

## A GENERALIZED MODELLING APPROACH TO ASSESS CLIMATE INFLUENCES ON HAND, FOOT, AND MOUTH DISEASE IN EAST COAST MALAYSIA

(*Pendekatan Pemodelan Teritlak untuk Mengkaji Pengaruh Iklim Terhadap Penyakit Tangan, Kaki dan Mulut di Pantai Timur Malaysia*)

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### ABSTRACT

Hand, foot, and mouth disease (HFMD) outbreaks in Asia have increased since the late 1990s, causing severe and often fatal consequences. Several statistical approaches, such as Generalized Linear Models (GLM) and Generalized Additive Models (GAM), have been used in numerous studies to examine the association between climate factors and HFMD cases. However, the results vary by country. In Malaysia, these issues require further research, as there are only a few studies employing GLM and GAM approaches that focus on HFMD cases and climate factors, particularly in the East Coast region. Therefore, this study explores the association between HFMD and climate factors on Malaysia's East Coast using GLM and GAM with Negative Binomial to identify the best model for interpreting HFMD cases. The findings show that climate factors affect HFMD differently across states in East Coast Malaysia. The results show that the GAM Negative Binomial model best represents these issues. The temperatures between 26°C and 28°C will decrease the risk of HFMD cases in Pahang over the next two weeks. Besides, temperatures ranging from 25 to 27°C and 28.5 to 30°C significantly increased HFMD risk in Terengganu over the next two weeks. Nevertheless, Kelantan found no correlation between climate and HFMD. These findings can help local health authorities in developing a climate-based early warning system to minimize HFMD outbreaks in Malaysia's East Coast Region.

**Keywords:** generalized linear model; generalized additive model; climate change; disease

### ABSTRAK

Penyakit tangan, kaki, dan mulut (HFMD) telah meningkat di negara Asia sejak akhir 1990-an, membawa kepada kesan negatif dan sering kali membawa maut. Beberapa pendekatan statistik, seperti Model Linear Teritlak (GLM) dan Model Tambahan Teritlak (GAM), telah digunakan dalam banyak kajian untuk mengkaji hubungan antara faktor iklim dan kes HFMD. Namun, hasil setiap kajian berbeza mengikut negara. Di Malaysia, isu ini memerlukan penyelidikan lanjut kerana hanya ada beberapa kajian yang menggunakan pendekatan GLM dan GAM Binomial Negatif digunakan untuk menentukan model terbaik bagi mewakili kes HFMD dan faktor iklim di Pantai Timur. Hasil kajian ini menunjukkan bahawa faktor iklim mempengaruhi kes penyakit HFMD secara berbeza mengikut negeri di Pantai Timur Malaysia. Kajian ini menunjukkan bahawa model GAM Binomial Negatif adalah yang terbaik untuk mentafsir isu ini. Suhu antara 26°C dan 28°C akan mengurangkan risiko kes HFMD di Pahang dalam tempoh dua minggu akan datang. Selain itu, suhu antara 25 hingga 27°C dan 28.5 hingga 30°C secara signifikan meningkatkan risiko kes HFMD di Terengganu dalam tempoh dua minggu akan datang. Manakala, tiada korelasi ditemui antara iklim dan kes penyakit HFMD di Kelantan. Hasil ini dapat membantu pihak berkuasa kesihatan tempatan membangunkan sistem amaran awal penyakit berasaskan perubahan iklim untuk mengurangkan wabak HFMD di Pantai Timur Malaysia.

Kata kunci: model linear teritlak; model tambahan teritlak; perubahan iklim; penyakit

## References

- Akaike H. 1998. Information theory and an extension of the maximum likelihood principle. In Parzen E., Tanabe K. & Kitagawa G. (eds.). *Selected Papers of Hirotugu Akaike*: 199-213. New York, NY: Springer.
- Allison P.D. 1999. *Multiple Regression: A Primer*. Thousand Oaks, CA: Pine Forge Press.
- Alsop J., Flewett T.H. & Foster J.R. 1960. "Hand-foot-and-mouth disease" in Birmingham in 1959. *British Medical Journal* **2**(5214): 1708-1711.
- Breslow N.E. 1984. Extra-Poisson variation in log-linear models. *Journal of the Royal Statistical Society: Series C (Applied Statistics)* **33**(1): 38-44.
- Cameron A.C. & Trivedi P.K. 1990. Regression-based tests for overdispersion in the Poisson model. *Journal of Econometrics* **46**(3): 347-364.
- Cameron A.C. & Trivedi P.K. 2013. *Regression Analysis of Count Data*. 2nd Ed. Cambridge, UK: Cambridge University Press.
- Chai T. & Draxler R.R. 2014. Root mean square error (RMSE) or mean absolute error (MAE)? – Arguments against avoiding RMSE in the literature. *Geoscientific Model Development Discussions* **7**: 1247–1250.
- Chang H.-L., Chio C.-P., Su H.-J., Liao C.-M., Lin C.-Y., Shau W.-Y., Chi Y.-C., Cheng Y.-T., Chou Y.-L., Li C.-Y., Chen K.-L & Chen K.-T. 2012. The association between enterovirus 71 infections and meteorological parameters in Taiwan. *PLoS ONE* **7**(10): e46845.
- Chen C., Lin H., Li X., Lang L., Xiao X., Ding P., He P., Zhang Y., Wang M. & Liu Q. 2014. Short-term effects of meteorological factors on children hand, foot and mouth disease in Guangzhou, China. *International journal of Biometeorology* **58**(7): 1605-1614.
- Chen K.-T., Chang H.-L., Wang S.-T., Cheng Y.-T. & Yang J.-Y. 2007. Epidemiologic features of hand-foot-mouth disease and herpangina caused by enterovirus 71 in Taiwan, 1998–2005. *Pediatrics* **120**(2): e244-e252.
- Chen S., Liu X., Wu Y., Xu G., Zhang X., Mei S., Zhang Z., O'Meara M., O'Gara M.C., Tan X. & Li L. 2019. The application of meteorological data and search index data in improving the prediction of HFMD: a study of two cities in Guangdong Province, China. *Science of The Total Environment* **652**: 1013-1021.
- Cheong Y.L., Burkart K., Leitão P.J. & Lakes T. 2013. Assessing weather effects on dengue disease in Malaysia. *International Journal of Environmental Research and Public Health* **10**(12): 6319-6334.
- Chowdhury M.Z.I. & Turin T.C. 2020. Variable selection strategies and its importance in clinical prediction modelling. *Family Medicine and Community Health* **8**(1): e000262.
- Du Z., Xu L., Zhang W., Zhang D., Yu S. & Hao Y. 2017. Predicting the hand, foot, and mouth disease incidence using search engine query data and climate variables: an ecological study in Guangdong, China. *BMJ Open* **7**(10): e016263.
- Du Z., Zhang W., Zhang D., Yu S. & Hao Y. 2016. The threshold effects of meteorological factors on hand, foot, and mouth disease (HFMD) in China, 2011. *Scientific Reports* **6**(1): 36351.
- Flewett T.H., Warin R.P. & Clarke S.K. 1963. Hand, foot, and mouth disease' associated with Coxsackie A5 virus. *Journal of Clinical Pathology* **16**(1): 53-55.
- Guo Z., Wang Y., Li Y. & Zhou L. 2023. Impact of meteorological factors on the incidence of hand-foot-mouth disease in Yangzhou from 2017 to 2022: a time series study. *Frontiers in Public Health* **11**: 1278516.
- Hardin J.W. & Hilbe J.M. 2007. *Generalized Linear Models and Extensions*. 2nd Ed. College Station, Texas: Stata Press.
- Hastie T. & Tibshirani R. 1986. Generalized additive models. *Statistical Science* **1**(3): 297-318.
- Hastie T. & Tibshirani R. 1990. Exploring the nature of covariate effects in the proportional hazards model. *Biometrics* **46**(4): 1005-1016.
- Hastie T. & Tibshirani R. 1995. Generalized additive models for medical research. *Statistical Methods in Medical Research* **4**(3): 187-196.
- Hastie T., Tibshirani R. & Friedman J. 2009. *The Elements of Statistical Learning: Data Mining, Inference, and Prediction*. 2nd Ed. New York: Springer.
- Hii Y.L., Rocklöv J. & Ng N. 2011. Short-term effects of weather on hand, foot, and mouth disease. *PLoS ONE* **6**(2): e16796.
- Hilbe J.M. 2011. *Negative Binomial Regression*. 2nd Ed. Cambridge, UK: Cambridge University Press.
- Hyndman R.J. 2014. Measuring forecast accuracy. In Gilliland M., Tashman L. & Sglaivo U. (eds.). *Business Forecasting: Practical Problems and Solutions*: 177-183. Hoboken: John Wiley & Sons.
- Ibrahim A.O., Chen W.Y., Robert J.Z., Yi C.C. & Majid M.A. 2024. An Investigation of the Effect of Weather on Hand-Foot Mouth Disease Incidence in Sarawak. In Thiruchelvam V., Alfred R., Ismail Z.I.B.A., Haviluddin H. & Baharum A. (eds.). *Proceedings of the 4th International Conference on Advances in Computational Science and Engineering, ICACSE 2023. Lecture Notes in Electrical Engineering*: 407-419. Singapore: Springer Nature Singapore.

*A Generalized Modelling Approach to Assess Climate Influences on HFMD in East Coast Malaysia*

- Kim B.I., Ki H., Park S., Cho E. & Chun B.C. 2016. Effect of climatic factors on hand, foot, and mouth disease in South Korea, 2010-2013. *PLoS ONE* **11**(6): e0157500.
- Krzanowski W.J. 1998. *An Introduction to Statistical Modelling*. Hoboken: Wiley.
- Kutner M.H., Nachtsheim C.J., Neter J. & Li W. 2005. *Applied Linear Statistical Models*. 5th Ed. New York, NY: McGraw-Hill.
- Leong P.F., Labadin J., Abd Rahman S.B. & Juan S.F.S. 2011. Quantifying the relationship between the climate and hand-foot-mouth disease (HFMD) incidences. In *2011 Fourth International Conference on Modeling, Simulation and Applied Optimization*, pp. 1-5.
- Li Y., Zhang J. & Zhang X. 2014. Modeling and preventive measures of hand, foot and mouth disease (HFMD) in China. *International Journal of Environmental Research and Public Health* **11**(3): 3108-3117.
- Liao Y., Ouyang R., Wang J. & Xu B. 2015. A study of spatiotemporal delay in hand, foot and mouth disease in response to weather variations based on SVD: a case study in Shandong Province, China. *BMC Public Health* **15**: 71.
- Lin H., Zou H., Wang Q., Liu C., Lang L., Hou X. & Li Z. 2013. Short-term effect of El Nino-Southern Oscillation on pediatric hand, foot and mouth disease in Shenzhen, China. *PLoS ONE* **8**(7): e65585.
- Ma E., Lam T., Wong C. & Chuang S.K. 2010. Is hand, foot and mouth disease associated with meteorological parameters? *Epidemiology & Infection* **138**(12): 1779-1788.
- Marquardt D.W. 1970. Generalized inverses, ridge regression, biased linear estimation, and nonlinear estimation. *Technometrics* **12**(3): 591-612.
- McCullagh P. 1989. *Generalized Linear Models*. 2nd Ed. New York: Routledge.
- Melnick J.L. 1984. Enterovirus type 71 infections: a varied clinical pattern sometimes mimicking paralytic poliomyelitis. *Reviews of Infectious Diseases* **6**: 387-390.
- Miller A. 2002. *Subset Selection in Regression*. 2nd Ed. Boca Raton, FL: Chapman and Hall.
- Ministry of Health Malaysia. 2012. Hand, foot, and mouth disease (HFMD): Nasihat kesihatan (advisory) HFMD. <http://www.moh.gov.my/english.php/pages/view/171> (23 February 2019).
- Nelder J.A. & Wedderburn R.W.M. 1972. Generalized linear models. *Journal of the Royal Statistical Society Series A: Statistics in Society* **135**(3): 370-384.
- Neter J., Kutner M.H., Nachtsheim C.J. & Wasserman W. 1996. *Applied Linear Statistical Models*. 4th Ed. New York, NY: McGraw-Hill.
- O'brien R.M. 2007. A caution regarding rules of thumb for variance inflation factors. *Quality & Quantity* **41**: 673-690.
- Onozuka D. & Hashizume M. 2011. The influence of temperature and humidity on the incidence of hand, foot, and mouth disease in Japan. *Science of the Total Environment* **410-411**: 119-125.
- Ooi M.H., Wong S.C., Lewthwaite P., Cardosa M.J. & Solomon T. 2010. Clinical features, diagnosis, and management of enterovirus 71. *The Lancet Neurology* **9**(11): 1097-1105.
- Robinson C.R., Doane F.W. & Rhodes A.J. 1958. Report of an outbreak of febrile illness with pharyngeal lesions and exanthem: Toronto, summer 1957—isolation of group A coxsackie virus. *Canadian Medical Association Journal* **79**(8): 615-621.
- Schmidt N.J., Lennette E.H. & Ho H.H. 1974. An apparently new enterovirus isolated from patients with disease of the central nervous system. *Journal of Infectious Diseases* **129**(3): 304-309.
- Siegel A.F. 2016. *Practical Business Statistics*. 7th Ed. Cambridge, Massachusetts: Academic Press.
- Sigauke C., Kumar S., Maswanganyi N. & Ranganai E. 2019. Reliable predictions of peak electricity demand and reliability of power system management. In Anand A. & Ram M. (eds.). *System Reliability Management: Solutions and Technologies*: 137-160. Boca Raton, FL: CRC Press.
- Thanh T.C., Giao N.T.Q., Khoa D.D. & Hien T.T. 2016. Effects of climate variations on hand-foot-mouth disease in Ho Chi Minh City. *Journal of Science and Technology* **54**(2A): 120-127.
- Underwood F.M. 2009. Describing long-term trends in precipitation using generalized additive models. *Journal of Hydrology* **364**(3-4): 285-297.
- US Centers for Disease Control and Prevention. 2021. About hand, foot, and mouth disease. <https://www.cdc.gov/hand-foot-mouth/index.html> (20 May 2021).
- Wahid N.A.A., Suhaila J. & Rahman H.A. 2021. Effect of climate factors on the incidence of hand, foot, and mouth disease in Malaysia: A generalized additive mixed model. *Infectious Disease Modelling* **6**: 997-1008.
- Wahid N.A.A., Suhaila J. & Sulekan A. 2020. Temperature effect on HFMD transmission in selangor, Malaysia. *Sains Malaysiana* **49**(10): 2587-2597.
- Wang S.M., Liu C.C., Tseng H.W., Wang J.R., Huang C.C., Chen Y.J., Yang Y.J., Lin S.J. & Yeh T.F. 1999. Clinical spectrum of enterovirus 71 infection in children in southern Taiwan, with an emphasis on neurological complications. *Clinical Infectious Diseases* **29**(1): 184-190.
- Wang S.M. & Liu C.C. 2009. Enterovirus 71: epidemiology, pathogenesis and management. *Expert Review of Anti-infective Therapy* **7**(6): 735-742.

- Wang L., Xu C., Wang J., Qiao J., Wu N. & Li L. 2023. Spatiotemporal associations between hand, foot and mouth disease and meteorological factors over multiple climate zones. *International Journal of Biometeorology* **67**(9): 1493-1504.
- Wood S. 2015. Package ‘mgcv’. R Package Version 1, 29.
- Wood S.N. 2017. *Generalized Additive Models: An Introduction with R*. 2nd Ed. New York: Chapman and Hall.
- World Health Organization. 2011. A Guide to clinical management and public health response for hand, foot and mouth disease (HFMD). WHO Regional Office for the Western Pacific.
- Wu H., Wang H., Wang Q., Xin Q. & Lin H. 2014. The effect of meteorological factors on adolescent hand, foot, and mouth disease and associated effect modifiers. *Global Health Action* **7**(1): 24664.
- Xing W., Liao Q., Viboud C., Zhang J., Sun J., Wu J.T., Chang Z., Liu F., Fang V.J., Zheng Y., Cowling B.J., Varma J.K., Farrar J.J., Leung G.M. & Yu H. 2014. Hand, foot, and mouth disease in China, 2008–12: an epidemiological study. *The Lancet Infectious Diseases* **14**(4): 308-318.
- Yang X., Wang J., Zhang G. & Yu Z. 2024. Short-term effects of extreme meteorological factors on hand, foot, and mouth disease infection during 2010–2017 in Jiangsu, China: A distributed lag non-linear analysis. *GeoHealth* **8**(4): e2023GH000942.

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