

Two Decades of Endemic Dengue in Bangladesh (2000-2022): Trends, Seasonality, and Impact of Temperature and Rainfall Patterns on Transmission Dynamics

Journal:	Journal of Medical Entomology
Manuscript ID	Draft
Manuscript Type:	Research
Date Submitted by the Author:	n/a
Complete List of Authors:	Hasan, Mohammad Nayeem; Shahjalal University of Science and Technology, Department of Statistics Khalil, Ibrahim; Government of the People's Republic of Bangladesh Ministry of Fisheries and Livestock, Department of Livestock Services Chowdhury, Muhammad Abdul Baker; University of Florida College of Medicine, Department of Neurosurgery Rahman, Mahbubur; University of London, Institute of Epidemiology; Ministry of Health and Family Welfare, Institute of Epidemiology Asaduzzaman, Md; Staffordshire University, Technologies, and Arts Billah, Masum; Staffordshire University, Technologies, and Arts Banu, Laila Arjuman; Bangabandhu Sheikh Mujib Medical University, Department of Anatomy Alam, Mahbub-UI; International Centre for Diarrhoeal Disease Research Bangladesh, Environmental Intervention Unit Ahsan, Atik; International Centre for Diarrhoeal Disease Research Bangladesh, Environmental Intervention Unit Traore, Tieble; WHO, Emergency Preparedness and Response Programme Uddin, Md. Jamal; Shahjalal University of Science and Technology, Department of Statistics; Daffodil International University, Department of General Educational and Development Galizi, Roberto; Keele University, School of Life Sciences Russo, Ilaria; Keele University, School of Medicine Zumla, Alimuddin; University College London, Division of Infection and Immunity; University College London Hospitals NHS Foundation Trust, Division of Infection and Immunity Haider, Najmul; Keele University, School of Life Sciences
 	Modeling/GIS, Risk Assessment, Economic Impact
Organism Keywords:	Aedes aegypti
Field Keywords:	Public Health Entomology

SCHOLARONE™ Manuscripts

- 1 Two Decades of Endemic Dengue in Bangladesh (2000-2022): Trends,
- 2 Seasonality, and Impact of Temperature and Rainfall Patterns on Transmission
- 3 **Dynamics**

- 5 Mohammad Nayeem Hasan¹, Ibrahim Khalil ², Muhammad Abdul Baker Chowdhury ³,
- 6 Mahbubur Rahman 4,5, Md Asaduzzaman 6, Masum Billah 6, Laila Arjuman Banu 7,
- 7 Mahbub-ul Alam⁸, Atik Ahsan ⁸, Tieble Traore ⁹, Md. Jamal Uddin^{1,10} Roberto Galizi ¹¹,
- 8 Ilaria Russo¹², Alimuddin Zumla¹³, Najmul Haider^{11*}

9

- ¹Department of Statistics, Shahjalal University of Science and Technology, Sylhet 3114,
- Bangladesh (MNH: nayeem5847@gmail.com, MJU: jamal-sta@sust.edu)

12

- ² Department of Livestock Services, Ministry of Fisheries and Livestock, Bangladesh, Dhaka,
- 14 Bangladesh (IK: <u>dribrahim.dls@gmail.com</u>)

15

- ³ Department of Neurosurgery, University of Florida College of Medicine, Gainesville, Florida
- 17 FL 32610, USA (MABC: <u>baker.chowdhury@neurosurgery.ufl.edu</u>)

18

- ⁴ The Royal Veterinary College, University of London, Hawkshead Lane, North Mymms,
- Hatfield, Hertfordshire, United Kingdom (Email: MR: <u>dr_mahbub@yahoo.com</u>)

21

- ⁵ Institute of Epidemiology, Disease Control and Research (IEDCR), Ministry of Health and
- Family Welfare, Mohakhali, Dhaka, Bangladesh (MR: dr mahbub@yahoo.com)

24

- ⁶ School of Digital, Technologies, and Arts, Staffordshire University, Staffordshire, UK (MB:
- 26 <u>masum.billah@staffs.ac.uk</u>, MA: <u>Md.Asaduzzaman@staffs.ac.uk</u>)

27

- ⁷ Department of Anatomy, Bangabandhu Sheik Mujib Medical University, Dhaka, Bangladesh
- 29 (dr.lailabanu@gmail.com)

53

⁸ Environmental Intervention Unit, International Centre for Diarrhoeal Diseases Research, 30 Bangladesh (ICDDR,B), Dhaka-1212, Bangladesh (MUA: mahbubalam@icddrb.org, AA: 31 32 atik.ahsan@icddrb.org) 33 ⁹ Emergency Preparedness and Response Programme, WHO Regional Office for Africa, Dakar 34 Hub, Daker, Senegal (TT: traoret@who.int) 35 36 ¹⁰ Department of General Educational and Development, Daffodil International University, 37 Dhaka, Bangladesh (MJU: jamal-sta@sust.edu) 38 39 ¹¹ School of Life Sciences, Faculty of Natural Sciences, Keele University, Keele, Staffordshire, 40 United Kingdom, ST5 5BG (NH: n.haider@keele.ac.uk, RG: r.galizi@keele.ac.uk). 41 42 ¹² School of Medicine, Faculty of Medicine and Health Sciences, Keele University, 43 Staffordshire, ST5 5BG, United Kingdom (IR: i.russo@keele.ac.uk) 44 45 ¹³ Division of Infection and Immunity, Centre for Clinical Microbiology, University College 46 London and NIHR-BRC, University College London Hospitals, London, United Kingdom (AZ: 47 a.zumla@ucl.ac.uk). 48 49 *Corresponding author (NH): Dr Najmul Haider, School of Life Sciences, Keele University, 50 Huxley Building, Room 122, Keele, Staffordshire, ST5 5BG, United Kingdom, Email: 51

n.haider@keele.ac.uk, Phone: (+44) 01782 734414

54	Abstract:
55	Background: The objectives of this study were to compare the dengue virus (DENV) infection,
56	deaths, case-fatality ratio, and meteorological parameters between the first and the recent decade
57	(2000-2010 vs. 2011-2022) and to understand the trends, seasonality, and impact of change of
58	temperature and rainfall pattern on transmission dynamics of dengue in Bangladesh
59	
60	Methods: For the period 2000-2022, dengue cases and death data from Bangladesh's Ministry of
61	Health and Family Welfare's website, and meteorological data from the Bangladesh Meteorological
62	Department were analyzed. A Poisson regression model was performed to identify the impact of
63	meteorological parameters on the monthly dengue incidence. A forecast of dengue cases was
64	performed using an autoregressive integrated moving average model.
65	
66	Results: Over the past 22 years, a total of 244,246 dengue cases were reported including 849 deaths
67	(Case fatality ratio [CFR] =0.34%). The mean annual number of dengue cases increased eight-fold
68	during the second decade, with 2216 cases during 2000-2010 vs. 18,321 cases during 2011-2022.
69	The mean annual deaths have doubled (21 vs. 46), but the overall CFR has decreased to one-third
70	(0.69% vs 0.23%). The annual temperature increased by 0.49 $^{\circ}$ C, and rainfall decreased by 314 mm
71	Monthly mean temperature (Incidence risk ratio [IRR]: 1.26), first-lagged rainfall (IRR: 1.08), and
72	second-lagged rainfall (IRR: 1.17) were significantly associated with monthly dengue incidence.
73	
74	Conclusions: The increased local temperature and unusual rainfall might have contributed to the
75	increased incidence of DENV infection in Bangladesh. Community engagement, vector control,
76	and destruction of mosquito habitats are key to controlling dengue.
77	
78	Keywords: Dengue, Bangladesh, Climate change, Temperature, Rainfall
79	
80	

82

83

84

85

86

87

88

89

90

91

92

93

94

95

96

97

98

99

100

101

102

103

Introduction:

Dengue fever is a mosquito-borne disease (MVD) caused by four distinct serotypes of the dengue virus (DENV) of the Flaviviridae family (WHO 2009). DENV is transmitted to humans by the bites of the female Aedes species mosquitoes including Ae. aegypti and Ae. albopictus (WHO 2009, CDC 2019). DENV is endemic in over 125 countries of the world and the number of cases globally reported to WHO continues to increase every year (WHO 2023a). Annually, an estimated 390 million dengue infections are recorded across the world, including 96 million clinical cases making DENV as one of the most important vector-borne diseases (VBDs) in the world (Murray et al. 2013, WHO 2023b). Most infections (>80%) are self-limiting with no or mild clinical manifestation resulting in lifelong immunity for serotype (WHO-Bangladesh 2022). However, infections with different serotypes, known as secondary or tertiary dengue infection, may result in severe dengue with a higher case-fatality ratio(Teo et al. 2023). Currently, South and Southeast Asia is considered to be the hotspot of DENV infection with more than 50% of cases recorded in the regions (WHO South-East Asia 2023). The first official DENV outbreak in Bangladesh was reported in 2000, and since then, dengue has become endemic in the country posing a significant health challenge (Sharmin et al. 2015). Over the past few years, the number of dengue cases has been steadily increasing with significant seasonal and regional variations. Analysis of data from 2000 to 2017 revealed that almost half of the dengue cases occurred during the monsoon season (May-August) and the post-monsoon season (September-December) (Mutsuddy et al. 2019). However, a shift in seasonal patterns has been observed since 2014, with dengue cases being reported during the pre-and-post monsoon season (Mutsuddy et al. 2019). During 2015-2017, the number of dengue cases during the pre-monsoon

season was more than seven times higher compared to the previous 14 years (Mutsuddy et al. 2019). The annual incidence of DENV infection started to increase sharply after introduction of the serotype DENV-3 in 2019 in Bangladesh (Haider et al. 2021).

Climate change including changes in precipitation, temperature, and humidity, as well as rapid unplanned urbanization, were identified as strong predictors of an ecological imbalance that has led to an increase in dengue cases in Bangladesh (Mutsuddy et al. 2019). This suggests that the dengue transmission season could eventually extend year-round, with a higher chance of outbreaks occurring at any time of the year. Identifying trends and seasonality in dengue cases can aid health authorities and relevant public and private administrations in effectively allocating resources to control the spread of the DENV through vector control. The objectives of our study were to: i) compare the annual and monthly cases in the first [2000-2010] and recent decade [2011-2022], ii) identify the trend and seasonality of dengue cases, iii) quantify the impact of climatic parameters for the monthly incidence of dengue cases in the country and iv) forecast the annual incidence of dengue cases for next decade.

Methods:

Data sources:

The data on the number of reported dengue-infected people have been extracted from the Directorate General of Health Services (DGHS)'s website from January 2000 to December 2022 (DGHS 2023). We used the definition of dengue cases used by the Ministry of Health and Family Welfare, Bangladesh, which was discussed in our earlier article (Ahsan et al. 2021). We collected three-hourly temperature and daily rainfall data from Bangladesh Meteorological

Department (BMD) over the period 2000–2022 (BMD 2023) for the meteorological station located in Mirpur, Dhaka.

Variables

The monthly number of dengue cases was used as the primary outcome variable. Two climatic variables- temperature and rainfall are used as the covariates for the regression analysis. In addition, two lagged variables rainfall in lag 1 and lag 2 have also been used as the predictors for the incidence of monthly dengue cases to capture the actual impact of those meteorological parameters. We also used monthly mortality data for comparison between two decades.

Statistical analysis

We analyzed the monthly dengue incidence and meteorological data for the period of 2000-2022. In the first stage, descriptive analysis was conducted to determine the characteristics of confirmed dengue cases and deaths with mean, and standard deviation in each year and each month for the entire period. Then, we compared dengue cases, deaths, and weather parameters in two decades (2000-2010 and 2011-2022) using paired sample t-test. Next, we calculated the monthly growth factor (GF) of dengue cases by dividing the number of dengue cases reported in each month by the number of dengue cases reported in the previous month and repeating this process for each month from 2000 to 2022 (Haider et al. 2021). The formula for the growth factor can be given by

147
$$GF_t = \frac{N_{t+1} + 1}{N_t + 1}$$

where N_t indicates the number of dengue cases in t^{th} month. To avoid the occurrence of zeros in some months, we added 1 to the total number of cases for each month. This allows us to obtain a real-valued measurement of the GF for the above equation. The distribution of GF was skewed; therefore, we used the first natural log transformation before the data was further examined. However, we have also performed a reverse transformation of the log (GF) values by exponentiating values to convert them to the original scale for ease of interpretation(Haider et al. 2021).

We performed forecasting using the autoregressive integrated moving average (ARIMA) model. The ARIMA model is a data-driven, exploratory strategy that enables us to fit a suitable model and forecast values. The ARIMA model consists of autoregressive (p) terms, differencing (d) terms, and moving average (q) operations, and it is denoted as ARIMA (p, d, q) (Kumar and Susan 2020). To select the appropriate autoregressive and moving average orders, the autocorrelation function (ACF) and partial autocorrelation function (PACF) were examined. Additionally, the differencing parameter, represented by "d," indicates the number of times the time series is different to achieve stationarity. An ARIMA (p, d, q) process refers to an autoregressive moving average (ARMA) model that has been differenced "d" times to obtain stationarity (Hasan et al. 2021). By removing high-frequency noise from the data, the model discovers local patterns by assuming that the time series values are linearly related. We also conducted a Mann-Kendall (M-K) trend analysis to determine possible upward or downward trends (Yue and Pilon 2004). We also performed the Sen's slope test to assess variations in annual dengue cases and deaths(Sen 1968).

We, then used a time series count generalized linear model (GLM), more specifically, a timeseries Poisson regression model to determine whether the climatic factors were associated with
the dengue cases over time (Sumi et al. 2021). The non-normality, heteroscedasticity, and nonlinearity that characterize count data can be fitted easily using GLMs. The time-series
observations may possess autocorrelation and they might be nonnegative integers, and thus GLM
is useful in overcoming both issues [20, 21, 22]. Monthly dengue cases were utilized as the
outcome variable in this model, along with data from the Bangladesh Meteorological Department
(BMD) on temperature and rainfall. To capture the actual impact of rainfall on dengue incidence
across time, we additionally employed two lagged variables of meteorological elements, mainly
rainfall in lag 1 and 2. After eliminating predictors with a higher multicollinear relationship, we
have arrived at average temperature, rainfall (in lag 1), and rainfall (in lag 2) as the final set of
predictors for the monthly dengue incidence in Bangladesh. We used the statistical program
RStudio, version 3.5.2.2 for the analyses (R Core Team 2022).

Results:

Between 2000 and 2022, Bangladesh reported a total of 244,246 dengue cases with an annual mean of 10,161 cases (± standard deviation [SD]=23,971) including 849 fatal outcomes indicating a case-fatality ratio (CFR) of 0.34%. Between 2000 to 2010, the mean annual number of dengue cases was 2,216 (±2,123) which has increased over eight folds compared to the following decade (2011-2022) at 18,321 (±31,778) (**Table 1**). Between these two periods, the mean number of annual deaths due to DENV infection has increased by 2.2 times (21.18 cases vs

193	46.58 cases). However, the CFR of DENV infection has decreased to almost one-third between
194	two decades (0.69 vs 0.23) (Table 1).
195	
196	The highest monthly average number of cases was recorded in August (n=3407 cases) and the
197	lowest was in March (n=6.7 cases) (Fig 1). The highest number of annual cases was reported in
198	2019 with 101,354. The highest number of deaths was recorded in 2022 with 281 deaths, which
199	was 35% of total deaths recorded in the past 23 years in Bangladesh (Fig 1). Most of the dengue-
200	related deaths were recorded after 2018, with more than 65% (n=550) deaths recorded during
201	this time (Fig 1).
202	
203	The average annual temperature was 26.35 °C (SD=0.49) during the first decade (2000-2010)
204	and 26.84 °C (SD=) during the recent decade (2011-2022) (Table 1). The increase of 0.49 ° C
205	temperature was equivalent to 4292 degree-hour/year of heat (365 days X 24 hours X 0.49 $^{\circ}$ C).
206	The annual rainfall has decreased by 314 mm between two decades (2078.66 mm vs. 1764.50
207	mm) (Table 1), of which 308 mm decreased during the monsoon (July-October) season and only
208	6 mm decreased during the non-monsoon period. Compared to the first decade (2000-2010), an
209	unusually higher amount of monthly precipitation was observed in the second decade (2011-
210	2022) with most of the months recording extreme rainfall (more than 3 rd quantile value of
211	monthly rainfall for the decade) shown as an outlier of the box plot (Fig 2).
212	
213	The overall mean GF from month to month was 1.37 (SD=0.86). However, in four months
214	(April-July), the monthly GF was above one (lower 95% confidence interval >1), while for the
215	rest of the years, the monthly GF was less than 1 (95% confidence interval crossed 1). More

217

218

219

220

221

222

223

224

225

226

227

228

229

230

231

232

233

234

235

DENV infection in Bangladesh (Fig 4).

than 77% (71/92) of months between April and July for the period 2000–2022 had mean monthly GF > 1 compared to only 16% (30/184) of months between August and March of the same period. June had the highest GF with a mean value of 3.47 indicating that cases would be more than three times higher in the next month (July). The lowest GF was recorded in December with a mean of 0.54 (95% CI: 0.40 to 0.69) indicating that cases in January would be halved compared to the number of cases recorded in December (Fig 3). In the M-K trend analysis, we found a positive trend of reported dengue cases (p < 0.001 and tau = 0.26). In Sen's slope test, the slope was 171.67 (95% CI: -46 to 687) indicating an upward trend in upcoming months (**Table** 2). In the GLM, the estimated effect of each variable is presented as the incidence risk ratio (IRR). The model suggests that dengue cases would rise by 26% for a one-degree centigrade (°C) temperature increase. For each additional centimeter (cm) of rainfall in the first lagged month, the number of dengue cases increased by 8% (IRR= 1.08 [95% CI: 1.08-1.09]), and in the second lagged month increase the cases by 17% [IRR=1. 17 (95% CI: 1. 17 -1.18)] (**Table 3**). In the ARIMA model, we detected an increasing trend for the first few years, which then started to decline. However, a stiff rise in cases was observed after 2018 except for 2020 (the first year

236

237

of the Covid-19 pandemic). The forecasted value showed a continuously increasing trend of

Discussions:

Dengue is currently a worrying and important public health challenge for Bangladesh. Our analysis showed that the number of DENV infection has increased eight times and deaths have doubled, and the CFR dropped to one-third between the first and second decade of this century in Bangladesh. Between these periods, the annual temperature increased by 0.49 °C, and annual rainfall decreased by 314 mm, despite changes in rainfall patterns with unusually early or late rainfall outside the typical monsoon season in Bangladesh (July-October) (Haider et al. 2014). The monthly growth factor remains above one significantly for four months (April to July) which overlaps the hot and humid period of the year. Monthly mean temperature, monthly first-lagged rainfall, and second-lagged rainfall played a critical role in monthly dengue incidence in Bangladesh.

The increase of 0.49 °C temperature adds approximately 4292-degree-hours equivalent heat per year in the country. This additional heat would favor VBD transmission. For dengue virus transmission, approximately 305-degree-hours equivalent heat is needed to accomplish the extrinsic incubation period in Aedes mosquitoes at 26° C (Focks et al. 1995). Thus, the additional 0.49°C temperature will add the burden of more than 14 generations of infectious mosquitoes in the environment of Bangladesh. An 8-fold increase in dengue cases is an indication of such changes in temperature in the country. Our model identified a significant role of monthly mean temperature with an additional 1 °C temperature increasing the monthly cases by 26%. Earlier studies showed that for every 1 °C increase in temperature, dengue cases increased by 61% in Australia, 12-22% in Cambodia, 5% in Vietnam, and 2.6% in Mexico (Soneja et al. 2021).

262

263

264

265

266

267

268

269

270

271

272

273

274

275

276

277

278

279

280

Rainfall facilitates mosquito breeding and plays an important role in mosquito-borne disease transmission. Although we found a 15% reduction in annual rainfall in the recent decade from the immediate past decade, we detected an increase in unusual high rainfall in pre-and-post monsoon season allowing an extended season for mosquitoes and other arthropod vectors. Our model showed that both the first and the second lagged month's rainfall increased monthly cases by 8% and 17%, respectively. These findings are consistent with earlier studies in Bangladesh that showed that peak dengue cases occurred two months after the peak rainfall(Salje et al. 2016) or an additional rainy day per month increased dengue cases by 6% in the succeeding month (Rahman et al. 2020). Similar findings were reported in Vietnam with the dengue incidence being associated with both first and second-lagged months (Cuong et al. 2011). In Timor-Leste, a 47% increase in dengue incidence was recorded with an additional 1 mm seasonal rainfall increase (Wangdi et al. 2018). These findings are biologically plausible as rainfall allows approximately two generations of dengue cases over a month. A generation interval is a time difference between a primary human infection and a second human infection originating from the first human case through two bites of the mosquitoes (Siraj et al. 2017). To accomplish a generation interval the virus and mosquito undergo several phases including intrinsic incubation period in humans, human-mosquito transmission (first bite), extrinsic incubation period in mosquitoes, blood meal digestion period, and finally mosquitoes-to-human transmission (2nd bite) (Siraj et al. 2017). Ideally, for DENV, the generation interval completes at around 16 days at 28-32 °C (Siraj et al. 2017).

282

281

Bangladesh's dengue season is characterized by hot and wet periods running between June to August. This is the period with the highest amount of rainfall in the country facilitating Aedes mosquito breeding in the country (Haider et al. 2023). The monthly mean growth factor above 1 for April – June indicates that for each of these months, the incidence of dengue cases will surpass the current month. Thus, we suggest starting vector control intervention in April in Bangladesh.

Globally and regionally in South and Southeast Asia, dengue cases are increasing. DENV infection increased by more than 46% between 2015 and 2019 in region (WHO South-East Asia 2023). In 2023, up until 31 May, a total of 1515,460 DENV infections were recorded in Brazil with 387 deaths(European CDC 2023). In Malaysia, a total of 43,619 DENV infections have been recorded by 21 May 2023(European CDC 2023). We found an increasing trend of DENV infection in Bangladesh. This increasing trend was much stiffer after the serotype DENV-3 was introduced in the country in 2018 (Ahsan et al. 2021). This increased trend is possibly linked with climate change in the region attributed to increased temperature and unusual rainfall, urbanization, population growth, inadequate water supply and storage practice, poor sewer, and waste management system, rise in global commerce and tourism (WHO South-East Asia 2023).

The case fatality ratio (CFR) of primary dengue infection is very low with an estimation of 0.018% - 0.1% (Huits and Schwartz 2021). However, the CFR of secondary or tertiary DENV infection is high, although precise estimates are not available, some studies show more than 1% and reaching up to 4% (Liu et al. 2020). Bangladesh's overall CFR of dengue infection (0.34%) seems slightly higher considering the overall CFR reported in other South and Southeast Asian

countries (WHO South-East Asia 2023). However, more than 65% of dengue-related deaths in

306

307

308

309

310

311

312

313

314

315

316

317

318

319

320

321

322

323

Bangladesh were recorded after the introduction of the serotype DENV-3 in 2019. Earlier, DENV serotype 1-3 has been reported over different years between 2000 and 2019 (Rahim et al. 2021) and DENV-4 had reappeared in 2022 (Haider et al. 2023). Thus, secondary, or tertiary infections are likely contributing to higher dengue-related deaths in Bangladesh. In addition, the CFR of the dengue virus infection might have been affected by a lack of active surveillance and missing the mild and asymptomatic cases, and not recording the cases outside the public hospital and few selected private hospitals in Bangladesh or weaker health care system in the country(Ahsan et al. 2021). In some years, the CFR was very high, for example, in the year 2003, the CFR was 2.1 (total cases 486), in the year 2000, 1.68 (total cases 5,551), and in 2022, 0.45 (total cases 62,382). On the other hand, the CFR of DENV infection decreased in the second decade in Bangladesh. This improvement is probably associated with improved access to the health care system, a better understanding of the treatment protocol including the availability of clinical management guidelines and training for the health care providers, better availability of Information, Education, and Communication (IEC) materials, community engagement and expansion of surveillance system to more hospitals in the surveillance system across the county in the recent years, and overall improvement of the economic condition of the country (Diseases Control Division (DGHS) 2013, WHO 2017, Albis et al. 2019). Two large dengue outbreaks occurred in Bangladesh in the year 2019 and 2022 both characterized by unusual weather patterns and the occurrence of two different serotypes. The

324

325

326

327

328

2019 outbreak was characterized by early rainfall of 120 mm in February compared to a monthly

mean of 20 mm precipitation, along with the introduction of a new serotype of DENV-3 in the

country (Ahsan et al. 2021). The 2022 outbreak was characterized by the late onset of rainfall

329

330

331

332

333

334

335

336

337

338

339

340

341

342

351

with 297 mm rainfall in October compared to a monthly mean of 156 mm, and thus prolongation of vector transmission season along with the introduction of a new serotype, DENV-4 in the country (Haider et al. 2023). The occurrence of a new serotype exposed a large naïve population in a densely populated country like Bangladesh. A large proportion of the population is already infected with one of the serotypes of DENV with more than 80% of people living in Dhaka having antibodies against DENV (Salje et al. 2016). Another study predicted an estimated 40 million people being infected with DENV nationally and 2.4 million annual infections (Salje et al. 2019). Thus, any subsequent infections raise the risk of developing severe dengue hemorrhagic fever through antibody-dependent enhancement (ADE) (Teo et al. 2023). The deaths of many people in the year 2022 when the new serotype DENV-4 was introduced were probably associated with secondary or tertiary DENV infection (Haider et al. 2023). Controlling vector-borne diseases in tropical countries where temperatures, humidity, and rainfall remain favorable for breeding mosquitoes during most periods of the year is a difficult task(Haider et al. 2023). Concerns were raised over the development of insecticide resistance (Al-Amin et al. 2020, Ahsan et al. 2021) and the failure of developing a successful dengue

rainfall remain favorable for breeding mosquitoes during most periods of the year is a difficult
task(Haider et al. 2023). Concerns were raised over the development of insecticide resistance
(Al-Amin et al. 2020, Ahsan et al. 2021) and the failure of developing a successful dengue
vaccine (Wang et al. 2017). The prospect of *Wolbachia*-related intervention is bright but still far
from applying on a national scale considering the expenses and technicalities associated with
this. In this situation, an integrated and holistic vector management plan while engaging the local
communities is key for controlling Aedes-borne diseases, especially in resource-limited
countries. Regular destruction of mosquito breeding sites and increasing surveillance for

detecting active cases are key in controlling dengue virus infection. Continuous active

surveillance of DENV infections will enable early detection of cases and outbreaks. Public health authorities will be able to identify areas where the disease is spreading, take immediate action to control mosquito populations, isolate infected patients, and implement public awareness campaigns to educate people about preventive measures. Early detection and response can help prevent the further spread of the disease and reduce its impact on individuals and communities.

Regular destruction of mosquito breeding habitats and increasing surveillance for detecting active cases should prioritize in controlling DENV infection in Bangladesh. Policymakers need to design an Aedes-borne disease management plan by considering a range of diseases that Aedes mosquito can transmit including Chikungunya, yellow fever, Zika virus, West Nile, Japanese Encephalitis, Eastern Equine Encephalitis, Ross River, Rift Valley fever, and the LaCrosse virus (Haider et al. 2023).

Our data should be viewed instead of several weaknesses of our study. We relied on the reported number of cases from the Ministry of Health and Family Welfare's website, which mainly relies on passive reporting systems from the selected health facilities in the country(Ahsan et al. 2021). These numbers seem to be an underestimation of actual cases. A modeling study based on the national seroprevalence of DENV antibodies predicted an annual infection of 2.4 million cases (Salje et al. 2019). However, dengue infection is underestimated globally as it is difficult to diagnose asymptomatic or mild cases that never reach healthcare settings. Although mild cases are missed more frequently, the severe and fatal cases would likely visit the hospital and thus be counted as numerators in our estimation. Thus, our estimation did not overlook the worst-case

scenario, that is, our estimation, for example, estimated the higher CFR rather than the lower possible estimates.

376

374

375

Conclusions:

378

379

380

381

382

383

384

385

386

387

388

389

390

391

392

393

394

395

377

Between the first (2000-2010) and the second decade (2011-2022), dengue cases have increased by 8.3 times, and annual deaths have increased by 2.2 times in Bangladesh. This growth of cases is partly explained by the influence of global warming with an increase of 0.49°C annual temperature as well as changes in duration and length of the rainy season. Unusual rain including early or late rain in and beyond the monsoon season likely contributed to extending the length of the dengue transmission season in Bangladesh. The monthly mean temperature, monthly total rainfall of the first-lagged month and second-lagged months showed a greater influence on monthly incidence of DENV infection in Bangladesh. The mean monthly growth factor remains significantly above one during April-July, which coincides with the hot and rainy season of the country indicating an earlier vector control would benefit the country. The ARIMA model forecasts a continuously increasing trend of DENV infection for the next decade in Bangladesh. We recommend an integrated and holistic vector management plan while engaging the local communities in the regular destruction of mosquito breeding sites and increasing surveillance for detecting active DENV-infected cases. Proactive surveillance, vector control, and vaccine rollout remain essential public health interventions. In the context of climate change, urbanization, trade, and the movement of people with vectors, there is a need to operationalize the One Health approach to address dengue fever and other vector-borne diseases in Bangladesh and beyond.

396

397 **Acknowledgments:** 398 399 We are grateful to the Ministry of Health and Family Welfare of Bangladesh for publicly sharing 400 the dengue cases and deaths data. We acknowledge Bangladesh Meteorological Department for 401 sharing the meteorological data. NH, and AZ, are part of the PANDORA-ID-NET Consortium 402 (EDCTP 373 Reg/Grant RIA2016E-1609) funded by the European and Developing Countries 403 Clinical Trials Partnership (EDCTP2) programme. NH is a member of the International 404 Development Research Centre, Canada's grant on West African One Health Actions for 405 understanding, preventing, and mitigating outbreaks (109810-001). AZ is a National Institutes of 406 407 Health Research senior investigator, and a Mahathir Science Award and Pascoal Mocumbi Award 408 laureate. 409 410 411 Author contribution statement: NH ideated the study and all authors helped develop the study outline and protocol. MNH and IK collected the data. NH, MNH, MA and AZ analyzed the data. NH, IK and MNH 412 prepared the first draft manuscript and all authors contributed to several drafts and finalization of the 413 manuscript. All authors approved the final draft and submission of the manuscript. 414 415 **Financial Support:** There was no funding for this research. 416 417 **Conflict of interest:** The authors declare that they have no conflict of interest. Ethics statement: This study does not include individual-level data and thus does not require ethical 418 approval. We used publicly available data on Dengue cases and deaths. 419 420 Data availability statement: All the dengue data presented in this manuscript are publicly available on 421 Bangladesh's Ministry of Health and Family Welfare's Directorate General of Health Services website (https://dghs.gov.bd/). The meteorological data were purchased from Bangladesh Meteorological 422 Department and are restricted to use for research purposes only and anyone interested in these data can 423 request Bangladesh Meteorological Department (https://live3.bmd.gov.bd/). 424

426	References
427 428 429	Ahsan, A., N. Haider, R. Kock, and C. Benfield. 2021. Possible Drivers of the 2019 Dengue Outbreak in Bangladesh: The Need for a Robust Community-Level Surveillance System. J Med Entomol. 58: 37–39.
430 431 432	Al-Amin, H. M., F. T. Johora, S. R. Irish, M. R. H. Hossainey, L. Vizcaino, K. K. Paul, W. A. Khan, R. Haque, M. S. Alam, and A. Lenhart. 2020. Insecticide resistance status of Aedes aegypti in Bangladesh. Parasit Vectors. 13: 622.
433 434	Albis, M. L. F., S. K. Bhadra, and B. Chin. 2019. Impact evaluation of contracting primary health care services in urban Bangladesh. BMC Health Serv Res. 19: 854.
435 436	BMD. 2023. Climate and Weather Data Portal Bangladesh Meteorological Department. (http://www.bmddataportal.com/#/).
437 438	CDC. 2019. Transmission through mosquito bites. (https://www.cdc.gov/dengue/transmission/index.html).
439 440 441	Cuong, H. Q., N. T. Hien, T. N. Duong, T. V. Phong, N. N. Cam, J. Farrar, V. S. Nam, K. T. D. Thai, and P. Horby. 2011. Quantifying the Emergence of Dengue in Hanoi, Vietnam: 1998–2009. PLoS Negl Trop Dis. 5: e1322.
442	DGHS. 2023. DGHS. (https://old.dghs.gov.bd/index.php/bd/home/5200-daily-dengue-status-report).
443 444	Diseases Control Division (DGHS) . 2013 . National guidelines for clinical management of Dengue syndrome. Dhaka.
445	European CDC. 2023. Dengue worldwide overview. ECDC.
446 447 448	Focks, D. A., E. Daniels, D. G. Haile, and J. E. Keesling. 1995. A simulation model of the epidemiology of urban dengue fever: Literature analysis, model development, preliminary validation, and samples of simulation results. American Journal of Tropical Medicine and Hygiene.
449 450 451 452	Haider, N., YM. Chang, M. Rahman, A. Zumla, and R. A. Kock. 2021. Dengue outbreaks in Bangladesh: Historic epidemic patterns suggest earlier mosquito control intervention in the transmission season could reduce the monthly growth factor and extent of epidemics. Current Research in Parasitology & Vector-Borne Diseases. 1: 100063.
453 454 455	Haider, N., M. N. Hasan, I. Khalil, D. Tonge, S. Hegde, M. A. B. Chowdhury, M. Rahman, M. Hossain Khan, R. Ansumana, A. Zumla, and M. J. Uddin. 2023. The 2022 dengue outbreak in Bangladesh: hypotheses for the late resurgence of cases and fatalities. J Med Entomol.
456 457 458 459	Haider, N., M. S. Rahman, S. U. Khan, A. Mikolon, E. S. Gurley, M. G. Osmani, I. S. Shanta, S. K. Paul, L. Macfarlane-Berry, A. Islam, J. Desmond, J. H. Epstein, P. Daszak, T. Azim, S. P. Luby, N. Zeidner, and M. Z. Rahman. 2014. Identification and Epidemiology of a Rare HoBi-Like Pestivirus Strain in Bangladesh. Transbound Emerg Dis. 61: 193–198.

- 460 Hasan, M. N., N. Haider, F. L. Stigler, R. A. Khan, D. McCoy, A. Zumla, R. A. Kock, and M. J. Uddin. 2021.
- The Global Case-Fatality Rate of COVID-19 Has Been Declining Since May 2020. Am J Trop Med Hyg.
- 462 104: 2176–2184.
- Huits, R., and E. Schwartz. 2021. Fatal outcomes of imported dengue fever in adult travelers from nonendemic areas are associated with primary infections. J Travel Med. 28.
- Kumar, N., and S. Susan. 2020. COVID-19 Pandemic Prediction using Time Series Forecasting Models,
 pp. 1–7. *In* 2020 11th International Conference on Computing, Communication and Networking
- 467 Technologies (ICCCNT). IEEE.
- 468 **Liu, Y., K. Lillepold, J. C. Semenza, Y. Tozan, M. B. M. Quam, and J. Rocklöv. 2020**. Reviewing estimates of the basic reproduction number for dengue, Zika and chikungunya across global climate zones.
- 470 Environ Res.
- 471 **Murray, N. E. A., M. B. Quam, and A. Wilder-Smith**. **2013**. Epidemiology of dengue: past, present and future prospects. Clin Epidemiol. 5: 299–309.
- 473 Mutsuddy, P., S. Tahmina Jhora, A. K. M. Shamsuzzaman, S. M. G. Kaisar, M. N. A. Khan, and S.
- **Dhiman. 2019**. Dengue Situation in Bangladesh: An Epidemiological Shift in terms of Morbidity and
- 475 Mortality. Can J Infect Dis Med Microbiol. 2019.
- 476 **R Core Team**. **2022**. R: A language and environment for statistical computing. Vienna, Austria. URL https://www.R-project.org/.
- 478 Rahim, R., A. Hasan, N. Hasan, E. E. Nakayama, T. Shioda, and M. Rahman. 2021. Diversity of Dengue
- Virus Serotypes in Dhaka City: From 2017 to 2021. Bangladesh Journal of Medical Microbiology. 15:
- 480 23–29.
- Rahman, K. M., Y. Sharker, R. A. Rumi, M.-U. I. Khan, M. S. Shomik, M. W. Rahman, S. M. Billah, M.
- 482 Rahman, P. K. Streatfield, D. Harley, and S. P. Luby. 2020. An Association between Rainy Days with
- 483 Clinical Dengue Fever in Dhaka, Bangladesh: Findings from a Hospital Based Study. Int J Environ Res
- 484 Public Health. 17: 9506.
- Salje, H., I. Morales, E. S. Gurley, and S. Saha. 2016. Seasonal Distribution and Climatic Correlates of Dengue Disease in Dhaka, Bangladesh. Am J Trop Med Hyg. 94: 1359–1361.
- 487 Salje, H., K. K. Paul, R. Paul, I. Rodriguez-Barraquer, Z. Rahman, M. S. Alam, M. Rahman, H. M. Al-
- 488 Amin, J. Heffelfinger, and E. Gurley. 2019. Nationally-representative serostudy of dengue in
- Bangladesh allows generalizable disease burden estimates. Elife. 8.
- 490 **Sen, P. K. 1968**. Estimates of the Regression Coefficient Based on Kendall's Tau. J Am Stat Assoc. 63: 491 1379–1389.
- 492 **Sharmin, S., E. Viennet, K. Glass, and D. Harley**. **2015**. The emergence of dengue in Bangladesh:
 - 493 epidemiology, challenges and future disease risk. Trans R Soc Trop Med Hyg. 109: 619–627.
 - 494 Siraj, A. S., R. J. Oidtman, J. H. Huber, M. U. G. Kraemer, O. J. Brady, M. A. Johansson, and T. A.
 - 495 **Perkins**. **2017**. Temperature modulates dengue virus epidemic growth rates through its effects on
 - reproduction numbers and generation intervals. PLoS Negl Trop Dis.

497 498	Soneja, S., G. Tsarouchi, D. Lumbroso, and D. K. Tung. 2021 . A Review of Dengue's Historical and Future Health Risk from a Changing Climate. Curr Environ Health Rep. 8: 245–265.
499 500	Sumi, S. N., N. C. Sinha, and M. A. Islam. 2021. Generalized linear models for analyzing count data of rainfall occurrences. SN Appl Sci. 3: 481.
501 502	Teo, A., H. D. Tan, T. Loy, P. Y. Chia, and C. L. L. Chua . 2023 . Understanding antibody-dependent enhancement in dengue: Are afucosylated IgG1s a concern? PLoS Pathog. 19: e1011223.
503 504 505 506	Wang, T. T., J. Sewatanon, M. J. Memoli, J. Wrammert, S. Bournazos, S. K. Bhaumik, B. A. Pinsky, K. Chokephaibulkit, N. Onlamoon, K. Pattanapanyasat, J. K. Taubenberger, R. Ahmed, and J. V. Ravetch. 2017. IgG antibodies to dengue enhanced for FcγRIIIA binding determine disease severity. Science (1979). 355: 395–398.
507 508	Wangdi, K., A. C. A. Clements, T. Du, and S. V. Nery. 2018. Spatial and temporal patterns of dengue infections in Timor-Leste, 2005–2013. Parasit Vectors. 11: 9.
509 510	WHO . 2009 . DENGUE: GUIDELINES FOR DIAGNOSIS, TREATMENT, PREVENTION AND CONTROL. Geneva, Switzerland.
511 512	WHO . 2017 . Improving the quality of care in the public health system in Bangladesh: building on new evidence and current policy levers: Bangladesh Health Systems in Transition, Policy Notes. Dhaka .
513 514	WHO . 2023a . Dengue and severe dengue. https://www.who.int/news-room/fact-sheets/detail/dengue-and-severe-dengue. WHO.
515 516	WHO . 2023b . Dengue and severe dengue. (https://www.who.int/news-room/fact-sheets/detail/dengue-and-severe-dengue).
517	WHO South-East Asia. 2023. Dengue in the South-East Asia. WHO Regional office for South-East Asia.
518 519	WHO-Bangladesh. 2022. Dengue - Bangladesh. https://www.who.int/emergencies/disease-outbreak-news/item/2022-DON424.
520 521 522 523	Yue, S., and P. Pilon. 2004. A comparison of the power of the t test, Mann-Kendall and bootstrap tests for trend detection / Une comparaison de la puissance des tests t de Student, de Mann-Kendall et du bootstrap pour la détection de tendance. Hydrological Sciences Journal. 49: 21–37.
524	
525	

526	Tables and Figure Legends:
527	Tables:
528	Table 1: Comparison of dengue cases, deaths, and weather parameters between the first
529	(2000-20210) and the recent decade (2011-2022) in Bangladesh
530	Table 2: The Mann-Kendell trend test for reported dengue cases in Bangladesh, 2000-2022
531	Table 3: The incidence risk ratio (IRR) of monthly average temperature and total rainfall
532	for monthly incidence of Dengue cases in Bangladesh using time-series count Generalized
533	Linear Model for the period 2000-2022.
534	
535	Figures:
536	Fig 1: Top: Number of dengue cases and deaths over the period 2000-2022, Bangladesh.
537	Bottom: Number of monthly dengue cases and deaths recorded in Bangladesh.
538	
539	Fig 2: The boxplot compares monthly rainfall in Dhaka city, Bangladesh between two decades
540	(2000-2010 vs 2011-2022). The bottom and top of the box indicate the first and third quantiles,
541	the band inside the box is the median. The dots outside the box are individual outliers. Most of
542	the months in the second decade had outlier rainfall whereas in the first decade, only the cooler
543	months (Nov-Jan) had some extreme rainfall.
544 545	Fig 3: Top: Mean monthly growth factor for the period of 2000-2022. Bottom: The Monthly
546	growth factor for the individual year 2000-2022. The horizontal line indicates monthly growth
547	factor 1 (the same number of DENV infections in two subsequent months).
548	
549	Fig 4: The observed and forecasted number of dengue cases in Bangladesh using the
550	Autoregressive moving average (ARIMA) model including a 95% confidence interval.

Tables

Table 1:

	First decade (2000-	Recent decade (2011-	p-
	2010)	2022)	value
Mean annual dengue cases (±Standard	2216.64 (±2123.62)	18321.92	0.219
deviation [SD])		$(\pm 31,778.90)$	
Mean annual dengue deaths (±SD)	21.18 (±30.69)	46.58 (±90.90)	0.853
Mean Case-fatality ratio (± SD)	$0.69 (\pm 0.79)$	0.23 (±13)	0.08
Mean temperature °C (±SD)	26.35 (±0.49)	26.84 (±0.37)	< 0.001
Mean annual rainfall in mm (±SD)	2078.66 (±459.68)	1764.50 (±448.32)	0.188

Table 2:

Test		
Mann-Kendell trend analysis	Tau	p-value
·	0.26	0.139
Sen's Slop test		
•	Sen's Slope	95% Confidence
	-	Interval
	171.67	-46 to 687

Table 3:

	IRR (95% CI)	P-value
Average temperature	1.26 (1.258 – 1.265)	<0.001
Rainfall (lag 1) in centimeter	1.08 (1.079 – 1.086)	< 0.001
Rainfall (lag 2) in centimeter	1.17 (1.168 – 1.175))	< 0.001

Figures

Fig 1:

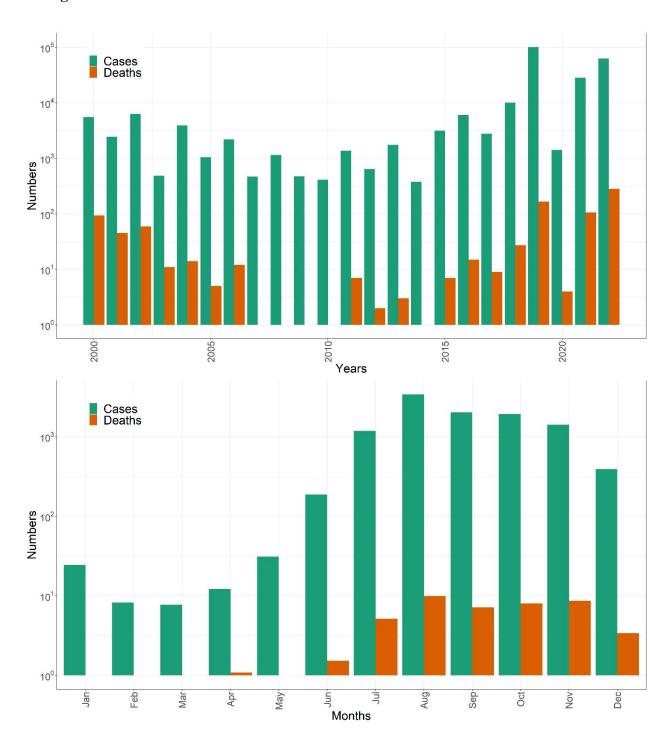
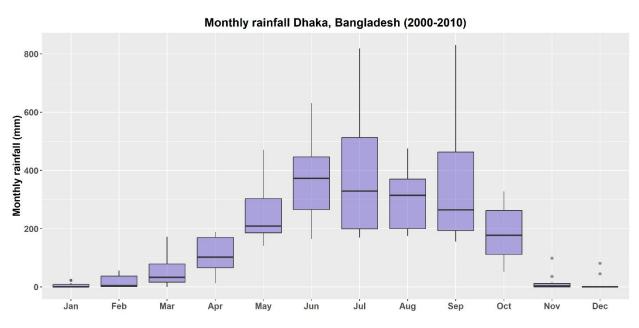


Fig 2:



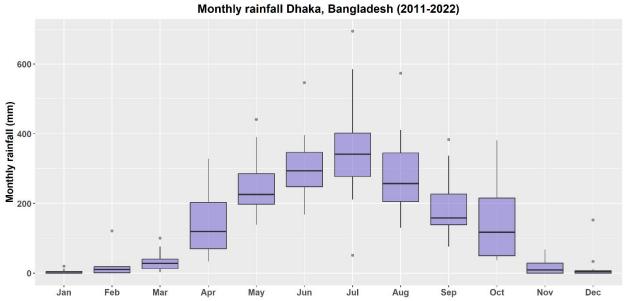


Fig 3:

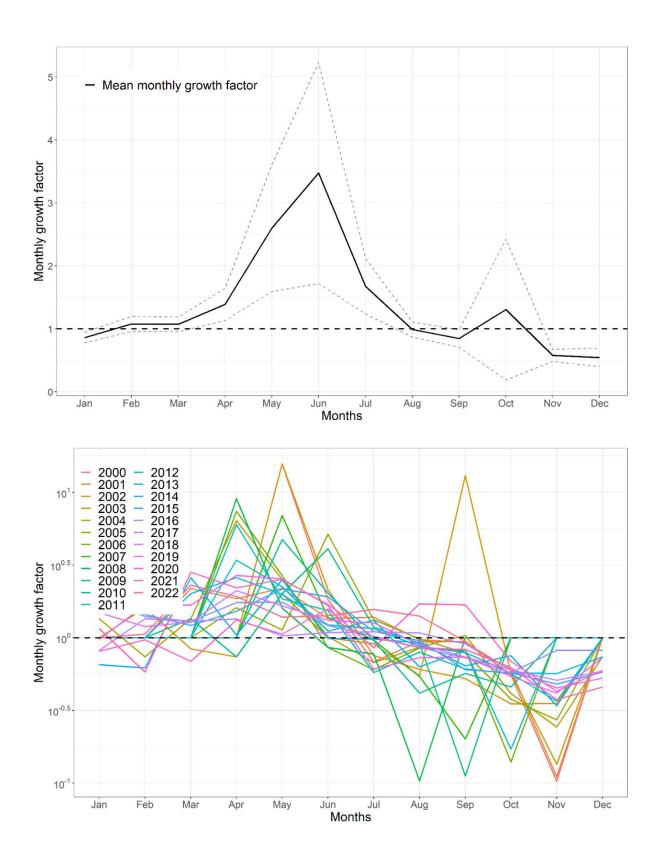
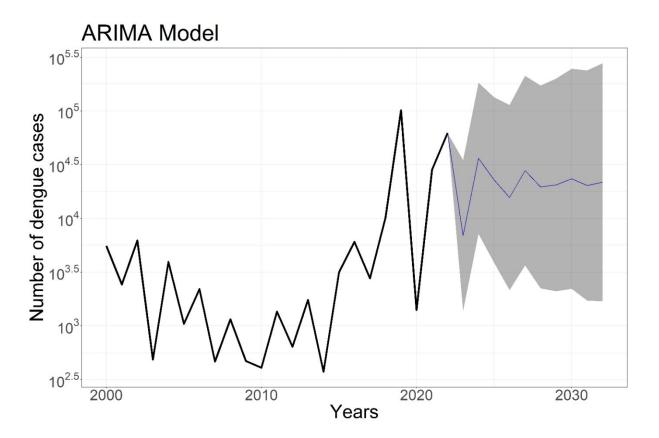


Fig 4:



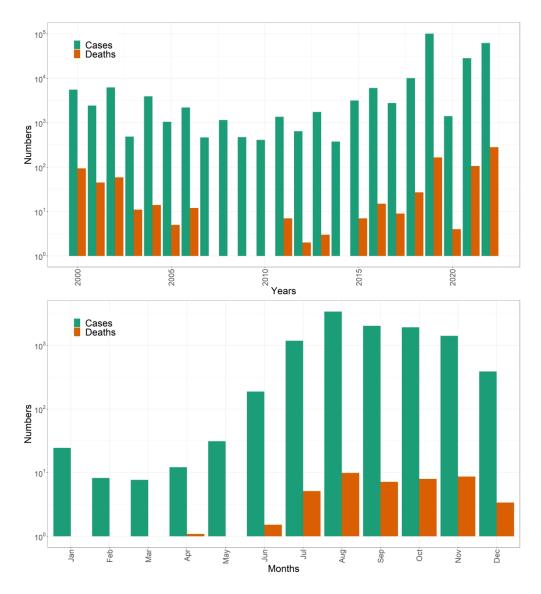


Fig 1: Top: Number of dengue cases and deaths over the period 2000-2022, Bangladesh. Bottom: Number of monthly dengue cases and deaths recorded in Bangladesh.

457x508mm (300 x 300 DPI)

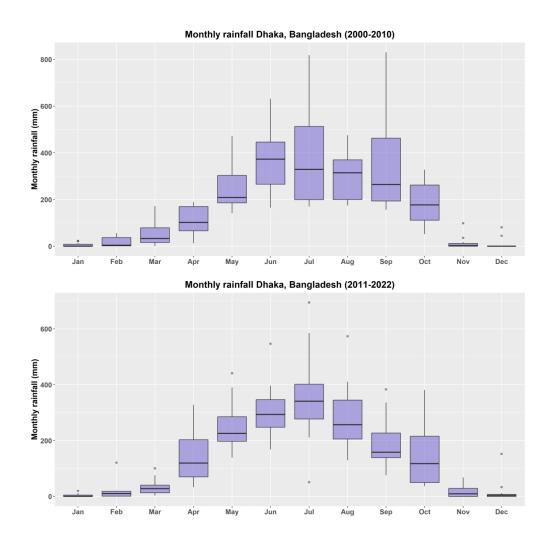


Fig 2: The boxplot compares monthly rainfall in Dhaka city, Bangladesh between two decades (2000-2010 vs 2011-2022). The bottom and top of the box indicate the first and third quantiles, the band inside the box is the median. The dots outside the box are individual outliers. Most of the months in the second decade had outlier rainfall whereas in the first decade, only the cooler months (Nov-Jan) had some extreme rainfall.

304x304mm (300 x 300 DPI)

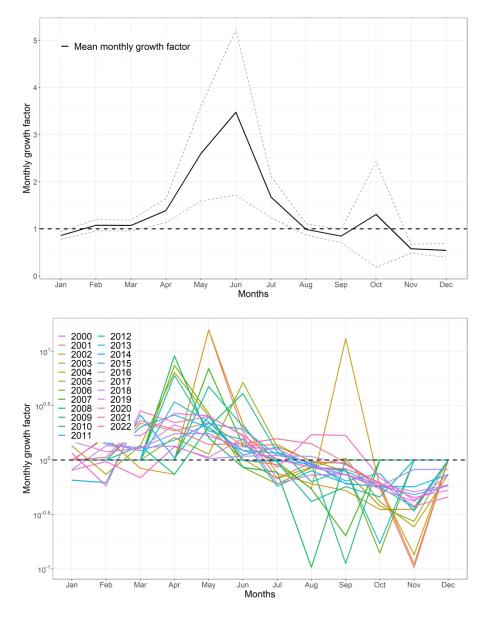


Fig 3: Top: Mean monthly growth factor for the period of 2000-2022. Bottom: The Monthly growth factor for the individual year 2000-2022. The dotted horizontal line indicates monthly growth factor 1 (same number of cases in two subsequent months).

304x406mm (300 x 300 DPI)

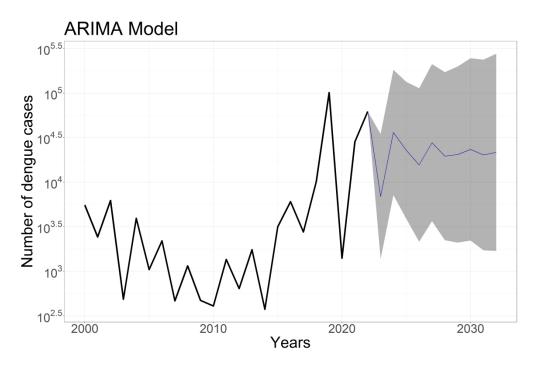


Fig 4: The observed and forecasted number of dengue cases in Bangladesh using the Autoregressive moving average (ARIMA) model including a 95% confidence interval.

457x304mm (300 x 300 DPI)