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Quantifying the success of measles vaccination campaigns in the Rohingya refugee camps

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Abstract

In the wake of the Rohingya population's mass migration from Myanmar, one of the world's largest refugee settlements was constructed in Cox's Bazar, Bangladesh to accommodate nearly 900,000 new refugees. Refugee populations are particularly vulnerable to infectious disease outbreaks due to many population and environmental factors. A large measles outbreak, with over 1700 cases, occurred among the Rohingya population between September and November 2017. Here, we estimate key epidemiological parameters and use a dynamic mathematical model of measles transmission to evaluate the effectiveness of the reactive vaccination campaigns in the refugee camps. We also estimate the potential for subsequent outbreaks under different vaccination coverage scenarios. Our modeling results highlight the success of the vaccination campaigns in rapidly curbing transmission and emphasize the public health importance of maintaining high levels of vaccination in this population, where high birth rates and historically low vaccination coverage rates create suitable conditions for future measles outbreaks.

Keywords

Measles; Vaccination; Mathematical modeling; Rohingya; Bangladesh

1. Background

Since violence in Myanmar intensified in August 2017, nearly 900,000 Rohingya refugees have migrated from the Rakhine State of Myanmar and settled in refugee settlements in Cox's Bazar, Bangladesh (Inter Sector Coordination Group, 2019). Mass population

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Author contributions

TC performed the data analyses and drafted the manuscript. COB and ASM designed the research. All authors revised the manuscript and approved the final version.

Declaration of interests

None.

Appendix A. Supplementary data

Supplementary data associated with this article can be found, in the online version, at <https://doi.org/10.1016/j.epidem.2020.100385>.

displacement has been connected to adverse public health outcomes due to a plethora of risk factors, including heightened insecurity, reduced healthcare access, and inadequate living conditions (Hammer et al., 2018). Despite global gains in reducing the burden of vaccine-preventable infectious diseases (Greenwood, 2014), low vaccination coverage rates and overcrowding conditions in refugee camps often give rise to large-scale outbreaks in these settings (Goma Epidemiology Group, 1995; Porter et al., 1990; Kouadio et al., 2010; Golicha, 2018; Shannon et al., 2019).

Given the crowded conditions in the refugee camps and the low rates of measles immunization among the Rohingya population, a large measles outbreak broke out in the Rohingya refugee camps in Cox's Bazar in late 2017, with over 1700 cumulative suspected cases of measles reported from September to November 2017 (WHO, 2017a). Measles is a highly contagious respiratory infection caused by a paramyxovirus (Centers for Disease Control and Prevention, 2015). Transmission occurs through respiratory droplets or direct contact with secretions from the nose or throat of infected individuals (WHO, 2020). Two vaccination campaigns were conducted in response to the outbreak; the first campaign ran from September 16, 2017 to October 4, 2017 and delivered the measles and rubella (MR) vaccine to 135,519 children aged 6 months to 15 years old (WHO, 2017b). A second vaccination campaign ran from November 18, 2017 to December 2, 2017 and vaccinated 323,940 children in the same age group (WHO, 2017e).

We first evaluate the success of these vaccination campaigns in curbing the measles outbreak in the camps, by quantifying changes in the effective reproductive number (R_e) and estimating the number of cases averted. Our results highlight the success of the reactive vaccination campaigns—a remarkable public health achievement given the very low rates of prior immunization among the Rohingya population.

We find that although these campaigns achieved high coverage, future measles outbreaks are still likely. First, the Rohingya population has historically low vaccination coverage rates; a survey conducted in March 2018 found that 42.9% of children under the age of four had not received even a single dose of an injectable vaccine in Myanmar, and only 2.8% of them had received five or more doses (Bhatia et al., 2018). Second, the birth rate among the refugee population is high, which rapidly replenishes the susceptible population. UNICEF estimates that approximately 60 Rohingya babies were born in Cox's Bazar each day in the nine months following the initial wave of migration in August 2017 (UNICEF, 2018). We use a mathematical model to estimate the potential for subsequent measles outbreaks under different vaccination coverage scenarios. Our findings have important public health implications, as greater understanding of the dynamics of measles in Cox's Bazar can help inform future resource prioritization and vaccination targets.

2. Methods

Daily case data was obtained from the World Health Organization's (WHO) reported cases of measles and rubella in Cox's Bazar from September 6, 2017 to November 25, 2017 (WHO, 2017a). The data include 1708 confirmed and suspected cases of measles and rubella during this time period. All suspected cases are included in this analysis given the disease's

characteristic rash (Centers for Disease Control and Prevention, 2015), which facilitates syndromic reporting of measles. During an early blood sampling of 89 suspected cases of measles during the outbreak, 84% of blood samples tested positive for measles-specific immunoglobulin M (IgM), and only 1% of samples tested positive for rubella-specific IgM (WHO, 2017a). The suspected cases are therefore assumed to be measles rather than rubella. We also assume that all suspected cases occurred among the refugee population given the WHO's finding that only 1% of cases occurred among the host community population (WHO, 2017a).

2.1. Estimation of R_e

We first estimated the daily effective reproductive number for the measles outbreak, in order to estimate the basic reproductive number—a key epidemiological parameter. The basic reproductive number is defined as the expected number of secondary cases from a typical infectious case in a completely susceptible host population (Gay, 2004). The reproductive number quantifies the transmissibility of an infection, thereby informing key attributes such as the herd immunity threshold.

We estimated the effective reproductive number at time t using the approach developed by Wallinga and Teunis (2004), where knowledge of the serial interval distribution and incident cases is used in a likelihood-based estimation procedure that considers pairs of cases. The serial interval distribution of measles used in this analysis was based on a meta-analysis of existing studies of measles, and was assumed to be normally distributed with a mean of 11.9 days and standard deviation of 2.6 days (Vink et al., 2014). Table 1 reports all parameters used in this analysis and their sources. The *EpiEstim* package in *R* was used to estimate the daily effective reproductive number over the course of the outbreak. The 95% credible intervals were estimated using 1000 simulations and 14-day time windows to increase precision in estimates of the daily effective reproductive number (Cori, 2019).

2.2. Mechanistic measles model

Based on our estimated R_e , we estimated the basic reproductive number, R_0 , in order to simulate the dynamics of measles using a mechanistic transmission model. To account for uncertainty at the start of outbreak, we used a range of effective reproductive numbers, R_e to estimate R_0 according to the relationship, where s_0 is the fraction of the population susceptible at the start of the outbreak and R_e is the effective reproductive number at the start of the epidemic. Historical vaccination rates of the Rohingya population in Myanmar and the age distribution of the Rohingya population informed the shape of a triangular distribution assumed for the parameter s_0 (Table 1). The R_e range was chosen based on when the epidemic curve suggests the measles outbreak took off. Effective reproductive numbers between 1.4 and 3.1 correspond to values calculated using 14-day intervals that start between days 6 and 16 into the outbreak. Here, we assume mass-action and that the population is well-mixed (Gay, 2004). Latin Hypercube Sampling (LHS) was used to capture uncertainty around parameter estimates by efficiently sampling from the joint distribution of parameter inputs (Stein, 1987). Using this method, 1000 values of s_0 and R_e were sampled from their assumed probability distributions (Table 1). The maximum estimated basic reproductive number was restricted to 20 based on the ranges for R_0

commonly reported in the literature (Guerra et al., 2017) A gamma distribution was fit to the resultant 1000 estimated values of R_0 . The median and 2.5% and 97.5% percentile of the reproductive number are reported, and a histogram of estimated values is shown to reflect estimates based on the LHS procedure (Supplementary Fig. 1).

A deterministic, Susceptible-Infectious-Recovered (SIR) model, which incorporates birth and death rates, was used to model the size of measles outbreaks in the refugee camps under the scenario in which reactive vaccination had not been implemented in 2017 and under different vaccination scenarios for the next outbreak (Fig. 1). The birth and death rates, which are assumed to be equal, were based on estimates for the number of children born among the Rohingya refugee population in Cox's Bazar in 2018 (Save the Children, 2018; UNFPA, 2017) (Table 1). The model assumes random mixing in the population and does not consider age-dependent heterogeneity in transmission that occurs in other settings (Schenzle, 1984).

The transmission coefficient, β , was estimated according to the relationship $\beta = R_0/ND$ where N is the total population (*i.e.*, $S + I + R$) and D is the average duration of infectiousness, which was assumed to be 9 days (Table 1). A measles-specific death rate was not included in the model since it was assumed to be negligible, as there were only 3 measles-attributed deaths from August 25, 2017 to December 9, 2017 (WHO, 2017d,e). Immunity from both natural infection and vaccination was assumed to be life-long. Two doses of the MMR vaccine have been shown to confer immunity for at least 20 years (Kontio et al., 2012). The model uses one infected case as the initial value for the number of individuals in the infected compartment, $I(0)$. $S(0)$ is either 1) drawn from a triangular distribution[0.05, 0.25, 0.15] assumed for the proportion of the population estimated to be susceptible at the start of the 2017 outbreak, s_0 , or 2) drawn from a uniform distribution[0.05, 0.15] assumed for the proportion of the population estimated to be susceptible at the start of the next outbreak, s_U . The initial value for the recovered compartment was calculated as $R(0) = N - S(0) - I(0)$.

2.2.1. Estimating the impact of vaccination on the 2017 outbreak—The population included in this analysis was the 579,661 Rohingya that migrated to Cox's Bazar after August 25, 2017 and settled in the refugee camps (Inter Sector Coordination Group, 2017). The proportion of the population that was estimated to be susceptible at the start of the outbreak, s_0 , which was also used to estimate R_0 , was estimated by considering prior vaccination rates and the age distribution of the Rohingya population (Table 1). The model runs for 365 days to reflect one year following the start of the outbreak. LHS was used to take into account uncertainty in parameter estimates; 1000 values for each parameter were sampled from the assumed parameter probability distributions (Table 1). A histogram of values for the estimated final size of the 2017 epidemic in the absence of vaccination is shown to reflect uncertainty in estimates (Supplementary Fig. 2).

2.2.2. Modeling the impact of vaccination on the next outbreak—To model the impact of vaccination on subsequent measles epidemics, we assumed a total Rohingya population size of 870,534 as of April 2019 (Inter Sector Coordination Group, 2019). The SIR model described above (Fig. 1) was used to estimate the size of subsequent epidemics

based on varying assumptions for vaccination coverage rates. The model incorporates infection rates during the 2017 outbreak and the vaccination coverage rates of the campaigns conducted in 2017 by reducing the proportion of the population that is susceptible (s) through the formula previously described by Gastañaduy et al. (2018)

$$s = V_C(1 - V_E) + s_U(1 - V_C) \quad (1)$$

where V_C is 1-dose coverage of the Measles, Mumps, and Rubella (MMR) vaccine, V_E is the median effectiveness of 1 dose of MMR vaccine (94%) (McLean et al., 2013), and s_U is the proportion of unvaccinated individuals who are susceptible. This approach assumes vaccination is implemented as one-time interventions between the end of the 2017 outbreak and now. In order to inform the distribution assumed for s_U , binomial probability calculations were first used to estimate the probability of receiving 0, 1, or 2 vaccine doses during the two vaccination campaigns. The number of individuals estimated to have received 0 doses was then added to the number of births estimated to have occurred from the end of the vaccination campaigns to present day (1.5 years later). Table 1 reports all parameters and calculations used as inputs in the mathematical model. LHS was again used to take into account uncertainty in parameter estimates.

For this analysis, the SIR model was run for two years to estimate the impact of vaccination on the next outbreak. The cumulative number of cases over two years was calculated for vaccination coverage rates of 10%, 50%, and 90% of the at-risk population. Cumulative cases were estimated as the cumulative proportion of the population that moves into the infectious compartment in the SIR framework. Distributions of our estimates for the cumulative number of cases at two years under the different vaccination rate scenarios are presented. To contrast these results with the scenario of the next measles outbreak in the absence of vaccination, the same model used to estimate the size of the 2017 outbreak in the absence of vaccination was used with two changes: (1) the total population size was increased to 870,534 (Inter Sector Coordination Group, 2019), and (2) the proportion of the population that was assumed to be susceptible at the start of the outbreak was estimated as s_u instead of s_0 (Table 1) to incorporate the two vaccination campaigns conducted in 2017 and births that have occurred since December 2017.

3. Results

3.1. Quantifying changes to R_e

During the 2017 measles outbreak in Cox's Bazar, the total number of measles cases from September 6, 2017 to November 25, 2017 was 1708. The number of reported measles cases reached a peak of 86 incident cases on November 18, 2017, followed by a decline in cases (Fig. 2A). Current measles transmission appears to be ongoing but limited in the refugee camps, with 323 suspected cases reported from January 1, 2019 to May 4, 2019 (WHO, 2019a).

The daily effective reproductive number ranged from approximately 1.4 (2.5% percentile 1.1 to 97.5% percentile 1.7) at 21 days after the start of the outbreak to a maximum of 4.3 (2.5% percentile 4.0 to 97.5% percentile 4.8) at 34 days, or the week ending on October 10, 2017

(Fig. 2B). The first measles vaccination campaign conducted in response to the outbreak appears to have been highly successful in curbing the transmission of measles, as the daily effective reproductive number declined shortly after the end of the two-week campaign. The second vaccination campaign, meanwhile, was conducted starting in December, when the daily effective reproductive number was already below 1. By the 72nd day of the outbreak, or mid-November, the effective reproductive number had already dipped below 1. After restricting the maximum estimated basic reproductive number to 20, the median basic reproductive number is estimated to be 13.7 with a standard deviation of 3.3 (Supplementary Fig. 1).

3.2. Estimating the impact of vaccination on the 2017 outbreak

If the two vaccination campaigns had not been implemented in response to the 2017 outbreak, our model estimates a median final epidemic size of approximately 79,404 measles cases (standard deviation (sd) of 29,073). This result reflects the high transmissibility of measles, and suggests that after one year from one infectious individual entering the Rohingya population in September 2017, in the absence of interventions, almost all susceptible individuals-or equivalently, approximately 14% of the total population-would have contracted measles. Considering that our model therefore indicates a near 100% attack rate among the susceptible population, the fact that only approximately 1700 cumulative cases occurred from September to the end of November 2017 is a testament to the tremendous success of the vaccination campaigns that were implemented by the WHO, the Bangladesh Ministry of Health, Family and Welfare (MoHFW), and their partners in response to the measles outbreak (WHO, 2017a, 2019b; UNICEF, 2017). Our results suggest that nearly 77,000 cases were averted as a result of the reactive vaccination campaigns conducted in late 2017 (Fig. 3). The swift response of WHO, the MoHFW, and partner organizations, considering the challenging circumstances of accommodating over 650,000 new arrivals by December 2017, is a remarkable public health achievement (WHO, 2017c).

3.3. Modeling the impact of vaccination on the next outbreak

When our model is applied to the current population size under different vaccination scenarios, the model estimates that the median number of measles cases after two years would range from approximately 74,500 cases if 90% of the population is vaccinated to 90,000 cases under the scenario of a 10% vaccination coverage rate (Fig. 4A). These results contrast with the larger median estimate of approximately 93,700 cases, or 11% of the total Rohingya population, predicted for the next measles outbreak in this setting under the scenario of no vaccination (Supplementary Fig. 3). Vaccination therefore has the potential to avert between approximately 4000 and 19,000 cases over two years depending on the coverage rate. Fig. 4B shows estimates in terms of boxplots to reflect the uncertainty in our model's results due to parameter uncertainty.

4. Discussion

Our analysis highlights the immense success of the two large-scale vaccination campaigns in averting measles cases during the 2017 outbreak in Cox's Bazar and supports the importance of maintaining high vaccination coverage rates in the population going forward. Our

findings on the estimate of the basic reproductive number in this setting (median = 13.7; sd = 3.3) are in line with previous estimates reported in the literature (Guerra et al., 2017).

We find that nearly 77,000 cases were averted over the course of one year due to vaccination in the 2017 outbreak. The daily effective reproductive number declined from its maximum value of 4.3 at 34 days after the start of the outbreak to below 1 by the 72nd day of the outbreak. When we apply our model to estimate the impact of vaccination on subsequent outbreaks, depending on the coverage rate in the population, vaccination could avert between 4000 and 19,000 cases over two years. The near 100% attack rate found among susceptibles in the absence of vaccination reinforces the importance of maintaining vaccination as a key public health priority among the Rohingya community in Cox's Bazar.

Our study has several important limitations. The estimate for the basic reproductive number may be biased due to two possible ways in which the serial interval distribution from the literature may overestimate the serial interval distribution in this specific setting. First, it is possible that other intervention measures such as contact tracing and isolation of infectious cases were implemented during the outbreak, which would have the effect of shortening the serial interval for such individuals. However, press articles or WHO reports do not mention these interventions, and it may be reasonable to assume that these interventions were not widely used given their infeasibility in this setting. Secondly, the contraction of the serial interval described by Kenah et al. (2008) in high transmission settings is not considered in this analysis. Instead, a constant serial interval distribution is assumed. For these reasons, the serial interval used in this analysis may represent an overestimation of the actual serial interval in this setting, which would result in an overestimation of the basic reproductive number given the positive correlation between the two measures. Future analyses could examine how the effective reproductive number at time t would change if a hazard-based estimator described by Kenah et al. (2008) was instead used in order to address potential serial interval contraction.

Second, only a limited subset of cases at the beginning of the outbreak was sent for lab confirmation. By assuming that all confirmed and suspected cases were measles cases at the beginning of the outbreak, our methods may overestimate the reproductive number and attack rate. Reporting rates and how these rates changed over time, however, are also not considered in this analysis. One possible scenario is that identification of measles was worse at the start of the epidemic relative to later in the epidemic, and that fewer measles cases were therefore reported in the early phase of the epidemic. If this were the case, the initial case counts may underestimate the true extent of measles transmission at the start of the epidemic, which would affect the calculation for the reproductive number. Assumptions about changes in the reporting rate could be incorporated in future work, such that the reported daily cases are inflated by a different fraction across time, for example using a method similar to that used by White et al. (2009) in their investigation of the 2009 H1N1 pandemic.

Additionally, our mathematical model does not explicitly take into account maternal immunity, which would reduce the proportion of the population susceptible to measles. However, when we tested a wider range for the proportion susceptible to measles in the next

outbreak, our results did not change substantially. Due to lack of data on contact patterns in refugee camps and measles cases by age group, we were also unable to include age-dependent heterogeneities in our model. If transmission was concentrated among children, the epidemic may have taken off more quickly, but the overall impact of age-contact heterogeneities on the epidemic is unclear without additional information. Our mathematical model also makes the assumption that the two vaccination campaigns in 2017 were conducted at random. If officials instead targeted under-reached areas of the camps during the second vaccination campaign, our estimate for s_U , the proportion of the population assumed to be susceptible at the start of the next outbreak, would be too high, and our estimates of the final size of the next outbreak would be an overestimate.

Lastly, our results pertain to a specific outbreak among the Rohingya population in Cox's Bazar and may not generalize to other populations. Importantly, our assumption of homogeneous mixing may not hold in other settings.

Despite these limitations, this analysis provides estimates for the transmissibility of measles in the setting of Cox's Bazar as well as quantifies the impact of vaccination. Starting in December 2017, the WHO and its partners implemented MR vaccination at border check points (WHO, 2017f). From February to December 2018, approximately 29,000 MR doses were delivered to children (WHO, 2018). The WHO currently supports the Ministry of Health in routine measles surveillance (WHO, 2019c). These efforts are highly commendable and should be recognized.

Importantly, however, our findings emphasize the necessity of maintaining ongoing surveillance and of consistently implementing preventive interventions like vaccination. Recently, attention in the refugee camps has been focused primarily on diarrheal diseases in Cox's Bazar (WHO, 2019d). At the same time, according to the WHO's Early Warning, Alert, and Response System (EWARS) in Cox's Bazar, approximately 10 to 20 cases of suspected measles or rubella have been reported to EWARS each week since the start of 2019, signaling ongoing measles transmission (WHO, 2019e).

Against this backdrop of competing public health concerns, it will be important to sustain routine vaccination of vaccine-preventable childhood infections like measles given the high birth rate in the camps and evidence that measles transmission continues to linger. Estimates of the basic reproductive number and size of the next outbreak from this analysis reinforce the public health importance of maintaining the high vaccination coverage rates that are known to be necessary for preventing outbreaks of highly-transmissible infections such as measles.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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References

- Bhatia A, et al., 2018 The Rohingya in Cox's Bazar: when the stateless seek refuge. *Health Hum. Rights* 20, 105–122. [PubMed: 30568406]
- Centers for Disease Control and Prevention, 2015 Pinkbook: Measles. <https://www.cdc.gov/vaccines/pubs/pinkbook/meas.html>.
- Cori A, 2019 EpiEstim Package. <https://www.rdocumentation.org/packages/EpiEstim/versions/1.1-2>.
- Gastañaduy PA, et al., 2018 Impact of public health responses during a measles outbreak in an Amish community in Ohio: modeling the dynamics of transmission. *Am. J. Epidemiol* 187, 2002–2010. [PubMed: 29635277]
- Gay NJ, 2004 The theory of measles elimination: implications for the design of elimination strategies. *J. Infect. Dis* 189, S27–S35. [PubMed: 15106086]
- Golicha Q, 2018 Cholera outbreak in Dadaab Refugee Camp, Kenya–November 2015–June 2016. *MMWR* 67 <https://www.cdc.gov/mmwr/volumes/67/wr/mm6734a4.htm>.
- Goma Epidemiology Group, 1995 Public health impact of Rwandan refugee crisis: what happened in Goma, Zaire, in July, 1994? *Lancet* 345, 339–344. [PubMed: 7646638]
- Greenwood B, 2014 The contribution of vaccination to global health: past, present and future. *Philos. Trans. R. Soc. B: Biol. Sci* 369.
- Guerra FM, et al., 2017 The basic reproduction number (R0) of measles: a systematic review. *Lancet Infect. Dis* 17, e420–e428. [PubMed: 28757186]
- Guzek J, Siddiqui R, White K, Van Leeuwen C, Onus R, 2017 Health Survey in Kutupalong and Balukhali Refugee Settlements, Cox's Bazar, Bangladesh. Tech. Rep. https://www.msf.org/sites/msf.org/files/coxsabazar_healthsurveyreport_dec2017_final1.pdf.
- Hammer CC, Brainard J, Hunter PR, 2018 Risk factors and risk factor cascades for communicable disease outbreaks in complex humanitarian emergencies: a qualitative systematic review. *BMJ Global Health* 3, e000647.
- Inter Sector Coordination Group, 2017 Situation Report: Rohingya Refugee Crisis, Cox's Bazar | 05 12 2017. Tech. Rep. https://reliefweb.int/sites/reliefweb.int/files/resources/171205_iscg_sitrep_one_pager_final.pdf.
- Inter Sector Coordination Group, 2019 Situation Report: Rohingya Refugee Crisis, Cox's Bazar | 30 4 2019. Tech. Rep. https://reliefweb.int/sites/reliefweb.int/files/resources/sitrep_april_2019.pdf.
- Kenah E, Lipsitch M, Robins JM, 2008 Generation interval contraction and epidemic data analysis. *Math. Biosci* 213, 71–79. [PubMed: 18394654]
- Kontio M, Jokinen S, Paunio M, Peltola H, Davidkin I, 2012 Waning antibody levels and avidity: implications for MMR vaccine-induced protection. *J. Infect. Dis* 206, 1542–1548. [PubMed: 22966129]
- Kouadio IK, Kamigaki T, Oshitani H, 2010 Measles outbreaks in displaced populations: a review of transmission, morbidity and mortality associated factors. *BMC Int. Health Human Rights* 10, 5.
- McLean HQ, Fiebelkorn AP, Temte JL, Wallace GS, 2013 Prevention of measles, rubella, congenital rubella syndrome, and mumps, 2013: summary recommendations of the Advisory Committee on Immunization Practices (ACIP). *MMWR Recomm. Rep* 62, 1–34.
- Porter JDH, Gastellu-Etchegorry M, Navarre I, Lungu G, Moren A, 1990 Measles outbreaks in the Mozambican refugee camps in Malawi: the continued need for an effective vaccine. *Int. J. Epidemiol* 19, 1072–1077. [PubMed: 2083992]
- Save the Children, 2018 At Least 48,000 Rohingya Babies Set to be Born in Overcrowded Refugee Camps in Bangladesh in 2018. <https://www.savethechildren.org/us/about-us/media-and-news/2018-press-releases/at-least-48-000-rohingya-babies-set-to-be-born-in-overcrowded-re>.
- Schenzle D, 1984 An age-structured model of pre- and post-vaccination measles transmission. *IMA J. Math. Appl. Med. Biol* 1, 169–191. [PubMed: 6600102]
- Shannon K, et al., 2019 Cholera prevention and control in refugee settings: successes and continued challenges. *PLOS Negl. Trop. Dis* 13, e0007347. [PubMed: 31220084]

- Stein M, 1987 Large sample properties of simulations using Latin hypercube sampling. *Technometrics* 29, 143–151.
- UNFPA, 2017 Republic of Korea Pledges US\$ 500,000 to scale up UNFPA’s response to the Rohingya refugee crisis in Bangladesh. <http://bangladesh.unfpa.org/en/news/republic-korea-pledges-us-500000-scale-unfpa%E2%80%99s-response-rohingya-refugee-crisis-bangladesh>.
- UNICEF, 2017 Bangladesh Steps Up Vaccination for New Rohingya Arrivals as Measles Cases Rise. <https://www.unicef.org/press-releases/bangladesh-steps-vaccinationnew-rohingya-arrivals-measles-cases-rise>.
- UNICEF, 2018 More Than 60 Rohingya Babies Born in Bangladesh Refugee Camps Every Day. <https://www.unicef.org/press-releases/more-60-rohingya-babies-born-bangladesh-refugee-camps-every-day-unicef>.
- Vink MA, Bootsma MCJ, Wallinga J, 2014 Serial intervals of respiratory infectious diseases: a systematic review and analysis. *Am. J. Epidemiol* 180, 865–875. [PubMed: 25294601]
- Wallinga J, Teunis P, 2004 Different epidemic curves for severe acute respiratory syndrome reveal similar impacts of control measures. *Am. J. Epidemiol* 160, 509–516. [PubMed: 15353409]
- White LF, et al., 2009 Estimation of the reproductive number and the serial interval in early phase of the 2009 influenza A/H1N1 pandemic in the USA. *Influenza Other Respir. Viruses* 3, 267–276. [PubMed: 19903209]
- World Health Organization, 2017a Mortality and Morbidity Weekly Bulletin (MMWB) Cox’s Bazar, Bangladesh, Volume No 8: 03 12 2017. Tech. Rep <http://www.searo.who.int/bangladesh/mmwbvol8.pdf>.
- World Health Organization, 2017b Mortality and Morbidity Weekly Bulletin (MMWB) Cox’s Bazar, Bangladesh, Volume No 1: 15 10 2017. Tech. Rep http://www.searo.who.int/bangladesh/mmwb_cxb_vol1_15oct2017_rev.pdf.
- World Health Organization, 2017c Situation Report #08, Refugee Crisis from Myanmar to Bangladesh, 10 19, 2017. Tech. Rep. http://www.searo.who.int/about/administration_structure/hse/20171026_searo_sitrep09.pdf?ua=1.
- World Health Organization, 2017d Mortality and Morbidity Weekly Bulletin (MMWB) Cox’s Bazar, Bangladesh, Volume No 5: 12 11 2017. Tech. Rep <http://www.searo.who.int/mediacentre/emergencies/bangladesh-myanmar/mmwb-vol5-12november2017.pdf?ua=1>.
- World Health Organization, 2017e Mortality and Morbidity Weekly Bulletin (MMWB) Cox’s Bazar, Bangladesh, Volume No 7: 26 11 2017. Tech. Rep <http://www.searo.who.int/bangladesh/mmwbvol7.pdf>.
- World Health Organization, 2017f Weekly Situation Report #7, Rohingya Refugees, 23 12 2017. Tech. Rep. <http://www.searo.who.int/bangladesh/weeklysitrep07cxbban.pdf?ua=1>.
- World Health Organization, 2018 Weekly Situation Report #57, Rohingya Refugees, 27 12 2018. Tech. Rep. <http://www.searo.who.int/bangladesh/weeklysitrep57cxbban.pdf>.
- World Health Organization, 2019a Health Sector Bulletin #9 Rohingya Crisis in Cox’s Bazar, Bangladesh. Tech. Rep <http://www.searo.who.int/bangladesh/health-sector-cxb-bangladesh-no9.pdf>.
- World Health Organization, 2019b Early Warning, Alert and Response System (EWARS) Epidemiological Highlights, Week 52. Tech. Rep <http://www.searo.who.int/bangladesh/ewarsw522018.pdf>.
- World Health Organization, 2019c Bi-Weekly Situation Report #10, Rohingya Refugees, 23 5 2019. Tech. Rep. <http://www.searo.who.int/bangladesh/biweeklysitrep10.pdf?ua=1&ua=1>.
- World Health Organization, 2019d Rohingya Crisis in Cox’s Bazar District, Bangladesh: Health Sector Bulletin #9. Tech. Rep <http://www.searo.who.int/bangladesh/health-sector-cxb-bangladesh-no9.pdf>.
- World Health Organization, 2019e Epidemiological Highlights, Week 16, 2019. Tech. Rep <http://www.searo.who.int/bangladesh/ewarsw162019.pdf>.
- World Health Organization, 2020 Disease and Epidemiology | Measles | Health Topics. <http://www.emro.who.int/health-topics/measles/disease-and-epidemiology.html>.

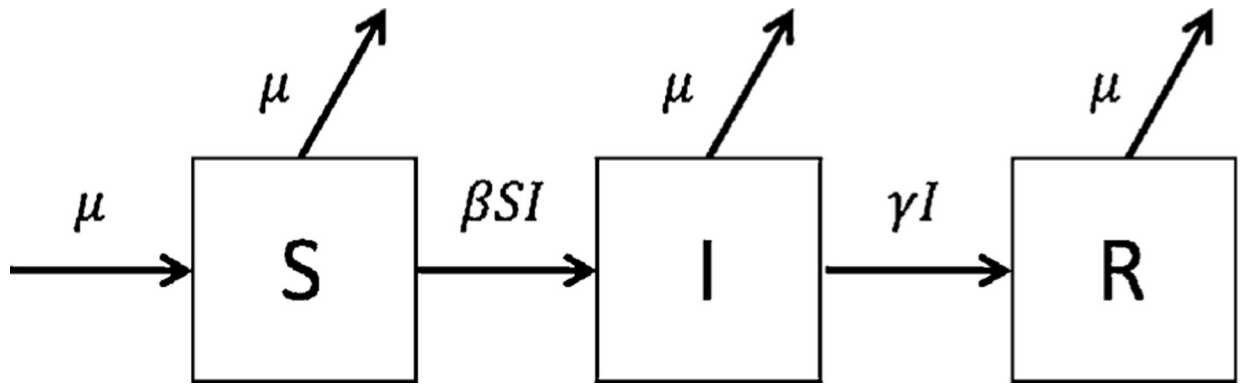


Fig. 1.

Schematic representation of disease states in a model of a measles outbreak in Cox's Bazar. Individuals are either susceptible to measles (S), infectious with measles (I), or recovered from and immune to measles infection (R), either due natural immunity from infection or prior vaccination. β represents the transmission coefficient, γ is the recovery rate, and μ is the birth and death rate. The model starts with one infected (and infectious) measles case (i.e., $I(0) = 1$) The number of recovered individuals at time = 0 was calculated as $R(0) = N - S(0) - I(0)$. The model tracks the number of individuals that move between the compartments each day and is run for one year using the ordinary differential equations (1)–(3).

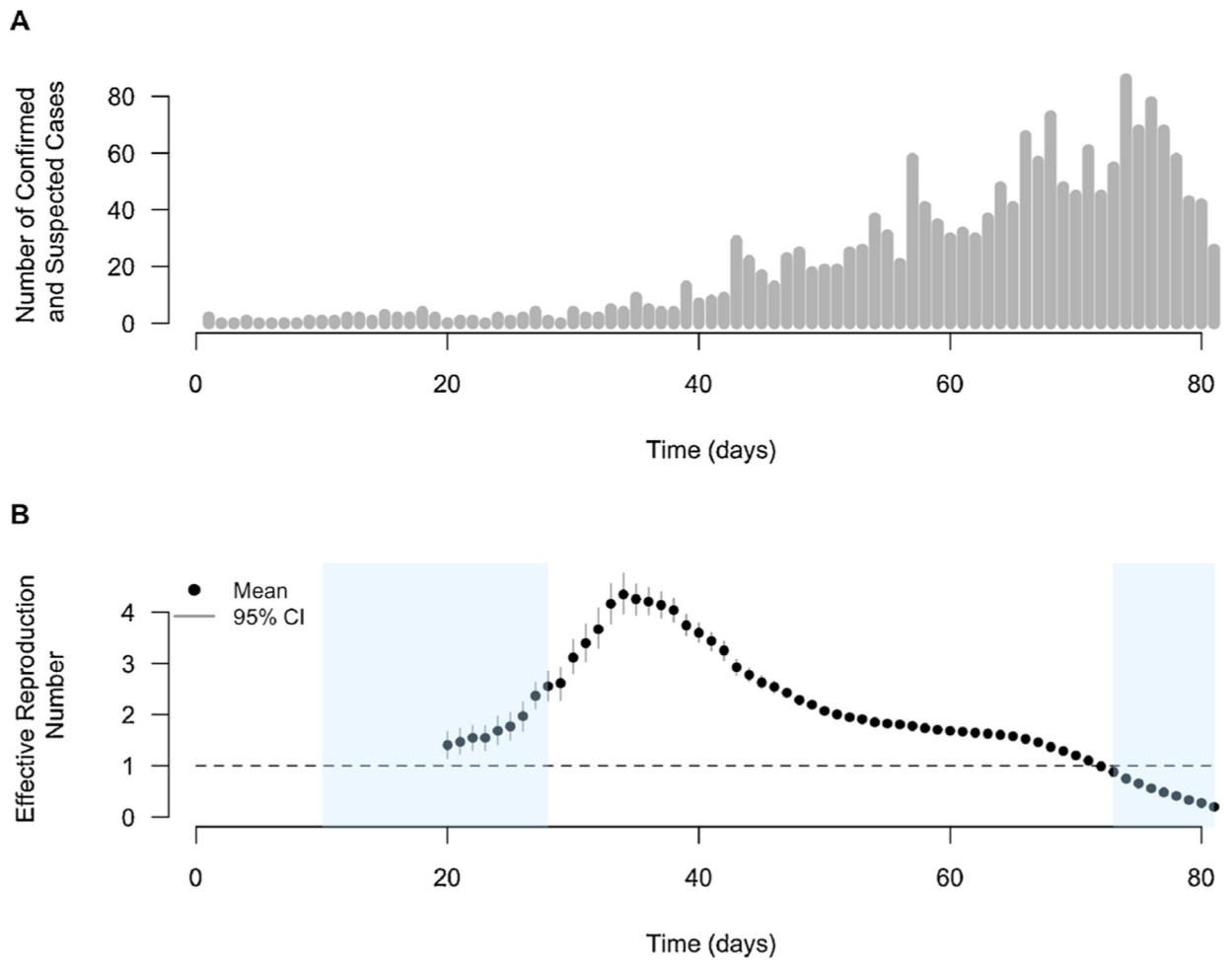


Fig. 2. (A) Confirmed and suspected measles and rubella cases in Cox's Bazar from September 6, 2017 to November 25, 2017. (B) Estimated daily effective reproductive number of measles in Cox's Bazar in the 2017 outbreak using the Wallinga and Teunis method over sliding 14-day windows. Time scale is days since the start of the outbreak on September 6, 2017. Blue shaded areas indicate vaccination campaigns (first campaign from September 16, 2017 to October 4, 2017 and second campaign from November 18, 2017 to December 2, 2017). The 95% credible intervals were estimated using 1000 simulations.

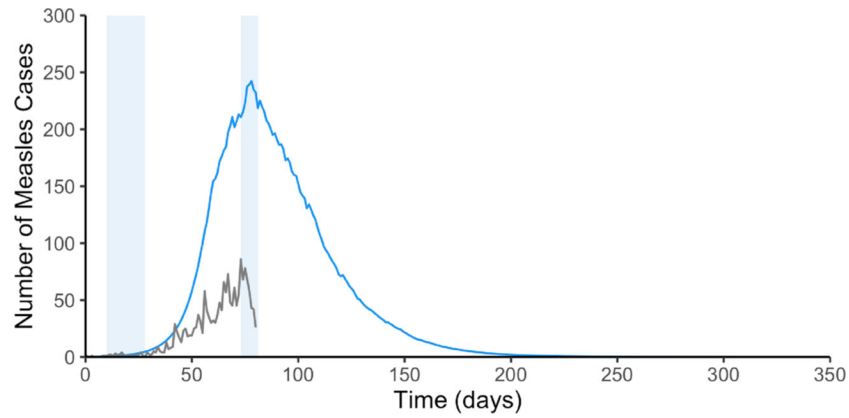


Fig. 3. Confirmed and suspected measles and rubella cases in Cox's Bazar from September 6, 2017 to November 25, 2017 shown in gray. Estimated median number of measles cases in the absence of vaccination over a one-year period beginning on September 6, 2017 shown in blue. The median is based on 1000 parameter combinations using LHS. Light blue shaded areas indicate vaccination campaigns (first campaign from September 16, 2017 to October 4, 2017 and second campaign from November 18, 2017 to December 2, 2017).

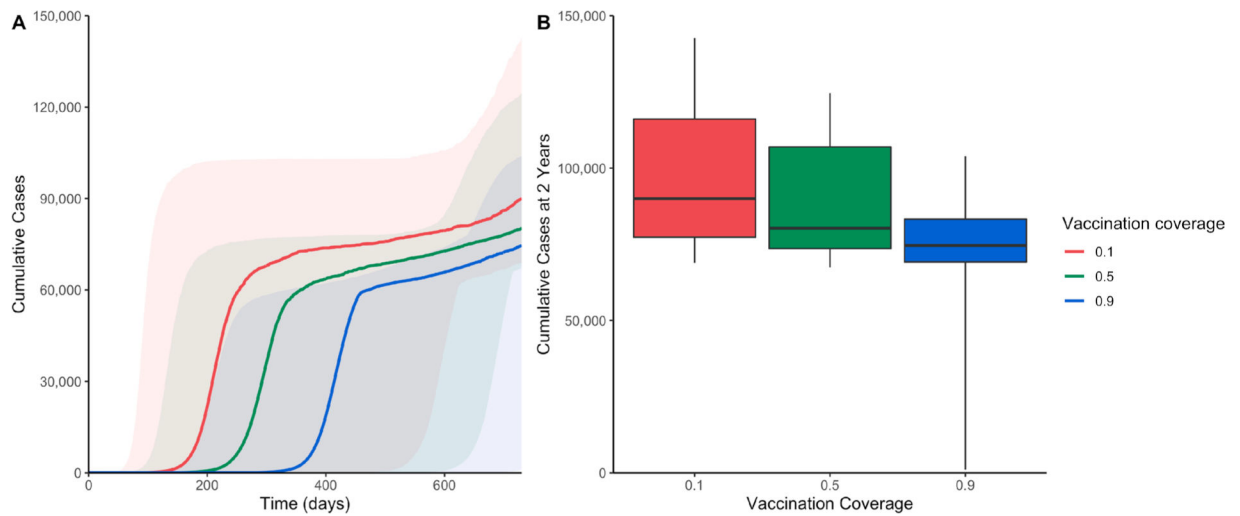


Fig. 4.

(A) Cumulative number of measles cases over time under varying scenarios of vaccination coverage rates. Shaded areas represent the range between estimates' 10% and 90% percentiles from LHS sampling of 1000 parameter combinations. (B) Boxplots of the cumulative number of measles cases at 2 years (730 days) using 1000 LHS parameter combinations under the same vaccination coverage rate scenarios. Maximum and minimum values represent 90% and 10% percentiles, respectively.

Table 1

Summary of input parameters.

Parameter	Value/distribution	Notes	Source
<i>Estimation of R_e</i>			
Measles serial interval distribution	$N(11.9, 2.6^2)$	An estimated serial interval distribution reported from a household transmission study from Providence, R.I. by Chapin (1925)	Vink et al. (2014)
Susceptible population at start of 2017 outbreak (%), S_0	Triangle(0.05,0.25,0.15)	Given the low vaccination rates in Myanmar (Bhatia et al., 2018), we assumed that most Rohingya adults were already immune to measles due to a resultant low average age of first infection. This assumption seems reasonable given that 83% of suspected cases were among children <5 years old, and 96% of suspected cases were among children <15 years old during the 2017 outbreak (WHO, 2017a). Approximately 50% of the Rohingya population in Cox's Bazar is <15 years old (Bhatia et al., 2018; Guzek et al., 2017). 72% of measles cases in the 2017 outbreak as of November 18, 2017 had no history of measles vaccination (WHO, 2017a).	Bhatia et al. (2018), WHO (2017a), MSF (Guzek et al., 2017)
<i>Mechanistic measles model</i>			
Basic reproductive number, R_0	Gamma(shape=16.2, rate=1.2)	Fit gamma distribution to histogram of R_0 values, which were estimated in the analysis using the relationship $R_e = R_0 * S_0$	Estimated in analysis
Rohingya population size, December 2017	579,661	The total influx of Rohingya into Cox's Bazar from August 25, 2017 to December 5, 2017, excluding those that settled in the host community	ISCC (Inter Sector Coordination Group, 2017)
Rohingya population size, April 2019	870,534	The number of Rohingya refugees identified in the refugee camps as of April 2019, excluding those registered before August 31, 2017	ISCC (Inter Sector Coordination Group, 2019)
Birth rate (/10,000/day), μ	1.45	Save the Children estimated that 48,000 Rohingya children would be born in Cox's Bazar in 2018 (Save the Children, 2018), which is in line with UNEFPA's estimate from December 2017 that 10,000 Rohingya women would give birth from January to March 2018 (UNFPA, 2017).	Save the Children (2018), UNFPA (2017)
Death rate (/10,000/day), μ	1.45	Assumed to be the same as the birth rate	Assumption
Recovery rate (days ⁻¹), γ	0.11	1/average duration of infectiousness (9 days)	McLean et al. (2013)
Transmission coefficient (days ⁻¹), β	-	R_0 /(average duration of infectiousness * population size)	Estimated in analysis
Susceptible population at start of next outbreak (%), S_u	Uniform(0.05,0.15)	<ul style="list-style-type: none"> 349,603 children <15 years old were targeted for second vaccination campaign (WHO, 2017a). If 135,519 children were vaccinated in the first round and 323,940 in the second round, 38.8% (135,519/349,603) and 92.7% (323,940/349,603) of children were vaccinated in these campaigns, respectively Use binomial calculations to estimate probability of child receiving 0, 1, and 2 MR doses as 4.5%, 59.6% and 35.9%, respectively If 4.5% of children received 0 doses, approximately 15,715 children were not vaccinated by the end of the vaccination campaigns Add estimated 72,000 births (48,000*1.5) that have occurred since the vaccination campaigns; annual estimate of 48,000 is the same estimate from Save the Children (2018) that informs the birth rate 	Assumption

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Parameter	Value/distribution	Notes	Source
		•	Estimate current proportion of children susceptible as $(15, 715 + 72, 000)/870, 534 = 0.10$ to inform the middle of the uniform distribution range