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## RESEARCH ARTICLE

## Epidemiological review of dengue fever in Yemen

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**Abstract**

Dengue fever is a mosquito borne disease that is spread rapidly in new governorates in Yemen in last ten years. The large scale of dengue outbreak in Alhodeidah governorate is due to the high vector circulation (*Aedes aegypti*), poor infrastructure, and excesses of raining period more than other cities in Yemen, ineffective vector and infectious disease surveillance and low level of education are the main reason for the increase incidence rate of dengue fever in Yemen. Current review was focused on highlighting the development spreading and the current status of the dengue fever disease in Yemen. In conclusion, dengue fever was spreading in different region in Yemen with high distribution in Alhodeidah, Taiz, Aden and Al-Mukalla. More vector controlling and preventive measurement and management should be considered.

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## INTRODUCTION

Dengue fever one of the most arboviral infections for human in recent decade. Recently increase of distribution and incidence of dengue fever (DF) and dengue hemorrhagic (DEH) and it's emerging in new countries is not fully understood(1).this disease is a mosquito-borne infection in which it doesn't transmitted directly from human to human (2). It is an important disease in areas where mosquitoes breed under optimal conditions (3). There are three form of the dengue disease, dengue fever (DF), dengue hemorrhagic fever (DHF) or dengue shock syndrome (DSS)(1). The classical dengue fever is mild from and usually patient recovered shortly after illness. However, I some cases (DF) can lead to DHF or DSS which can be life threatening(4). The mosquito is the main transmitted vector of dengue virus to humans through the bites of infected female of *Aedes aegypti* and *Aedes albopictus*. The transmission of dengue Virus has increased dramatically in recent years and spreading in urban and semi-urban endemic settings, especially in Americas, Western Pacific, South and South-east Asia and the (5).Infection with one serotype give memory immunity for the same serotype long life but not for the other serotypes. (6).

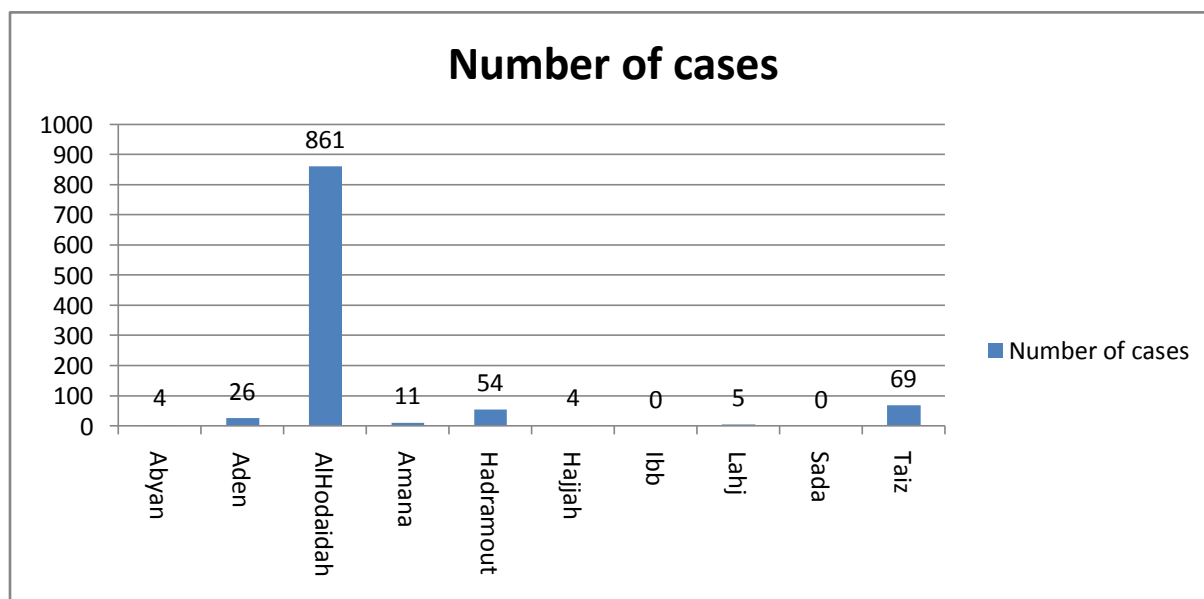
## ETIOLOGY:

Dengue virus was first isolated in 1943(7). It is RNA virus belong to *Flaviviridae* family which cause the most common arboviral, vector born disease(8). There are four closely dengue virus serotypes, dengue virus-1 to dengue virus-4. The closest antigenic relationship exists between type1 and 4. Dengue virus has four serotypes dengue virus from 1-4. Sequencing of dengue viral RNA has further verified strain variation within a serotype allowing viruses to be classified into genetically distinct groups within serotypes called genotypes (9). Based on the analysis of the envelope protein there are at least four genotypes (1 to 4) (3).

## DISTRIBUTION:

Dengue virus was widely distributed in Asia, Pacific, Medial East, Central and South America and Africa(10). In Yemen, dengue was first recorded in pandemic outbreak reported by Hirsch between 1870 and 1873(11). After that the pandemic dengue fever occur in Alhodeidah governorate in 1954 which affected 98% from the population and in 1984 traveler returned from Yemen confirmed to have dengue antibodies(12). Dengue has been a significant health problem especially in the coastal areas. Dengue fever was first reported and confirmed in Shabwah governorate and it widely distributed in nautical governorates as see in figure2(13, 14). Aden, Alhodeidah and Almukalla is the highest governorate affected by dengue fever in 2014(14). however the highest governorate was Aden, Alhodeidah and Taiz as in figure. Dengue virus serotype 3 was isolated from an outbreak of viral hemorrhagic fever in Al-Mukalla city Hadaramut governorate republic of Yemen. This study recommended using IgM and NS1-antigen tests to confirm acute dengue particularly under the adverse field conditions, where proper storage and transportation of specimens are missing, which substantially reduce the sensitivity of the RT-PCR for detecting dengue RNA(15). Another study, conducted in Taiz governorate, showed high prevalence of Pulmonary manifestations among the complicated cases of dengue fever either Dengue hemorrhagic fever or dengue shock syndrome (2). Moreover DENV 1 was confirmed in Alhodeidah governorate whereas DENV 3 was confirmed in Hadramout governorate in 2010 outbreak (16, 17). all three dengue viral serotyp1-3 was confirmed to be in Yemen by using PCR but Dengue virus type 2 was the predominant serotype in Yemen(18).

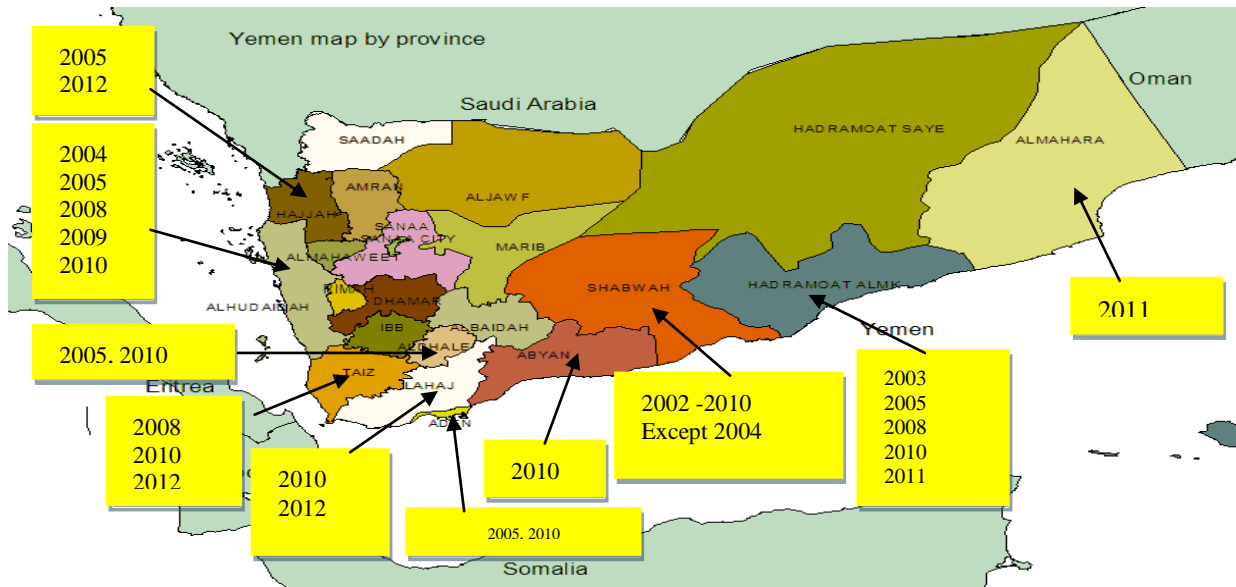
In Yemen, climate variability and change combined with demographic and socioeconomic factors (such as population increase social instability and poverty) may all aggravate the problem posed by dengue. Over the last decade, the temperatures and rainfall on the Arabian Peninsula have been unusually high, with severe floods in Yemen, Saudi Arabia and Oman. Such conditions are not only likely to favor mosquito breeding perhaps explaining the epidemics and high incidences of dengue fever recently observed in Yemen (19, 20) but may also allow mosquito vectors to move up to altitudes that were once too cool for their survival(21). Al-hodeidah governorate was the epidemic foci for several vector-borne outbreaks in the past including dengue fever in 2005 and 2012 (22).



**Figure 1** Distribution of dengue fever by governorate from week (1-24) in 2014 (14).

## RESERVOIR:

Human/ Mosquitoes, *Aedes aegypti* and *Ae. Albopictus*, cycle were the main reservoir for the Dengue virus whereas the Zoonotic cycle includes monkey/ mosquito *Aedes* species like *Ae. Scutellaris*, *Ae. Niveaus* and *Ae. fuscifer-taylori* complex in Polynesia, Malaysia and Africa respectively are involve to monkey/mosquito transmission(23). Non human primate don't seem to represent a significant reservoir of Dengue virus(24).



**Figure 3** Reported Dengue/CHIK Outbreaks in Yemen (13, 14).

## INCUBATION PERIOD AND PERIOD COMMUNICABILITY:

Incubation period ranged from 3-14 days. The mosquito will be able to transmitted the disease after 8-18days after getting viremic blood meal and remain so for life(23).

## CLINICAL SIGNS:

The dengue fever has three clinical manifestation forms. The mild form is acute febrile illness which characterized by headache, retro-ocular pain, body aches and arthralgia in more than 90% of apparent cases, combined by nausea or vomiting, amaculopapular rash, fever remitting, relapse, pain in the muscles and bones so it called 'break bone fever'(25). DHF/DSS forms of dengue fever are, the more severe forms, but it is less common dengue fever. It is characterized by fever, thrombocytopenia $<20 \times 10^3$ , hemorrhagic manifestation and evidence of increased vascular permeability with leakage of intravascular fluid into the intestinal space (26-28). The DHF forms mainly affected the children and is occasionally accompanied by a shock syndrome with case fatality 50% (25). The sever forms of dengue infection usually showed after infection by more than one virus serotypes ,e.g. serotype 1 and serotype 2, which characterized by fever vomiting enlarged of liver petechiae persist for 5-10days (25, 29). All three forms of dengue fever have been reported in Yemen with most common manifestation as shown in figure 3 (15, 16).

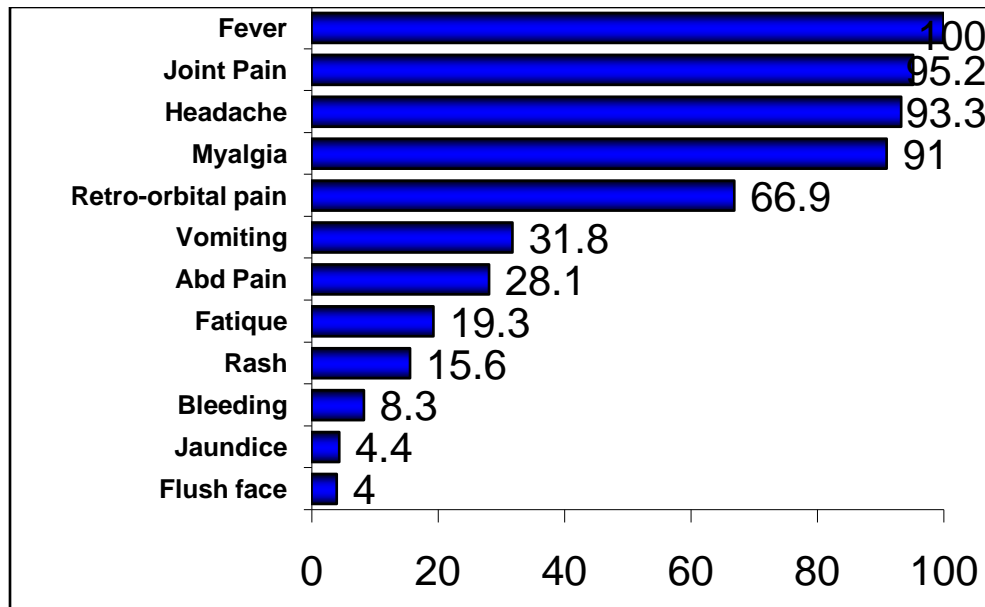


Figure 3 Most symptoms recorded in suspected dengue fever in Yemen (14).

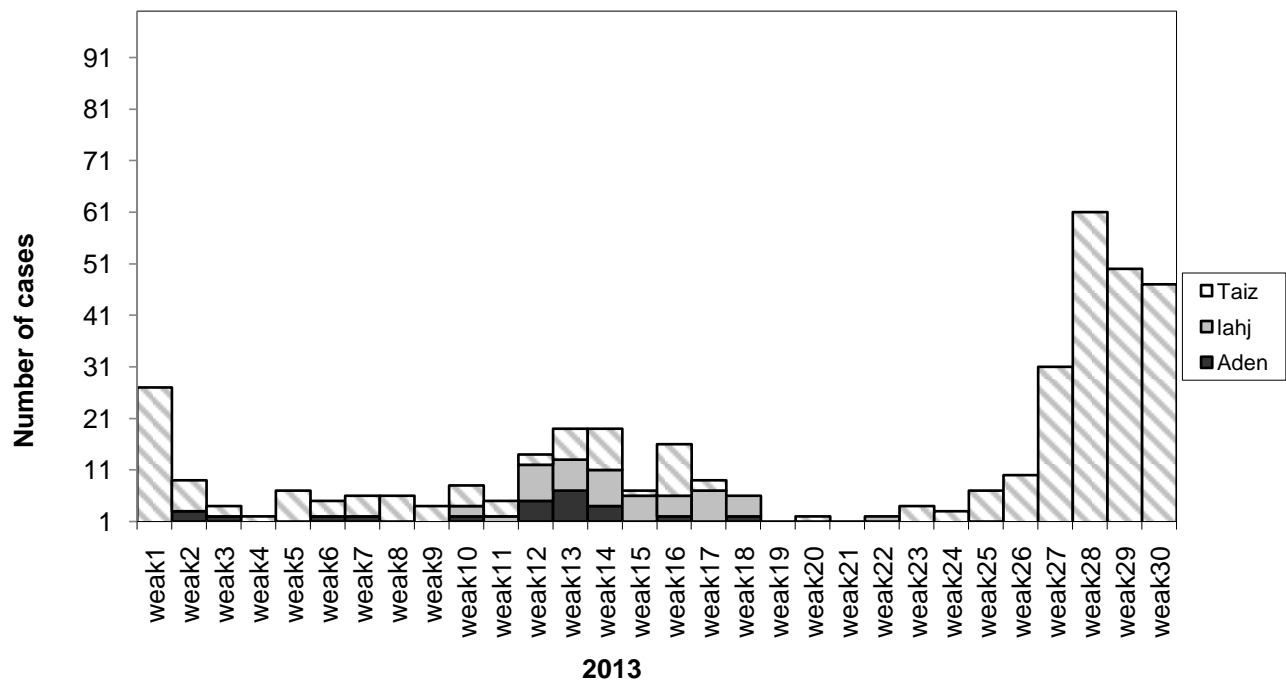


Figure 2 Epidemiological curve of dengue fever in first 30 weeks of 2013 at Taiz, lahj and Aden governorates (14).

## DIAGNOSIS:

Diagnosis of dengue fever is mainly depending on clinical symptoms, hypovolemia and isolation of the virus(30). Diagnosis of DENV serotyping was depending on IgG or IgM ELISA technique. Recently RT-PCR was used to diagnosis the four serotypes of DENV in connection with group specific primers(31, 32). One step RT-PCR was also developed for rapid typing of different dengue virus serotypes(4). In Yemen diagnosis still based on IgM ELISA. Blood specimens were tested according to WHO (1997) guideline for confirmation using IgG and IgM-ELISA at Yemeni Ministry of Public Health and Population (MOPHP) and other samples was Sent to Thai National Institute of Health or tested at U.S.NAMRU-3 for confirming stereotypes and genotype by RT-PCR(16, 33). DENV 1 was confirmed in 2011 outbreak in Alhodeidah governorate which showed 4(2.9%) from suspected cases (34). The differential diagnosis from other likely febrile diseases should be considered and differentiated depending on clinical sing and local prevalence of that diseases which including other arboviral diseases, measles, rubella, adenovirus infection, enterovirus infection, viral hepatitis, typhoid, malaria, leptospirosis and bacteria sepsis(35).

## TREATMENT:

There is no specific antiviral therapy used for treatment of dengue fever. However, replacement of fluid, blood transfusion and plasma protein transfusion are essential for treatment of DHF and DSS cases with continuous testing of electrolytic(30).

## CONTROL AND PREVENTION:

The general strategy for control dengue fever is a vector control which including physical controlling by elimination of specific breeding area of *Aedes spp.*, landscape management and continuous surveillance risk area(36). Biological control strategy was also used to control *Aedes aegypti* like using carnivorous fish (*Gambusia affinis*) and personal protection methods like covering skin with clothes at dusk and repellent(30, 36, 37). Chemical controlling of mosquitoes like insecticide and larvicides is useful in outbreak to reduce incidence rate of vector borne diseases(38). treated bed net was used also for controlling of mosquitoes which including 2% permethrin incorporated into polyethylene fibers of bed net which showed effectiveness in reduction of mosquito population in Haiti (39). Surveillance using remote sensing technique and satellite mapping are useful methods for prevent of dengue fever outbreak's before it happened(40). The vaccine with tetravalent attenuated live virus is the prophylaxis methods for dengue fever. This vaccine is inducing neutralizing antibody for all four serotypes. However this vaccine doesn't sufficiently tested(30). Community health education for prevention and controlling procedure of dengue fever has significant impact to reduce risk of dengue fever disease(41).

## CONCLUSION:

Dengue fever was spreading in new governorates in Yemen in last ten years. The large scale of dengue outbreak in Alhodeidah governorate is due to the high vector circulation (*Aedes aegypti*), poor infrastructure, and excesses of raining period more than other cities in Yemen, ineffective vector and infectious disease surveillance and low level of education are the main reason for the increase incidence rate of dengue fever in Yemen. In conclusion, dengue fever was spreading in different region in Yemen with high distribution in Al-Hodeidah, Taiz, Aden and Al-Mukalla. More vector controlling and preventive measurement and management should be considered. Sentinel-based surveillance systems for acute febrile illness should be enhanced in areas with high incidence of malaria and dengue fever. More genotyping, biodiversity, molecular characterization and climate changing studies needed to be conducted on dengue virus in Yemen.

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