Risk factors associated with an outbreak of dengue fever/dengue haemorrhagic fever in Hanoi, Vietnam

D. T. T. TOAN¹*, L. N. HOAT¹, W. HU², P. WRIGHT³ and P. MARTENS⁴

¹ Hanoi Medical University, Institute of Training for Preventive Medicine and Public Health, Vietnam ² School of Public Health and Social Work, Institute of Health and Biomedical Innovation, Queensland University of Technology, Australia

³*The Medical Committee Netherlands-Vietnam*

⁴ International Centre for Integrated Assessment and Sustainable Development, Maastricht University, Maastricht, The Netherlands

Received 17 June 2014; Accepted 15 September 2014; first published online 13 October 2014

SUMMARY

Dengue fever/dengue haemorrhagic fever (DF/DHF) appears to be emerging in Hanoi in recent years. A case-control study was performed to investigate risk factors for the development of DF/DHF in Hanoi. A total of 73 patients with DF/DHF and 73 control patients were included in the study. The risk factor analysis indicated that living in rented housing, living near uncovered sewers, and living in a house discharging sewage directly into to ponds were all significantly associated with DF/DHF. People living in rented houses were 2·2 times more at risk of DF/DHF than those living in their own homes [adjusted odds ratio (aOR) 2·2, 95% confidence interval (CI) 1·1–4·6]. People living in an unhygienic house, or in a house discharging sewage directly to the ponds were 3·4 times and 4·3 times, respectively, more likely to be associated with DF/DHF (aOR 3·4, 95% CI 1–11·7; aOR 4·3, 95% CI 1·1–16·9). These results contribute to the understanding of the dynamics of dengue transmission in Hanoi, which is needed to implement dengue prevention and control programmes effectively and efficiently.

Key words: Dengue fever, dengue haemorrhagic fever, risk factors, Hanoi.

INTRODUCTION

Dengue fever/dengue haemorrhagic fever (DF/DHF) is a fast-spreading vector-borne disease associated with a significant public health impact. Dengue is receiving attention all over the world for its epidemic expansion and high mortality rate [1]. Before 1970, DF/ DHF was detected in only nine countries but has now spread to over 100 countries. The global number of

(Email: dothithanhtoan@hmu.edu.vn)

reported DF/DHF infections has been stated to be around 96 million per year. However, many cases go unreported and it is estimated that the real number could be as high as 390 million (95% confidence interval 284–528) cases [2].

Vietnam is located in the heart of the endemic area for DF/DHF. It is recognized as a major cause of mortality and morbidity in Vietnam and ranks among the top ten communicable disease in terms of overall health burden [3]. In 1958, the first DHF case was described in Vietnam and the first reported outbreak occurred in southern Vietnam in 1963, resulting in 116 deaths [4]. The estimated morbidity and mortality rates between 1979 and 2005 were

^{*} Author for correspondence: Dr D. T. T. Toan, Biostatistics and Medical Informatics Department, Institute of Training for Preventive Medicine and Public Health, Hanoi Medical University, Vietnam.

33–462/100 000 and 0.1-2.7/100 000, respectively, per year. Although dengue transmission occurs in both rural and urban areas in Vietnam, 73% of the population live in rural areas and therefore the majority of DF/DHF cases and deaths are from these areas [5].

Hanoi in Northern Vietnam is a large city where DF/DHF appears to be emerging as a major public health concern. In 2009, Hanoi experienced its largest ever recorded outbreak. In this study we set out to investigate which risk factors are associated with DF/DHF in patients admitted to hospitals in Hanoi. Identifying the risk factors can help to design and apply effective preventive and control strategies.

MATERIALS AND METHODS

This study was performed in Hanoi at the National Hospital of Tropical Diseases, the Hospital of Hanoi Medical University and the Infectious Diseases Department of Bach Mai Hospital, from August 2009 to March 2010. Approval from the hospital administration was obtained before approaching the patients. We performed a prospective matched case-control study on patients who were diagnosed with DF/DHF; control patients were from the same hospital but negative for DF/DHF. The criteria for notification of DF were based on the guidelines of the Ministry of Health, 1999, on surveillance, diagnosis and treatment of dengue. The guidelines stipulate that individuals are suspected to have dengue when they have acute febrile illness (≥ 38 °C) of 2–7 days' duration, with ≥ 2 of the following non-specific manifestations of DF: headache, retro-orbital pain, myalgia, arthralgia, rash, haemorrhagic manifestations, and leucopenia [6]. The control group was matched to the study group for sex and age (± 5 years). The characteristics of the 73 DF/DHF patients and the 73 control patients are shown in Table 1.

All cases and control patients were interviewed during their period of hospitalization by two specially trained interviewers who administered a standard questionnaire. The questionnaire included information on demographic characteristics of the patients such as age, education level and occupation as well as variables related to their residence: location of house, type of house, source of water supply, and type of water containers in the house. The questionnaire had been pre-tested before being used in the study.

 χ^2 and Mann–Whitney tests were used to test for differences in demographic characteristics between DF/ DHF patients and control patients. A multivariable backward stepwise logistic regression model was applied to identify the risk factors for DF/DHF. The independent variables consisted of: age, education, occupation, having been in an epidemic area within past 6 months; place of living: in urban or rural area, in a rented house or near an open sewer; and living conditions such as unhygienic environment, other family member had DF/DHF during past 6 months, presence of mosquitoes, presence of larvae in water containers, directly discharging sewage to ponds, and being flooded during the rainy season. The significance level was set at P < 0.2. Associations are expressed in terms of odds ratios (ORs) with 95% confidence intervals (CIs). All analyses were performed using Stata statistical software, v. 12.1 (StataCorp., USA).

RESULTS

The characteristics of the 73 DF/DHF patients and 73 control patients are shown in Table 1. The mean age of the case group and the control group were comparable. The distribution of education level and occupation were quite similar between the case and control groups; most individuals had college education and many were students.

Table 2 displays variables related to the residence of the study population. In both groups, three quarters resided in the inner districts of Hanoi. More than half of the participants in the case group were living in rented houses, significantly higher than in the control group, with only one third. People in the DF/ DHF group were living in smaller houses with a larger household size compared with those in the control group. In particular, a significantly higher proportion of people in the case group were living in homes near suitable breeding areas for mosquitoes, such as open sewers.

Table 3 presents the adjusted and unadjusted ORs and CIs of variables affecting DF/DHF included in the final logistic regression model. The analysis revealed that living in a rented house, living near open sewers and untreated water discharging directly into nearby ponds/lakes were all significantly associated with DF/DHF. Living in a rented house increased risk by 2·2 times (aOR 2·2, 95% CI 1·1– 4·6). Living in an unhygienic house or one directly discharging sewage into ponds increased risk by 3·4 times and 4·3 times, respectively (aOR 3·4, 95% CI 1–11·7; aOR 4·3, 95% CI 1·1–16·9). Detecting mosquitoes in the house or living near an open sewer constituted a very high risk (aOR 6·3, 95% CI 0·7–59; aOR 6·9,

| Characteristic | | DF/DHF (<i>n</i> = 73) | Controls $(n = 73)$ | Р |
|----------------|--------------------|-------------------------|---------------------|--------|
| Sex | Male | 36 (49.32%) | 36 (49.32%) | |
| | Female | 37 (50.68%) | 37 (50.68%) | >0.05* |
| Age (years) | Mean ± s.d. | 28.1 ± 9.4 | 27.4 ± 9.9 | >0.05† |
| | Range | 17–75 | 18–78 | |
| Education | Elementary school | 9 (12·3%) | 10 (13.7%) | |
| | Middle school | 10 (13.7%) | 4 (5.5%) | >0.05* |
| | High school | 11 (15.1%) | 6 (8.2%) | |
| | University/college | 43 (58.9%) | 53 (72.6%) | |
| Occupation | Office worker | 17 (23.3%) | 24 (32.9%) | >0.05* |
| | Business person | 7 (9.6%) | 12 (16.4%) | |
| | Home worker | 4 (5.5%) | 7 (9.6%) | |
| | Pupil/student | 36 (49.3%) | 22 (30.1%) | |
| | Other | 9 (12.3%) | 8 (11.0%) | |

Table 1. Demographic characteristics of dengue fever/dengue haemorrhagic fever (DF/DHF) and control patients

* By χ^2 test.

† By Mann-Whitney test.

Table 2. Housing variables of dengue fever/dengue haemorrhagic fever (DF/DHF) and control patients

| Variables | | DF/DHF ($n = 73$) | Controls ($n = 73$) | Р |
|-------------------------------------|--------------------------------|---------------------|-----------------------|--------|
| Living area | Inner district | 54 (74.0%) | 55 (75.3%) | >0.05* |
| - | Outer district | 19 (26.0%) | 18 (24.7%) | |
| Accommodation status | Rented | 41 (56.2%) | 25 (34.3%) | <0.05* |
| | Owned | 32 (43.8%) | 48 (65.7%) | |
| Type of housing | Brick construction | 43 (58.9%) | 35 (47.9%) | |
| | Temporary house | 7 (9.6%) | 7 (9.6%) | |
| | Old condominium | 11 (15.1%) | 15 (20.6%) | >0.05* |
| | New condominium | 0 (0.0%) | 3 (4.1%) | |
| | Other | 12 (16.4%) | 13 (17.8%) | |
| Area of household (m ²) | Mean ± s.d. | 54.1 ± 47.1 | 68.2 ± 69.9 | >0.05† |
| | Range | 8-200 | 9-350 | |
| Number of people in household | Mean ± s.d. | 4.2 ± 2.3 | 4.9 ± 4.3 | >0.05† |
| | Range | 1–10 | 1–19 | |
| House with water storage containers | Yes (water tank without cover) | 22 (10.7%) | 21 (1.3%) | >0.05* |
| | No | 53 (70.8%) | 54 (72.0%) | |
| House with water storage containers | Yes | 11 (15.1%) | 13 (17.8%) | >0.05* |
| infested with larvae and/or pupae | No | 52 (71.2%) | 56 (76.7%) | |
| 1 1 | Don't know | 10 (13.7%) | 4 (5.5%) | |
| House environment | Pond/lake/river | 16 (19.3%) | 21 (24.1%) | >0.05* |
| | Open sewer | 33 (39.7%) | 13 (14.9%) | <0.01* |
| | Garbage collection point | 7 (36.8%) | 12 (63.2%) | >0.05* |

* By χ^2 test.

† By Mann–Whitney test.

95% CI 0.9–71.9, respectively), but the difference between the case and control groups was not significant.

DISCUSSION

The results of this study revealed that DF/DHF in Hanoi appeared mainly in downtown districts and

much less frequently in suburban districts, which is consistent with the statistical reports on the DF epidemic in 2009. The highest morbidity rate was found in the 15–30 years age group; at higher ages the rate became very low. Dung & Cam reported in 2003 that more than 90% of dengue morbidity was in individuals aged between 15 and 25 years [7]. This could

| Independent variables | Grouping | Crude OR (95% CI) | Logistic regression aOR (95% CI) |
|---|----------|----------------------|--|
| Living in rented house | Yes | 2.2 (1.1-4.8) | 2.2 (1.1-4.6) |
| Other family member had DF/DHF during past 6 months | Yes | 2 (0.9–4.6) | 1.96 (0.9-4.3) |
| Unhygienic house | Yes | 3.3 (0.9–12) | 3.4 (1–11.7) |
| Mosquitoes in house | Yes | 6.8 (0.7–64.2) | 6.3 (0.7–59.5) |
| Near open sewer | Yes | 6.9 (0.7–67.1) | 7.9 (0.9–71.9) |
| Discharging sewage directly into ponds | Yes | 5.9 (1.3–26.4) | 4.3 (1.1–16.9) |

Table 3. Crude and adjusted logistic regression oddsratios on selected variables

aOR, Adjusted odds ratio; CI, confidence interval; DF, dengue fever; DHF, dengue haemorrhagic fever.

be related to differences in lifestyle, time spent outdoors near vectors, sleeping without mosquito nets or other aspects of inadequate disease prevention among the young. Another explanation could be that the disease can create lifelong immunity for the individual, so that older persons who have been exposed more often may have more resistance, decreasing the morbidity rate.

Considering how many cases were young people, it is perhaps not surprising that around half of the positive cases were students. The second dominant group was office workers. This result differs from the result of a study in Kamphaeng Phet, Thailand [8] where 65% of the patients were farmers. The discrepancy is probably due to a different job distribution in the two regions studied. Students may also be prominent in the cases in our study because many students live in rented accommodation, which was another risk factor for dengue. The combination of cramped rented living accommodation, an unsanitary environment, infrequent cleaning, little knowledge of DF, and little interest in prevention activities may explain why the morbidity rate in the rental group is rising.

People living near stagnant water like ponds, lakes and rivers or open sewers, or favourable mosquito breeding places like garbage collection points, had higher rates of morbidity. This is consistent with other national and international studies [9-17]. In Brazil [10] the epidemic was associated with proximity to uncontrolled waterways and stagnant water in tanks, gutters, and cans. In Pakistan, poor condition of the house, such as uncovered toilet water tank or leaking water pipes, was a highly significant risk factor for the presence of Aedes foci [10]. In Vietnam, a study in Binh Thanh, Dong Thap [11] found that while most families stored their drinking water, only three quarters of them covered their water storage containers. Nowadays, although Hanoi residents seldom use water storage containers like jars or pots, the large area of natural water surfaces available makes an annual epidemic almost unavoidable [12]. Because dengue-spreading mosquitoes spend three stages of their life-cycle in water, it is recommended to reduce mosquito living sites, e.g. long-term stagnant water storage inside and outside the house, open sewers, and natural water surfaces.

According to around half of the respondents, their homes and the surrounding environment are not really clean. Although the difference between case and control groups was not statistically significant, it did reflect how the unsanitary, polluted living environment in the modern environment of the capital is favourable for the vector and the disease.

Environmental factors have a powerful influence on the appearance of DF/DHF. Simple interventions could help; in our study, people living in houses with uncovered water tanks were 7.9 times more likely to get DF. Those in houses discharging sewage directly into ponds had a 5.9 times higher risk than those in houses with sanitary sewage systems. For civil authorities it is important to note that people who stay in rented accommodation had double the risk of those owning their homes. These influences of the environmental factors have also been found in other studies. A 2001 study in Brazil revealed that people living in a slum area had a nearly 10 times higher risk of DF [10]. In An Giang, Vietnam, people living in homes with gardens or water tanks had a threefold higher risk [12].

This study was conducted soon after Hanoi had experienced heavy flooding in August 2009 which reduced the selection and recall bias often found in a case-control study. However, our data came from the three major hospitals for infectious diseases in Hanoi, and may have missed factors related to cases that did not present to these hospitals. Our logistic regression model was set up to identify factors that may influence dengue transmission dynamics. In a complex epidemiological setting, transmission dynamics involve interactions among people, dengue viruses, vectors and ecosystems, where biotic and abiotic determinants have both direct and indirect influences on transmission. Our model cannot account for all possible variables. More information is needed to further unravel the predictors of dengue transmission in Hanoi, which will help to inform planning of prevention and control measures.

CONCLUSIONS

This integrated analysis of the eco-social determinants of DF transmission risk contributes to improved understanding of the dynamics of dengue transmission in Hanoi. Our study shows various well-known risk factors of DF which are components of human behaviour, can be addressed through modifications of municipal services. The social and demographical contexts and lifestyles of people in Hanoi will significantly influence transmission and all information on this population will support dengue prevention and control programmes.

ACKNOWLEDGEMENTS

We thank Dr Nguyen Nhat Cam, Vice-Director of Hanoi Preventive Medicine Center and his staff for the assistance on collecting the epidemiological data. This study was funded by the Netherlands Higher Education (NPT) project on 'Strengthening teaching and research capacity in preventive medicine in Vietnam'.

REFERENCES

- 1. Gubler DJ. Dengue and dengue hemorrhagic fever. Clinical Microbiology Reviews 1998; 11: 480–496.
- Bhatt S, et al. The global distribution and burden of dengue. Nature 2013; 496: 504–507.
- Tien TKN, et al. Predictive indicators for forecasting epidemic of dengue/dengue haemorrhagic fever through epidemiological, virological and entomological surveillance. Dengue Bulletin 1999; 23: 34–39.

- Ha DQ, Ninh TU. Virological surveillance of dengue haemorrhagic fever in Vietnam, 1987–1999. *Dengue Bulletin* 2000; 24: 18–23.
- Tien TKN, et al. Epidemiological analysis of deaths associated with dengue haemorrhagic fever in southern Vietnam in 1999–2000. Dengue Bulletin 2001; 25: 28–32.
- 6. Vietnam Ministry of Health. Guidelines on the surveillance, diagnosis, and treatment of dengue hemorrhagic fever. Hanoi, Vietnam, 1999.
- Dung NT, Cam NN. Dengue fever/dengue hemorrhagic fever in Hanoi – 2003. *Journal of Vietnam Preventive Medicine* 2005; 1: 73–77.
- Koenraadt CJ, et al. Dengue knowledge and practices and their impact on Aedes aegypti populations in Khamphaeng Phet, Thailand. American Journal of Tropical Medicine and Hygiene 2006; 74: 692–700.
- Nguyen ND. Knowledge, attitude and practice on dengue/dengue hemorrhagic fever prevention among mothers in Ho Chi Minh City, Vietnam. *Journal of Ho Chi Minh City's Medical Practice* 1999; 2: 119–124.
- Heukelbach J, et al. Risk factors associated with an outbreak of Dengue fever in a favela in Fortaleza, northeast Brazil, *Tropical Medicine & International Health* 2001; 6: 635–642.
- Huong LTT, et al. Knowledge, attitude and practice on dengue prevention among residences in Binh Thanh commune, Thanh Binh District, Dong Thap province, Vietnam. Journal of Preventive Medicine 2009; 11: 36–42.
- Be PV, et al. Evaluation on knowledge, attitude and practice of people about dengue fever in two communes of An Giang province, Vietnam in 2004. Journal of Preventive Medicine 2005; 3: 56–64.
- Nga NT. Epidemiology of dengue fever/dengue hemorrhagic fever in Hai Ba Trung District, Hanoi City. Master thesis of Hanoi Medical University, Vietnam, 8/1982.
- Toan DTT, et al. Hot spot detection and spatiotemporal dispersion of dengue fever in Hanoi, Vietnam. Global Health Action 2013; 6: 18632.
- Itrat A, et al. Knowledge, awareness and pratices regarding Dengue fever among the adult population of Dengue hit cosmopolitan. PLoS ONE. Published online: 9 July 2008. doi:10.1371/journal.pone.0002620.
- Nalongsack S, et al. Knowledge, attitude and practice regarding dengue among people in Pakse, Laos. Nagoya Journal of Medical Science 2009; 71: 29–37.
- Syed M, et al. Knowledge, attitudes and practice regarding dengue fever among adults of high and low socioeconomic groups, *Journal of Pakistan Medical* Association 2010; 60: 243–247.