



University of Pennsylvania
ScholarlyCommons

Statistics Papers

Wharton Faculty Research

10-2013

The Causal Effect of Malaria on Stunting: A Mendelian Randomization and Matching Approach

Hyunseung Kang
University of Pennsylvania

Benno Kreuels


Ohene Adjei

Ralf Krumkamp

Jürgen May

See next page for additional authors

Follow this and additional works at: http://repository.upenn.edu/statistics_papers

 Part of the [Biostatistics Commons](#), [Epidemiology Commons](#), and the [Vital and Health Statistics Commons](#)

Recommended Citation

Kang, H., Kreuels, B., Adjei, O., Krumkamp, R., May, J., & Small, D. (2013). The Causal Effect of Malaria on Stunting: A Mendelian Randomization and Matching Approach. *International Journal of Epidemiology*, 42 (5), 1390-1398. <http://dx.doi.org/10.1093/ije/dyt116>

This paper is posted at ScholarlyCommons. http://repository.upenn.edu/statistics_papers/562
For more information, please contact repository@pobox.upenn.edu.

The Causal Effect of Malaria on Stunting: A Mendelian Randomization and Matching Approach

Abstract

Background Previous studies on the association of malaria and stunted growth delivered inconsistent results. These conflicting results may be due to different levels of confounding and to considerable difficulties in elucidating a causal relationship. Randomized experiments are impractical and previous observational studies have not fully controlled for potential confounding including nutritional deficiencies, breastfeeding habits, other infectious diseases and socioeconomic status.

Methods This study aims to estimate the causal effect between malaria episodes and stunted growth by applying a combination of Mendelian randomization, using the sickle cell trait, and matching. We demonstrate the method on a cohort of children in the Ashanti Region, Ghana.

Results We found that the risk of stunting increases by 0.32 (P -value: 0.004, 95% CI: 0.09, 1.0) for every malaria episode. The risk estimate based on Mendelian randomization substantially differs from the multiple regression estimate of 0.02 (P -value: 0.02, 95% CI: 0.003, 0.03). In addition, based on the sensitivity analysis, our results were reasonably insensitive to unmeasured confounders.

Conclusions The method applied in this study indicates a causal relationship between malaria and stunting in young children in an area of high endemicity and demonstrates the usefulness of the sickle cell trait as an instrument for the analysis of conditions that might be causally related to malaria.

Keywords

malaria, stunting, children, Mendelian randomization, matching

Disciplines

Biostatistics | Epidemiology | Statistics and Probability | Vital and Health Statistics

Author(s)

Hyunseung Kang, Benno Kreuels, Ohene Adjei, Ralf Krumkamp, Jürgen May, and Dylan Small

The Causal Effect of Malaria on Stunting: A Mendelian Randomization and Matching Approach,

Hyunseung Kang^{1*}, Benno Kreuels^{2,3*}, Ohene Adjei⁴, Ralf Krumkamp³,
Jürgen May³, Dylan S. Small^{1\$}

¹Department of Statistics, The Wharton School, University of Pennsylvania, Philadelphia, PA, USA; ² Section for Tropical Medicine, I. Medical Department, University Medical Center Eppendorf, Hamburg, Germany; ³ Infectious Disease Epidemiology, Bernhard Nocht Institute for Tropical Medicine, Hamburg, Germany; ⁴ Kumasi Center for Collaborative Research in Tropical Medicine, Kumasi, Ghana

*These authors contributed equally to this work.

^{\$}Corresponding author: Dylan S. Small, Department of Statistics, The Wharton School, University of Pennsylvania, 3730 Walnut Street, Philadelphia, PA, USA 19104. Phone: 215.898.8222. Fax: 215.898.1280. E-mail: dsmall@wharton.upenn.edu

Summary

Background: Previous studies on the association of malaria and stunted growth delivered inconsistent results. These conflicting results may be due to different levels of confounding and due to considerable difficulties in elucidating a causal relationship. Randomized experiments are impractical and previous observational studies have not fully controlled for potential confounding including nutritional deficiencies, breast-feeding habits, other infectious diseases and socioeconomic status.

Methods: This study aims to estimate the causal effect between malaria episodes and stunted growth by applying a combination of Mendelian randomization, using the sickle cell trait, and matching. We demonstrate the method on a cohort of children in the Ashanti Region, Ghana.

Results: We found that the risk of stunting increases by 0.32 (p-value: 0.004, 95% CI: (0.09,1.0)) for every malaria episode. The risk estimate based on Mendelian randomization substantially differs from the multiple regression estimate of 0.02 (p-value: 0.02, 95% CI: (0.003,0.03)). In addition, based on the sensitivity analysis, our results were reasonably insensitive to unmeasured confounders.

Conclusions: The method applied in this study indicates a causal relationship between malaria and stunting in young children in an area of high endemicity and demonstrates the usefulness of

sickle cell trait as an instrument for the analysis of conditions that might be causally related to malaria.

Keywords: malaria, stunting, children, Mendelian randomization, matching

MeSH Terms: Mendelian Randomization Analysis, Sickle Cell Trait, Malaria, Infants, Growth Disorders, Ghana