# Chikungunya virus infection in travellers to Australia

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hikungunya virus is a mosquito-borne alphavirus infection first described in Tanzania in 1952. The name Chikungunya means "that which bends up" in the Makonde language of the Newala Province, Tanzania.

We report eight cases of Chikungunya virus infection in travellers to Australia between February 2006 and September 2007. As clinicians became aware of the Indian Ocean epidemic, unwell travellers returning from affected countries were tested for Chikungunya virus. Before this epidemic, only one case of Chikungunya virus infection had been reported in Australia; this involved importation of the virus from Indonesia to Darwin in 1989. <sup>1</sup>

## **Data extraction**

The eight cases of Chikungunya virus infection were identified after reviewing the database of the Victorian Infectious Diseases Reference Laboratory between February 2006 and September 2007. The review was undertaken to raise awareness among medical practitioners of Chikungunya virus infection. Chikungunya virus was diagnosed by performing a generic alphavirus reverse transcriptase polymerase chain reaction (RT-PCR) on serum.<sup>2</sup> The diagnosis was confirmed in all eight patients by sequencing the alphavirus RNA product.

The treating clinicians of the eight patients were contacted. The medical records and investigations performed for each patient were analysed. Seven patients presented to tertiary referral hospitals in Melbourne. One patient presented to a local doctor in regional Victoria.

#### Clinical records

The first patient diagnosed with Chikungunya virus infection was a 59-year-old Mauritian man who presented to hospital 5 days after travelling from Mauritius for the 2006 Commonwealth Games. He reported a 2-day history of fevers, rigors, and headaches with associated arthralgia, conjunctivitis (Box 1), and left lower limb erythema. Investigations revealed leukopenia, thrombocytopenia, and mildly abnormal liver function tests. Blood taken at admission was positive for alphavirus RT-PCR, and sequencing confirmed Chikungunya virus. The patient received supportive treatment and had fully recovered on review 1 week later. A detailed description of his illness has been reported previously.<sup>2</sup>

Six subsequent patients had travelled from Sri Lanka, and the most recent patient travelled from India in June 2007 (Box 2). All patients described fevers, rigors and headaches. Six patients presented with arthralgia, five with rash, and two had conjunctivitis. Investigations revealed six patients had leukopenia, three had thrombocytopenia, and three patients had abnormal liver function tests with raised serum alanine aminotransferase levels. All patients recovered. Clinician awareness of the outbreak resulted in early diagnosis for all eight patients.

Acute serology was performed on four patients. All were IgG and IgM negative. Seroconversion was demonstrated with convalescent serology performed on one patient 8 weeks after presentation.

#### **ABSTRACT**

- We report eight recent cases of Chikungunya virus infection in travellers to Australia. Patients presented with fevers, rigors, headaches, arthralgia, and rash.
- The current Indian Ocean epidemic and Italian outbreak have featured prominently on Internet infectious disease bulletins, and Chikungunya virus infection had been anticipated in travellers from the outbreak areas.
- Diagnosis was by a generic alphavirus reverse transcriptase polymerase chain reaction with confirmatory sequencing.
- Prompt diagnosis of Chikungunya virus infections is of public health significance as the mosquito vectors for transmission exist in Australia. There is potential for this infection to spread in the largely naïve Australian population.

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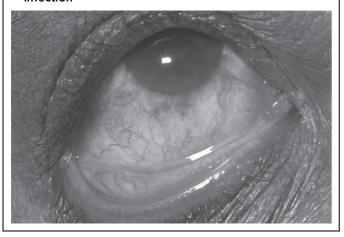
### **Discussion**

These eight patients illustrate the typical clinical presentation of Chikungunya virus infection. The illness is of abrupt onset, characterised by high fevers accompanied by rigors, arthralgia or arthritis, myalgia and headache. A maculopapular rash occurs on the second to fifth day in 50% of patients. The incubation period ranges from 2 to 12 days, and the febrile period lasts typically 3–7 days. Patients generally make a full recovery. Patients may experience persistent joint symptoms. Serious complications are rare. Neuroinvasive disease (encephalitis) and deaths have been reported during the outbreak in Réunion. Laboratory tests reveal a mild leukopenia and relative lymphocytosis. No specific drug therapy is available, and treatment is supportive. Chikungunya virus infection is believed to confer lifelong immunity. No licensed Chikungunya virus vaccine is currently available.

Laboratory diagnosis in these patients was by detecting viral RNA by RT-PCR. The viraemic phase of alphaviruses is 2–8 days, so the success of RT-PCR to detect the virus depends on early presentation and the clinical suspicion of a possible alphavirus infection. Diagnosis of Chikungunya virus infection may be made serologically at major virology reference laboratories (Westmead Hospital, Sydney, NSW; PathWest Laboratory Medicine WA, Perth, WA; and Queensland Health Scientific Services, Brisbane, QLD).

Chikungunya virus is an arthropod-borne alphavirus classified within the family Togaviridae. It is geographically distributed in Africa, India, and South-East Asia, and periodic outbreaks have been described in the past 50 years. The most recent epidemic began in Kenya in 2004. By early 2005, there was an outbreak in the Comoro Islands, which promptly spread to other Indian Ocean islands: Réunion, Mauritius, Seychelles and Madagascar (Box 3). As of September 2006, Réunion had an estimated 266 000 cases, including a reported 248 deaths (population, 770 000). The epidemic subsequently spread to India, Sri Lanka, the Maldives and Malaysia, with reports of imported cases in Europe, the United States, and Hong Kong. 5,9-11 The outbreak is ongoing in India,

# 1 Conjunctivitis in a patient with Chikungunya virus infection



with an estimated 1.4 million cases reported and attack rates as high as 45%.  $^{4,12-14}$ 

In August 2007, an outbreak of Chikungunya virus infection occurred in the Ravenna district in Italy. As of 13 September 2007, 254 cases had been reported. The first case was attributed to a traveller from India. This outbreak demonstrates the vulnerability of receptive countries to Chikungunya virus emergence. The mosquito vector, *Aedes albopictus*, is present in at least 12 European countries.<sup>15</sup>

This infection has potential implications for Australia. An outbreak similar to that seen in Italy could occur in Australia, and

medical practitioners should consider the possibility of local transmission of Chikungunya virus from an infected patient if imported disease occurs. They should take measures to minimise that risk, particularly if a viraemic patient is at risk of being bitten by mosquitoes. Mosquitoes are the vector for Chikungunya virus, and humans are the major reservoir during epidemics. The mosquito Aedes aegypti has been the major vector in India, Asia and Africa, and A. albopictus is the likely vector in Réunion and Italy. 4,15,16 Chikungunya virus may also be spread by other mosquitoes of the genera Aedes and Culex. The known vectors are present in Australia — A. aegypti is found in northern Queensland and A. albopictus has been recovered at ports around Australia, 17,18 so it may be of particular importance that infected travellers be diagnosed promptly in these locations and isolated from mosquito exposure early in their illness. No local transmission has yet been reported in Australia.

Globalisation and ease of travel means these and other exotic infections will present increasingly frequently. The arrival of Chikungunya virus was anticipated through an awareness of the Indian Ocean epidemic. There is potential for other emerging arboviruses to become established in Australia. Clinical awareness is essential to recognise these new infections, and Internet infectious disease bulletins efficiently facilitate this. A detailed travel history, and consultation with microbiologists and virologists to guide appropriate investigation are required for diagnosis of these emerging pathogens.

With the Chikungunya virus epidemic ongoing in India, it is likely we will see more cases in Australia. The clinical features of Chikungunya virus infection can be similar to dengue fever and malaria, and these infections should also be considered in the

Age, sex	Year of presentation	Country of acquisition	Presenting symptoms	Laboratory findings	Outcome
59 M	2006	Mauritius	2-day history of fever, headache, rigors, arthralgia, conjunctivitis, rash	RT-PCR positive Thrombocytopenia Leukopenia Raised ALT	Fully recovered at 1 week
29 F	2006	Sri Lanka	3-day history of fever, headache, rigors, arthralgia, rash	RT-PCR positive Leukopenia	Fully recovered at 1 week
24 M	2006	Sri Lanka	1-day history of fever, headache, rigors, rash	RT-PCR positive Leukopenia Raised ALT	Fully recovered by Day 3
6 M	2006	Sri Lanka	1-day history of fever, headache, rigors, rash, arthralgia, conjunctivitis	RT-PCR positive Thrombocytopenia	Mild ankle pain at 1 week
50 F	2006	Sri Lanka	1-day history of fever, headache, rigors, rash	RT-PCR positive Leukopenia Raised ALT	Fully recovered at 3 weeks
55 M	2007	Sri Lanka	2-day history of fever, headache, rigors, arthralgia	RT-PCR positive Leukopenia	Fully recovered at 4 weeks
50 M	2007	Sri Lanka	2-day history of fever, headache, rigors, arthralgia	RT-PCR positive Thrombocytopenia Leukopenia	Fully recovered at 2 weeks
25 F	2007	India	1-day history of fever, headache, rigors, arthralgia	RT-PCR positive	Fully recovered at 1 week

RT-PCR positive = positive result on generic alphavirus reverse transcriptase polymerase chain reaction testing. Leukopenia = leukocyte count  $< 4.0 \times 10^9$ /L. Raised ALT = serum alanine aminotransferase level > 56 U/L.



differential diagnosis of febrile, returned travellers. Clinicians should be alert for cases of Chikungunya virus infection in returned travellers, and consider measures to minimise the potential of local transmission through infection of local mosquitoes by a person with viraemia, especially in northern Australia.

### **Competing interests**

None identified

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