative increase, the number of non-Japanese cases per foreign arrivals from all  
184 regions pooled and Africa rose 28-fold and 4-fold, respectively (Table 3).

185

## 186 **Discussion**

187 The COVID-19 pandemic saw a notable decrease in traveler volume to Japan and a drastic  
188 shift in their characteristics, suggesting that the travel situation was considerably affected by  
189 travel restrictions and other related measures. Notably, the annual number of all 15 imported  
190 infectious disease cases decreased, along with a marked decline in the number of arrivals.<sup>4</sup>  
191 This was consistent with pre-pandemic findings which showed a positive correlation between  
192 the passenger volume and the number of imported cases.<sup>7</sup> However, this decline differed by  
193 disease and the proportion of malaria cases increased, indicating a proportionately greater  
194 importance of malaria importation for Japan.

195         Moreover, seven of the 15 diseases showed two-fold or greater relative increase in  
196 the number of cases per arrival during vs. before the pandemic, implying that the relative  
197 “risk” of detecting the disease among arrivals actually increased (though strictly speaking,  
198 “risk” represents a crude ratio of the number of cases to that of arrivals in a given time period,  
199 and the denominator may have included those not considered to be at risk for some of the

200 diseases). Furthermore, while dengue showed a large decrease in both the case counts and as a  
201 proportion among the 15 diseases, the notification rate among travelers had doubled. Notably,  
202 in addition to the relative increase, the substantial absolute increase accounting for travelers  
203 was observed for amebiasis, malaria, and typhoid fever. Therefore, despite the decrease in the  
204 number of importations, the relative and absolute risk among travelers for several diseases  
205 showed an appreciable increase.

206         The ratio comparing the notification rate accounting for arrivals between the two  
207 periods indicates a relative change. As with the concept of risk difference,<sup>13</sup> on the other hand,  
208 considering the difference in notifications per arrival accounts for the absolute risk and can  
209 quantify the notification rate change in absolute terms. Its importance can be illustrated by an  
210 example among non-Japanese malaria importations. The ratio of the number of cases per  
211 million arrivals during the pandemic compared to that of the pre-pandemic period was 28 for  
212 all regions (pooled) vs. 4 for Africa. This suggests that the pandemic period increased the risk  
213 of malaria importation relatively more among those from all regions compared to those from  
214 Africa. However, when the difference was considered, the order was reversed, being 31 and  
215 1538 per million foreign national arrivals, respectively. Hence, during the pandemic, while the  
216 malaria notification rate showed a greater relative increase for all regions (pooled), the  
217 absolute risk of malaria importation increased more among those from Africa. Similarly,  
218 while the notification rate for giardiasis increased six-fold while that for dengue only doubled,  
219 the absolute change in the notification rate per million arrivals was 2 for giardiasis and 6 for  
220 dengue. Thus, information provided by the difference in case counts per arrival can also be  
221 useful for public health decision making.<sup>13</sup>

222         Given the ongoing high risks among travelers, the following pre-pandemic travel-  
223 related concerns may be present: first, some of these diseases including malaria and dengue  
224 may cause fatal outcomes especially in high-risk travelers<sup>14,15</sup>; second, physicians in non-

225 endemic countries are unfamiliar with certain infectious diseases and are less likely to include  
226 them for differential diagnosis, which may delay diagnosis and treatment. In addition to these  
227 previously highlighted issues, new concerns have been raised under the pandemic: co-  
228 infection with dengue and COVID-19 has been reported to be associated with severe and fatal  
229 outcomes<sup>16</sup>; physicians may overlook some infectious diseases by focusing on COVID-19<sup>17</sup>;  
230 there may also have been changes in healthcare-seeking behavior and challenges in accessing  
231 healthcare. Therefore, considering the continued high risk among travelers despite the decline  
232 in notified case counts, public health authorities should continue their efforts to ensure that  
233 patients receive early diagnosis and treatment to prevent serious outcomes. Such quantitative  
234 evaluation of the risks posed to travelers could help public health practitioners to effectively  
235 communicate with physicians, improving their awareness.

236 In the additional analysis of amebiasis, the proportion of asymptomatic cases that  
237 may have been diagnosed incidentally,<sup>11</sup> cases that took more than one year from presumed  
238 infection to diagnosis, and cases with an unknown time of infection increased. These results  
239 suggest that cases diagnosed during the pandemic involved a certain number of those infected  
240 before the pandemic and the data may not reflect the trend of amebiasis imported during the  
241 pandemic. Therefore, given the drastic decrease in the number of travelers, the case counts  
242 among arrivals could have resulted in an apparent increase.

243 For malaria, the following factors may have contributed to their increased  
244 notifications per arrival. The largest number of malaria cases were reported as infected in  
245 Africa. In 2020, Sub-Saharan Africa experienced a malaria epidemic,<sup>18</sup> which may have  
246 contributed to the absolute increase in the notification rate among all arrivals. The difference  
247 in the number of cases per arrival was found to be larger when restricted to non-Japanese than  
248 when including non-Japanese and Japanese. This indicates that the absolute risk greatly  
249 increased among the non-Japanese. Particularly, when stratified by region, the increase among

250 African national arrivals was larger than that among those from other regions. Regarding the  
251 characteristics of foreign national arrivals, the proportion of temporary visitors and those with  
252 non-working status decreased substantially during the pandemic. In contrast, the distribution  
253 of those with employment or residence visas had increased considerably. Given this context  
254 and known risk factors for malaria among travelers such as visiting friends and relatives  
255 (VFRs), long-term stay, and travel to endemic countries,<sup>19,20</sup> the proportion of high-risk  
256 arrivals including long-term residence in endemic areas and VFR returnees may have  
257 increased. This may have contributed to the substantial increase in notifications among  
258 African arrivals. In addition, it is possible that the increased proportion of foreign national  
259 travelers staying in Japan through the incubation period made them more likely to be detected  
260 domestically. Meanwhile, the proportion of African nationals among foreign national arrivals  
261 remained very low. Taken together, the risk of malaria among all travelers entering Japan  
262 could have been affected by travelers' characteristics and local epidemics rather than by  
263 change in the travel volume from Africa.

264 Based on these findings, we believed that the altered travel situation such as the  
265 demographic composition, length of stay, and destination/origin of travel may have affected  
266 the trends and distributions of imported infectious diseases per travelers in FY2020, although  
267 there may be variations in degree depending on individual diseases. Due to the  
268 implementation of strict border control measures throughout the world during the same  
269 period, such findings (e.g. malaria) may also have been observed in other countries and be of  
270 relevance. As the COVID-19 pandemic shifts towards an endemic phase, many countries,  
271 including Japan, are further relaxing travel restrictions. With such a drastic change in the  
272 travel context, not only describing trends in the number of cases but accounting for the  
273 number of travelers would be useful for public health authorities to assess risk for response.  
274 Furthermore, considering the difference in the notification rate among travelers can inform

275 them about meaningful absolute changes, contributing to their selection of diseases that  
276 should be prioritized for action, particularly when resources are overwhelmed by public  
277 health emergencies such as COVID-19.

278 Our study has several limitations. First, trends in distributions may not be captured to  
279 the same extent as that before the pandemic if health-seeking behaviors or testing capacities  
280 had changed. If the implementation of health monitoring for arrivals during their quarantine  
281 period<sup>21</sup> had facilitated detection of individuals with imported diseases other than COVID-19,  
282 case detection during the pandemic may have become more sensitive. Second, overestimation  
283 may have also occurred for diseases that can take a long time from infection to diagnosis,  
284 such as amebiasis. Third, non-Japanese cases of malaria may have been misclassified by  
285 assumptions based on their names. However, the misclassification was expected to have  
286 occurred equally before and during the pandemic and this would have had little impact.  
287 Fourth, in the analysis of the number of cases per arrival by region, the numerator was the  
288 number of cases by region of infection and the denominator was the number of travelers by  
289 nationality; thus, a proportion of the numerator may not have been included in the  
290 denominator.

291

## 292 **Conclusions**

293 Although the number of cases notified under the national infectious disease surveillance  
294 scheme decreased for all 15 imported infectious diseases during the COVID-19 pandemic in  
295 Japan, relative increase in cases per travelers was observed for several diseases. Moreover, the  
296 number of cases per travelers increased considerably for amebiasis, malaria, and typhoid  
297 fever. In the context of drastic shifts in travel patterns, it is essential to account for the number  
298 of travelers and consider changes in both relative and absolute terms. Our findings and these  
299 considerations are important for public health practitioners to communicate to physicians to

300 facilitate vigilance against imported infectious diseases.

301

### 302 **Acknowledgments**

303 This research was based on cases reported by the prefectural and municipal public health  
304 institutes (PHIs) and public health centers (PHCs) to the NESID system. Some of the cases  
305 were diagnosed based on microbiological tests performed at PHIs/PHCs. We gratefully  
306 appreciate staff members from the PHCs and PHIs, along with the reporting physicians, for  
307 providing information on the 15 infectious diseases included in this study.

308

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312 Welfare, Japan.

313

### 314 **Conflicts of interest**

315 The authors declare they have no conflict of interest with respect to this research study and  
316 paper.

317

### 318 **Data availability**

319 The travel data used in this article are available in “Statistics on Legal Migrants  
320 ([https://www.moj.go.jp/isa/policies/statistics/toukei\\_ichiran\\_nyukan.html](https://www.moj.go.jp/isa/policies/statistics/toukei_ichiran_nyukan.html))”. The national  
321 infectious disease surveillance data are included in published “Trends in Notification of  
322 Imported Cases among Select Notifiable Infectious Diseases in Japan  
323 (<https://www.niid.go.jp/niid/ja/route/transport.html>)” and “Infectious Diseases Weekly Report  
324 (<https://www.niid.go.jp/niid/ja/data.html>)”.

325

326 **Authors' contributions**

327 AK, MF, YA designed the study. AK, SO, HN, TS, YS, AT, HF, CI, SN, YF and KK verified  
328 and extracted the data. AK, TA, MF and YA analyzed and interpreted the data. AK wrote the  
329 first draft of the manuscript. All authors provided important comments on the draft  
330 manuscript. MF and TSu obtained the funding. YA, MF, TSu and MS provided administrative  
331 or material support. All authors read and approved the manuscript.

332

333



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**Figure 1 title:** Number of arrivals to Japan by month of arrival, by nationality, April 2016–March 2021

**Figure 2 title:** Number of imported infectious disease cases by month of diagnosis, by disease, Japan, April 2016–March 2021 (restricted to pre-selected 15 priority notifiable diseases)

**Figure 3 title:** Proportionate distribution of imported infectious disease cases by year of diagnosis, Japan, April 2016–March 2021 (restricted to pre-selected 15 priority notifiable diseases)

**Table 1:** Number of arrivals to Japan, April 2016–March 2021

	FY2016		FY2017		FY2018		FY2019		FY2020	
Total arrivals	41,529,240		46,603,677		50,096,935		45,812,601		697,618	
Japanese	17,455,991	42.0%	17,981,203	38.6%	19,237,954	38.4%	18,403,845	40.2%	348,741	50.0%
Non-Japanese	23,904,199	57.6%	28,443,905	61.0%	30,666,253	61.2%	27,223,118	59.4%	323,742	46.4%
Region of nationality										
Africa	38,042	(0.2%)	40,181	(0.1%)	44,001	(0.1%)	56,858	(0.2%)	3,837	(1.2%)
Asia	20,072,119	(84.0%)	24,170,161	(85.0%)	25,974,815	(84.7%)	22,328,118	(82.0%)	274,896	(84.9%)
Europe	1,506,059	(6.3%)	1,665,549	(5.9%)	1,831,814	(6.0%)	1,957,879	(7.2%)	19,140	(5.9%)
North America	1,642,910	(6.9%)	1,825,944	(6.4%)	2,014,017	(6.6%)	2,025,215	(7.4%)	13,332	(4.1%)
Oceania	513,686	(2.1%)	588,693	(2.1%)	639,750	(2.1%)	690,584	(2.5%)	2,164	(0.7%)
South America	130,163	(0.5%)	152,519	(0.5%)	161,025	(0.5%)	163,634	(0.6%)	10,343	(3.2%)
No nationality	1220	(0.0%)	858	(0.0%)	831	(0.0%)	830	(0.0%)	30	(0.0%)
SOFA personnel	169,050	0.4%	178,569	0.4%	192,728	0.4%	185,638	0.4%	25,135	3.6%

FY, fiscal year (from April to March of the subsequent year); SOFA, Status of Forces Agreement. % represents the proportion among total arrivals for the fiscal year (% in parentheses represent the proportion among non-Japanese (i.e. foreign) nationals).

**Table 2:** Numbers of imported infectious disease cases, imported infectious disease cases per 1,000,000 arrivals, and their ratios and differences during the pandemic compared to the pre-pandemic period, by disease, Japan, April 2016–March 2021

Main mode of transmission	Disease	Period <sup>a</sup>	Annual number of cases <sup>b</sup>	Ratio of the number of cases during the pandemic to the pre-pandemic period	Difference in the number of cases from the pre-pandemic period	Number of cases per million arrivals <sup>c</sup>	Ratio of the number of cases per million arrivals during the pandemic to the pre-pandemic period (95% CI)	Difference in the number of cases per million arrivals from the pre-pandemic period (95% CI)
Vector-borne (mosquito-borne)	Chikungunya	pre-pandemic	18	ref.	ref.	0.4	ref.	ref.
		pandemic	0	0.0	−18	0.0	0.0 (0.0 to 13.7)	−0.4 (−0.5 to −0.3)
	Dengue	pre-pandemic	299	ref.	ref.	6.5	ref.	ref.
		pandemic	9	0.0	−290	12.9	2.0 (0.9 to 3.8)	6.4 (−2.0 to 14.8)
	Malaria	pre-pandemic	56	ref.	ref.	1.2	ref.	ref.
		pandemic	16	0.3	−40	22.9	18.9 (10.6 to 31.4)	21.7(10.5 to 33.0)
	Zika virus infection	pre-pandemic	4	ref.	ref.	0.1	ref.	ref.
		pandemic	0	0.0	−4	0.0	0.0 (0.0 to 63.9)	−0.1 (−0.1 to 0.0)
Food-borne/ water-borne	Amebiasis	pre-pandemic	137	ref.	ref.	3.0	ref.	ref.
		pandemic	44	0.3	−93	63.1	21.2 (15.2 to 28.8)	60.1 (41.5 to 78.7)
	Cryptosporidiosis	pre-pandemic	2	ref.	ref.	0.0	ref.	ref.

		pandemic	0	0.0	-2	0.0	0.0 (0.0 to 224.1)	0.0 (-0.1 to 0.0)
	Giardiasis	pre-pandemic	23	ref.	ref.	0.5	ref.	ref.
		pandemic	2	0.1	-21	2.9	5.7 (0.7 to 21.1)	2.4 (-1.6 to 6.3)
	Hepatitis A	pre-pandemic	63	ref.	ref.	1.4	ref.	ref.
		pandemic	2	0.0	-61	2.9	2.1 (0.3 to 7.6)	1.5 (-2.5 to 5.5)
	Hepatitis E	pre-pandemic	21	ref.	ref.	0.5	ref.	ref.
		pandemic	4	0.2	-17	5.7	12.4 (3.3 to 33.0)	5.3 (-0.3 to 10.9)
	Paratyphoid fever	pre-pandemic	19	ref.	ref.	0.4	ref.	ref.
		pandemic	0	0.0	-19	0.0	0.0 (0.0 to 13.5)	-0.4 (-0.5 to -0.3)
	Shigellosis	pre-pandemic	92	ref.	ref.	2.0	ref.	ref.
		pandemic	1	0.0	-91	1.4	0.7 (0.0 to 4.0)	-0.6 (-3.4 to 2.3)
	Typhoid fever	pre-pandemic	33	ref.	ref.	0.7	ref.	ref.
		pandemic	7	0.2	-26	10.0	14.1 (5.6 to 29.9)	9.3 (1.9 to 16.8)
Air-borne/ droplet	Measles	pre-pandemic	58	ref.	ref.	1.2	ref.	ref.
		pandemic	0	0.0	-58	0.0	0.0 (0.0 to 0.0)	-1.2 (-1.4 to -1.1)
Droplet	Rubella	pre-pandemic	32	ref.	ref.	0.7	ref.	ref.
		pandemic	0	0.0	-32	0.0	0.0 (0.0 to 7.7)	-0.7 (-0.8 to -0.6)
Zoonosis	Leptospirosis	pre-pandemic	4	ref.	ref.	0.1	ref.	ref.
		pandemic	0	0.0	-4	0.0	0.0 (0.0 to 73.6)	-0.1 (-0.1 to 0.0)

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CI, confidence interval.

<sup>a</sup> “Pre-pandemic” represents the period April 2016–March 2020, and “pandemic” represents the period April 2020–March 2021; both are based on the time of disease diagnosis.

<sup>b</sup> For the "pre-pandemic" period, the average annual number of cases for the period April 2016–March 2020 is presented.

<sup>c</sup> 184,042,453 and 697,618 were used as the aggregate number of arrivals before and during the pandemic, respectively.

Accepted Version



**Table 3:** Numbers of non-Japanese imported malaria cases, non-Japanese imported malaria cases per 1,000,000 foreign national arrivals, and their ratios and differences during the pandemic compared to the pre-pandemic period, by region, Japan, April 2016–March 2021

Origin of travel	Period <sup>a</sup>	Japanese and non-Japanese		Ratio of the number of cases during the pandemic to the pre-pandemic period	Difference in the number of cases from the pre-pandemic period	Number of cases per million arrivals <sup>c</sup>	Ratio of the number of cases per million arrivals during the pandemic to the pre-pandemic period (95%CI)	Difference in the number of cases per million arrivals from the pre-pandemic period (95%CI)	
		Annual number of cases <sup>b</sup>	Non-Japanese Annual number of cases <sup>b</sup>						
All regions	pre-pandemic	56	31	ref.	ref.	1.1	ref.	ref.	
	pandemic	16	11	0.4	−20	32.2	28.4 (13.8 to 52.6)	31.1 (12.0 to 50.1)	
By region	Africa	pre-pandemic	42	25	ref.	ref.	547.2	ref.	ref.
		pandemic	12	8	0.3	−17	2085.0	3.8 (1.6 to 7.8)	1537.7 (88.9 to 2986.6)
	Asia	pre-pandemic	8	6	ref.	ref.	0.2	ref.	ref.
		pandemic	2	2	0.3	−4	7.3	29.3 (3.3 to 118.5)	7.0 (−3.1 to 17.1)
	Europe	pre-pandemic	0	0	ref.	ref.	0.0	ref.	ref.
		pandemic	0	0	N/A	0	0.0	N/A	0.0 (0.0 to 0.0)

North America	pre-pandemic	0	0	ref.	ref.	0.0	ref.	ref.
	pandemic	0	0	N/A	0	0.0	N/A	0.0 (0.0 to 0.0)
Oceania	pre-pandemic	2	0	ref.	ref.	0.0	ref.	ref.
	pandemic	0	0	N/A	0	0.0	N/A	0.0 (0.0 to 0.0)
South America	pre-pandemic	0	0	ref.	ref.	0.0	ref.	ref.
	pandemic	0	0	N/A	0	0.0	N/A	0.0 (0.0 to 0.0)

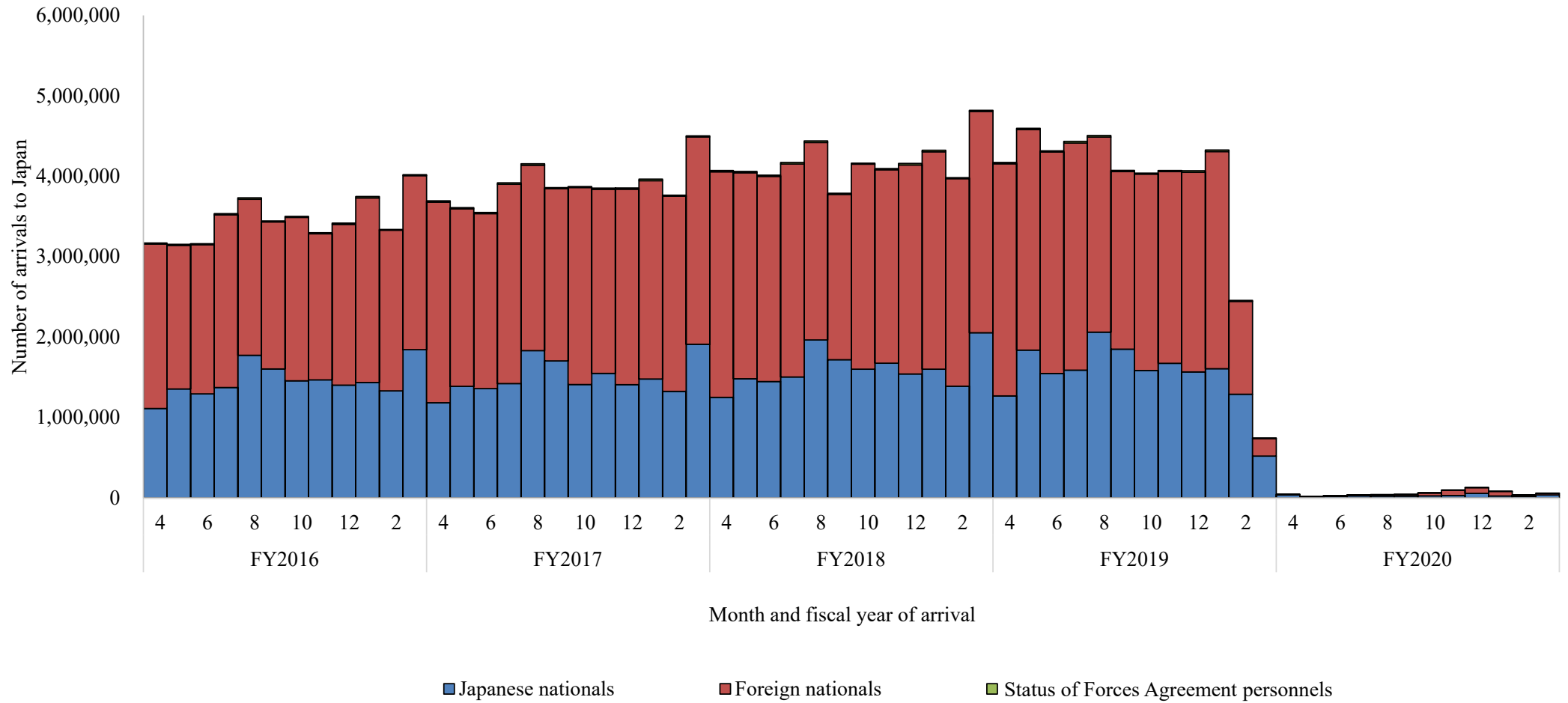
CI, confidence interval; N/A, not applicable.

<sup>a</sup>“Pre-pandemic” represents the period April 2016–March 2020, and “pandemic” represents the period April 2020–March 2021; both are based on the time of disease diagnosis.

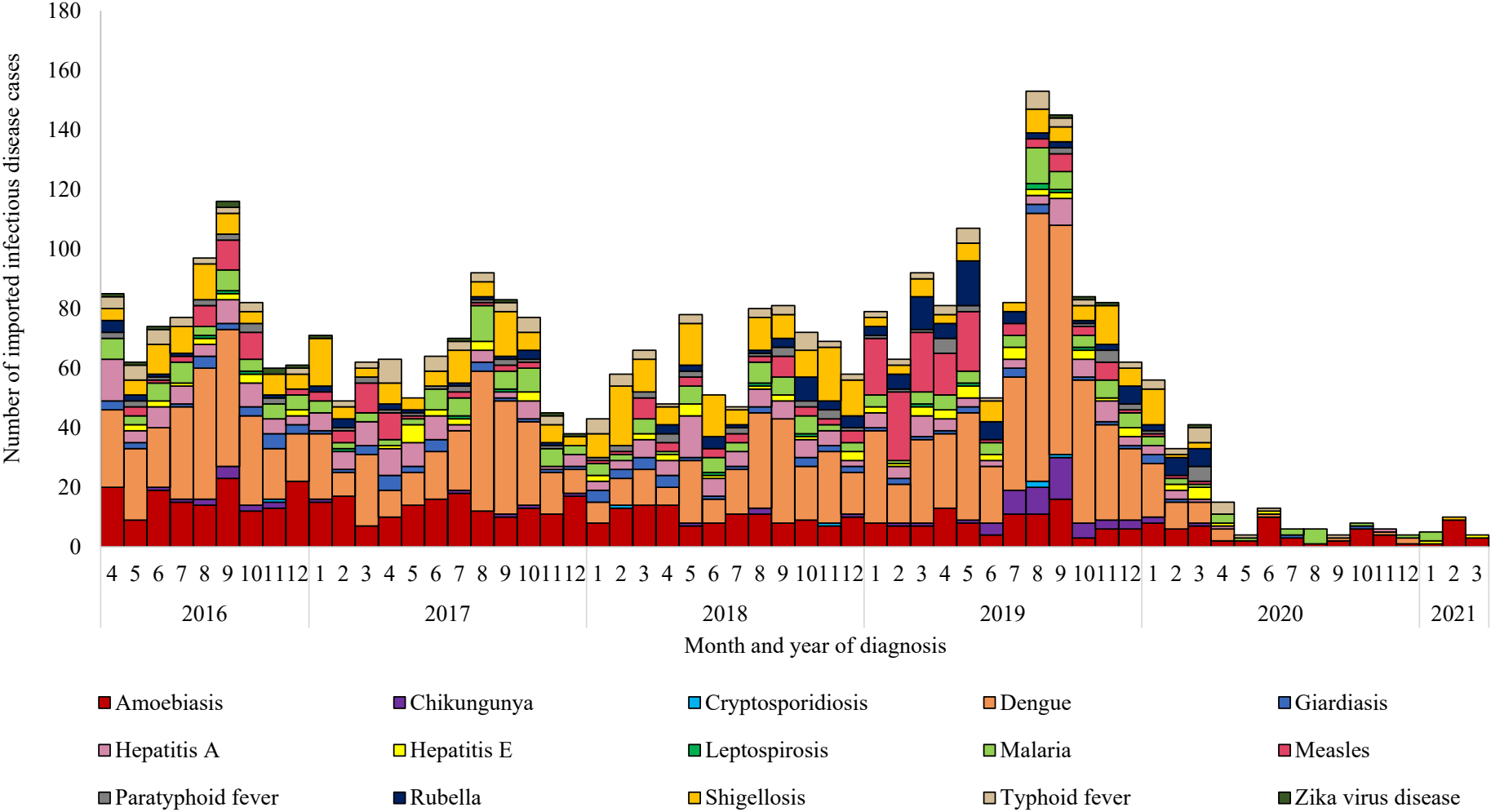
<sup>b</sup> For the "pre-pandemic" period, the average annual number of cases for the period April 2016–March 2020 is presented.

<sup>c</sup> 110,237,475 and 341,479 were used for the analysis in all regions as the aggregate number of foreign nationals arriving in Japan before and during the pandemic, respectively. For regional analysis, the following respective numbers were used: 179,082 and 3,837 for Africa; 92,545,213 and 274,896 for Asia; 6,961,301 and 19,140 for Europe; 1,877,022 and 13,332 for North America; 2,432,713 and 2,164 for Oceania; and 607,341 and 10,343 for South America.

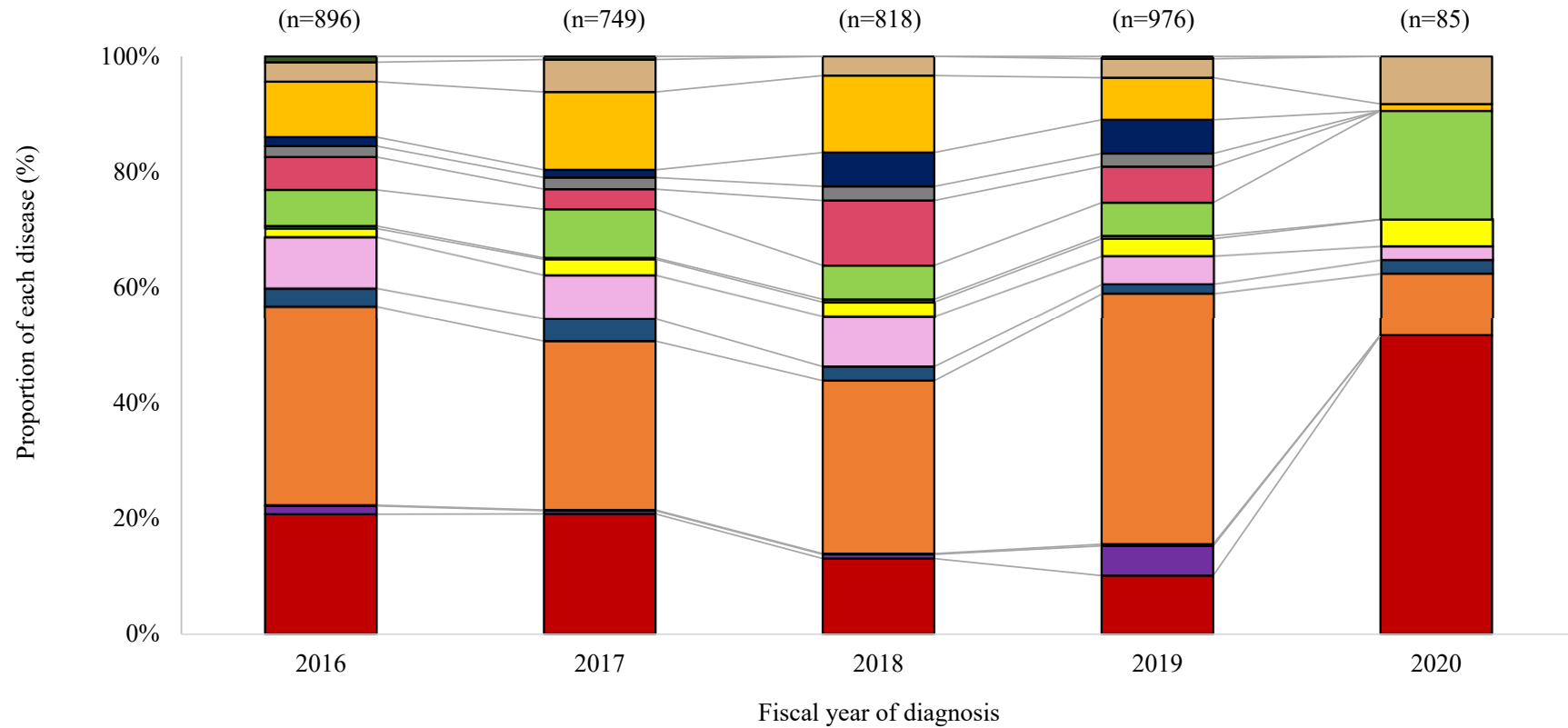
**Figure 1**



**Figure 2**



**Figure 3**



- Amoebiasis
- Chikungunya
- Cryptosporidiosis
- Dengue
- Giardiasis
- Hepatitis A
- Hepatitis E
- Leptospirosis
- Malaria
- Measles
- Paratyphoid fever
- Rubella
- Shigellosis
- Typhoid fever
- Zika virus disease