

DATA-DRIVEN STRATEGIES TO MODEL AND MITIGATE THE THREAT OF ZIKA

*Subhabrata Majumdar**

University of Florida Informatics Institute, Gainesville, FL, USA

ABSTRACT

The recent outbreak of Zika in Brazil and other Latin American countries has posed several challenges in the domain of public health. Although the outbreak is currently under control, because of the lack of vaccines and drugs till now Zika remains a threat. To effectively formulate strategies for preventing a future outbreak, it is imperative to collate and mine relevant data from different sources. In this direction, several data-driven approaches have been explored. This includes modelling the spread of the disease from past outbreaks, identifying vulnerable geographical areas based on the distribution of mosquito vectors, exploring the virus genome for better understanding of the disease mechanism, screening of chemical compounds as potential targets for drug and vaccine development, and predicting new animal reservoirs of the virus for proactive intervention in the virus life cycle. This chapter explores such strategies in detail: highlighting the major themes of contemporary research, providing technical discussions and outlining potential directions of future work.

Keywords: Zika; Infectious diseases; SIR (Susceptible-Infectious-Recovered) model; Zika drugs; machine learning; statistical model; Quantitative Structure Activity Relationship (QSAR); disease surveillance

* Currently at AT&T Labs Research. Corresponding Author Email: subho@research.att.com.

REFERENCES

- Agan, P. N. 2017. "Climate Change and Health Nexus: A Review." *J. earth Sci. Clim. change* 8 (12): 1000435.
- Agusto, F. B., S. Bewick, and W. F. Fagan. 2017. "Mathematical model of Zika virus with vertical transmission." *Infect. Dis. Model.* 23 (2): 244-267.
- Alicino et al, C. 2015. "Assessing Ebola-related web search behaviour: insights and implications from an analytical study of Google Trends-based query volumes." *Infect. Dis. Poverty* 4: 54.
- Andraud, M., N. Hens, C. Marais, and P. Beutels. 2012. "Dynamic Epidemiological Models for Dengue Transmission: A Systematic Review of Structural Approaches." *PLoS One* 7 (11): e49085.
- Arora, N., A. K. Banerjee, and M. L. Narasu. 2018. "Zika outbreak aftermath: status, progress, concerns and new insights." *Future Virol.* 13 (8): 539-556.
- Baca-Carrasco, D., and J. X. Velasco-Hernandez. 2016. "Sex, mosquitoes and epidemics: an evaluation of Zika disease dynamics." *Bull. Math. Biol.* 78: 2228-2242.
- Bakach, I, and J. Braselton. 2015. "A Survey of Mathematical Models of Dengue Fever." *J. Comput. Sci. Syst. Biol.* 8: 255-267.
- Balasubramanian et al, A. 2017. "Antiviral activities of selected antimalarials against dengue virus type 2 and Zika virus." *Antiviral Res.* (137) 141-150.
- Bauer, F. 2017. "Mathematical epidemiology: Past, present, and future." *Infect. Dis. Model.* 2 (2): 113-127.
- Bellan et al, S. E. 2015. "Statistical power and validity of Ebola vaccine trials in Sierra Leone: a simulation study of trial design and analysis." *Lancet Infect. Dis.* 15 (6): 703-710.
- Bhargava et al, S. 2017. "Identification of structural requirements and prediction of inhibitory activity of natural flavonoids against Zika virus through molecular docking and Monte Carlo based QSAR Simulation." *Nat. Prod. Res.* in press. doi:<https://doi.org/10.1080/14786419.2017.1413574>.
- Bhattacharya et al, M. K. 2013. "Dengue: A Growing Menace -- A Snapshot of Recent Facts, Figures & Remedies." *Int. J. Biomed. Sci.* 9 (2): 61-67.
- Bogoch et al, I. I. 2016a. "Anticipating the international spread of Zika virus from Brazil." *Lancet* 387 (10016): 335-336.
- Bogoch et al, I. I. 2016b. "Potential for Zika virus introduction and transmission in resource-limited countries in Africa and the Asia-Pacific region: a modelling study." *Lancet Infect. Dis.* 16 (11): 1237-1245.

- Brown et al, M. 2018. "Modeling Zika Virus Spread in Colombia Using Google Search Queries and Logistic Power Models." *bioRxiv*. doi:<https://doi.org/10.1101/365155>.
- Caminade et al, C. 2017. "Global risk model for vector-borne transmission of Zika virus reveals the role of El Niño 2015." *Proc. Natl. Acad. Sci. USA* 114 (1): 119-124.
- Capasso, V. 1993. *Mathematical Structures of Epidemic Systems*. Berlin: Springer-Verlag.
- Centers for Disease Control and Prevention 2017. *Laboratory Safety when Working with Zika Virus*. 4 27. <https://www.cdc.gov/zika/laboratories/lab-safety.html>.
- Chan et al, E. H. 2011. "Using Web Search Query Data to Monitor Dengue Epidemics: A New Model for Neglected Tropical Disease Surveillance." *PLoS Negl. Trop. Dis.* 5 (5): e1206.
- Chien et al, L.-C. 2018. "Surveillance on the endemic of Zika virus infection by meteorological factors in Colombia: a population-based spatial and temporal study." *BMC Infect. Dis.* 18 (1): 180.
- Chowell et al, G. 2014. "Synthesizing data and models for the spread of MERS-CoV, 2013: Key role of index cases and hospital transmission." *Epidemics* 9: 40-51.
- Chowell et al, G. 2016. "Using Phenomenological Models to Characterize Transmissibility and Forecast Patterns and Final Burden of Zika Epidemics." *PLoS Curr.* 8.
- Chretien, J. P., C. M. Rivers, and M. A. Johansson. 2016. "Make data sharing routine to prepare for public health emergencies." *PLoS Med.* 13: e1002109.
- Chretien, J.-P., S. Riley, and D. B. George. 2015. "Mathematical modeling of the West Africa Ebola epidemic." *eLife* 4: e09186.
- Devillers, J. 2018. "Repurposing drugs for use against Zika virus infection." *SAR QSAR Environ. Res.* 29: 103-115.
- Dey et al, S. 2017. "A Bioinformatics approach to designing a Zika virus vaccine." *Comput. Biol. Chem.* 68: 143-152.
- Dye, C., and N. Gay. 2003. "Modeling the SARS Epidemic." *Science* 300 (5627): 1884-1885.
- Ekins et al, S. 2016a. "Illustrating and homology modeling the proteins of the Zika virus." *F1000Res.* 5: 275.
- Ekins et al, S. 2016b. "Open drug discovery for the Zika virus." *F1000Res.* 5: 150.
- Elfiky, A. A., and A. M. Ismail. 2018. "Molecular docking revealed the binding of nucleotide/side inhibitors to Zika viral polymerase solved structures." *SAR QSAR Environ Res.* 29 (5): 409-418.

-
- Evans et al, M. V. 2017. "Data-driven identification of potential Zika virus vectors." *eLife* 6: e22053.
- Fitzgibbon, W.E., J.J. Morgan, and G.F. Webb. 2017. "An outbreak vector-host epidemic model with spatial structure: the 2015-2016 Zika outbreak in Rio De Janeiro." *Theor. Bio. Med. Model.* 14 (7): 1-17.
- Funk et al, S. 2016. "Comparative Analysis of Dengue and Zika Outbreaks Reveals Differences by Setting and Virus." *PLoS Negl. Trop. Dis.* 10 (12): e0005173.
- Ganeshkumar et al, P. 2018. "Dengue infection in India: A systematic review and meta-analysis." *PLoS Negl. Trop. Dis.* 12 (7): e0006618.
- Garcia-Luna et al, S. M. 2018. "Variation in competence for ZIKV transmission by *Aedes aegypti* and *Aedes albopictus* in Mexico." *PLoS Negl. Trop. Dis.* 12 (7): e0006599.
- Goswami et al, N. K. 2018. "Mathematical modeling of zika virus disease with nonlinear incidence and optimal control." *J. Phys. Conf. Ser.* (IOP Publishing) 1000: 012114.
- Guzzetta et al, G. 2016. "Assessing the potential risk of Zika virus epidemics in temperate areas with established *Aedes albopictus* populations." *Euro. Surveill.* 21 (15): 30199.
- Han et al, B. 2019. "Confronting data sparsity to identify potential sources of Zika virus spillover infection among primates." *Epidemics* 27: 59-65.
- Hethcote, H. 2000. "The Mathematics of Infectious Diseases." *SIAM Review* 42 (4): 599-653.
- Hijmans et al, R. 2005. "Very high resolution interpolated climate surfaces for global land areas." *Int. J. Climatol.* 25: 1965-1978.
- Holmes, E. C., A. Rambaut, and K. G. Andersen. 2018. "Pandemics: spend on surveillance, not prediction." *Nature* 558: 180-182.
- Huppert, A., and G. Katriel. 2013. "Mathematical modelling and prediction in infectious disease epidemiology." *Clin. Microbiol. Infect.* 19 (11): 999-1005.
- Jowkar, G. H., and E. G. Mansoori. 2016. "Perceptron ensemble of graph-based positive-unlabeled learning for disease gene identification." *Comput. Biol. Chem.* 64: 263-270.
- Keeling, M. J.; and L. Danon. 2009. "Mathematical modelling of infectious diseases." *Br. Med. Bull.* 92: 33-42.
- Kuhlman et al, C. J. 2018. "Hybrid Agent-based modeling of Zika in the united states." *2017 Winter Simulation Conference (WSC)*.
- Lo, D., and B. Park. 2018. "Modeling the spread of the Zika virus using topological data analysis." *PLoS ONE* 13 (2): e0192120.

- Matheson, T., B. Satterthwaite, and H. C. Highlander. 2017. "Modeling the spread of the Zika virus at the 2016 olympics." *Spora: A Journal of Biomathematics* 3: 29-44.
- May, R. M., and R. M. Anderson. 1991. *Infectious diseases of humans: dynamics and control*. Oxford: Oxford University Press.
- Meltzer et al, M. E. 2014. "Estimating the future number of cases in the Ebola epidemic--Liberia and Sierra Leone." *MMWR Suppl.* 63: 1-14.
- Mesci et al, P. 2018. "Modeling neuro-immune interactions during Zika virus infection." *Hum. Mol. Genet.* 27 (1): 41-52.
- Messina et al, J. P. 2016. "Mapping global environmental suitability for Zika virus." *eLife* 5 (1): e15272.
- Mlakar et al, J. 2016. "Zika virus associated with microcephaly." *New Eng. J. Med.* 374: 951-958.
- Musso et al, D. 2014. "Potential for Zika virus transmission through blood transfusion demonstrated during an outbreak in French Polynesia, November 2013 to February 2014." *Euro. Surveill.* 10 (19): 14.
- Nandy et al, A. 2016. "Characterizing the Zika Virus Genome – A Bioinformatics Study." *Curr. Comput. Aided Drug. Des.* 12: 87-97.
- NDTV. 2018. *3 Fresh Zika Cases In Rajasthan Takes Total To 126*. October 22. <https://www.ndtv.com/india-news/3-fresh-zika-cases-in-rajasthan-takes-total-to-126-1935892>.
- Nishiura et al, H. 2016a. "Transmission potential of Zika virus infection in the South Pacific." *Int. J. Infect. Dis.* 45: 95-97.
- Nishiura et al, H. 2016b. "Preliminary estimation of the basic reproduction number of Zika virus infection during Colombia epidemic, 2015–2016." *Trav. Med. Infect. Dis.* 14: 274-276.
- O'Reilly et al, K. M. 2018. "Projecting the end of the Zika virus epidemic in Latin America: a modelling analysis." *BMC Medicine* 16: 180.
- Padmanabhan, P., P. Seshaiyer, and C. Castillo-Chavez. 2017. "Mathematical modeling, analysis and simulation of the spread of Zika with influence of sexual transmission and preventive measures." *Lett. Bioinform.* 4 (1): 148-166.
- Pattanaik et al, A. 2018. "Discovery of a non-nucleoside RNA polymerase inhibitor for blocking Zika virus replication through in silico screening." *Antiviral Res.* 151: 78-86.
- Perkins et al, T. A. 2016. "Model-based projections of Zika virus infections in childbearing women in the Americas." *Nat. Microbiol.* 1 (9): 16126.
- Perkins et al, T. A. 2018. "What lies beneath: a spatial mosaic of Zika virus transmission in the 2015-2016 epidemic in Colombia." *bioRxiv* 276006. <https://www.biorxiv.org/content/early/2018/03/05/276006>.

- Ramharack, P., and M. E. S. Soliman. 2018. "Zika virus NS5 protein potential inhibitors: an enhanced in silico approach in drug discovery." *J Biomol. Struct. Dyn.* 36 (5): 1118-1133.
- Rock et al, K. 2014. "Dynamics of infectious diseases." *Rep. Prog. Phys.* 77: 26602.
- Rodriguez Barraquer et al, I. 2016. "Predicting intensities of Zika infection and microcephaly using transmission intensities of other arboviruses." *Bull. World Health Organ.* doi:10.2471/BLT.16.174128 .
- Saad-Roy, C. M., P. Van den Driessche, and J. Ma. 2016. "Estimation of Zika virus prevalence by appearance of microcephaly." *BMC Infect. Dis.* 16: 754.
- Sahoo, M., L. Jena, S. Daf, and S. Kumar. 2016. "Virtual Screening for Potential Inhibitors of NS3 Protein of Zika Virus." *Genomics Inform.* 14 (3): 104-111.
- Scata et al, M. 2016. "The impact of heterogeneity and awareness in modeling epidemic spreading on multiplex networks." *Sci. Reports* 6: 37105.
- Shutt et al, D. P. 2017. "Estimating the reproductive number, total outbreak size, and reporting rates for Zika epidemics in South and Central America." *Epidemics* 21: 63-79.
- Singh, A., and N. K. Jana. 2017. "Discovery of potential Zika virus RNA polymerase inhibitors by docking-based virtual screening." *Comp. Biol. Chem.* 71 (C): 144-151.
- Sinigaglia, A., S. Riccetti, M. Trevisan, and L. Barzon. 2018. "In silico approaches to Zika virus drug discovery." *Expert Opin. Drug Discov.* 13 (9): 825-835.
- Siraj et al, A. S. 2018. "Spatiotemporal incidence of Zika and associated environmental drivers for the 2015-2016 epidemic in Colombia." *Sci. Data* 5: 180073.
- Sorichetta et al, A. 2015. "High-resolution gridded population datasets for Latin America and the Caribbean in 2010, 2015, and 2020." *Sci. Data* 2: 150045.
- Srivastav, A. K., N. K. Goswami, M. Ghosh, and X.-Z. Li. 2018. "Modeling and optimal control analysis of Zika virus with media impact." *Int. J. Dynam. Control* in press. doi:https://doi.org/10.1007/s40435-018-0416-0.
- Suparit, P., A. Wiratsudakul, and C. Modchang. 2018. "A mathematical model for Zika virus transmission dynamics with a time-dependent mosquito biting rate." *Theor. Biol. Med. Model.* 15: 11.
- Suwanmanee, S., and N. Luplertlop. 2017. "Dengue and Zika viruses: lessons learned from the similarities between these Aedes mosquito-vectored arboviruses." *J. Microbiol.* 55 (2): 81-89.
- Vorou, R. 2016. "Zika virus, vectors, reservoirs, amplifying hosts, and their potential to spread worldwide: what we know and what we should investigate urgently." *Int. J. Infect. Dis.* 48: 85-90.

- Vynnycky, E., and R.G. White. 2010. *An Introduction to Infectious Disease Modelling*. Oxford: Oxford University Press.
- Wang, L., H. Zhao, S. M. Oliva, and H. Zhu. 2017. "Modeling the transmission and control of Zika in Brazil." *Sci. Rep.* 7: 7721.
- WHO Regional Office for Europe 2016. *Zika virus vectors and risk of spread in the WHO European Region*.
http://www.euro.who.int/__data/assets/pdf_file/0007/304459/WEB-news_competence-of-Aedes-aegypti-and-albopictus-vector-species.pdf?ua=1.
- Wiratsudakul, A., P. Suparit, and C. Modchang. 2018. "Dynamics of Zika virus outbreaks: an overview of mathematical modeling approaches." *PeerJ* 6: e4526.
- Yong, E. 2017. *Is It Possible to Predict the Next Pandemic?* Oct 25. <https://www.theatlantic.com/science/archive/2017/10/pandemic-prediction-challenge/543954/>.
- Yuan et al, S. 2017. "Structure-based discovery of clinically approved drugs as Zika virus NS2B-NS3 protease inhibitors that potently inhibit Zika virus infection in vitro and in vivo." *Antiviral Res.* 145: 33-43.
- Zanluca et al, C. 2015. "First report of autochthonous transmission of Zika virus in Brazil." *Mem. Inst. Oswaldo Cruz* 110: 569-572.
- Zhang et al, Q. 2017. "Spread of Zika virus in the Americas." *Proc. Natl. Acad. Sci. USA* 114: 433-443.
- Zinszer et al, K. 2017. "Reconstruction of Zika Virus Introduction in Brazil." *Emerg. Infect. Dis.* 23 (1): 91-94.