Following the identification of two autochthonous cases of dengue type 1 on 3 October 2012, an outbreak of dengue fever has been reported in Madeira, Portugal. As of 25 November, 1,891 cases have been detected on the island where the vector *Aedes aegypti* had been established in some areas since 2005. This event represents the first epidemic of dengue fever in Europe since 1928 and concerted control measures have been initiated by local health authorities.

Madeira is the main island of the Autonomous Region of Madeira, a Portuguese territory with 262,302 inhabitants [9]. It is located at around 1,000 km from mainland Portugal and around 500 km from the northern African coast. The other inhabited island of the Region is Porto Santo with 5,483 inhabitants [9]. The climate is temperate Mediterranean, with little temperature variation throughout the year [10]. *Ae. aegypti* was first identified on Madeira in 2005, in Santa Luzia parish of Funchal city [11].

**Epidemiological and clinical findings**

After the detection of the first two autochthonous dengue fever cases on 3 October 2012, active case finding was initiated and the following case definition has been used since:

A probable case should meet both clinical and epidemiological criteria outlined hereafter.

Clinical criteria comprised acute onset of fever and at least two of the following symptoms and signs: headache, retro-orbital pain, myalgia, arthralgia, exanthema, haemorrhagic manifestations or leucopenia.

Epidemiological criteria are residing in or having visited a dengue-affected area during the 21 days prior to onset of symptoms. Currently, the whole island is considered as ‘affected area’ which means that the epidemiological criterion is met for all cases occurring on Madeira.

A confirmed case is defined as a probable case with at least one of the following laboratory results: (i) presence of dengue-virus-specific IgM antibodies in blood or cerebrospinal fluid (CSF); (ii) significant increase in the concentration of dengue-virus-specific IgG antibodies (seroconversion), or; (iii) detection of dengue virus antigen in blood or cerebrospinal fluid.
virus nucleic acid in blood or CSF with RT-PCR or other Nucleic Acid Amplification Test (NAAT).

Pregnant women, patients with chronic diseases and with immunodeficiency were considered at particular risk of getting the disease and of possible complications.

As of 25 November 2012 (calendar week 47), seven weeks after the first autochthonous cases were identified, 1,891 probable cases of dengue fever have been reported, of which 966 were laboratory confirmed. Of the total number of 1,891 cases, 777 (41.1%) were male and 1,114 (58.9%) were female. The median age of the 1,891 cases was 39 years (range 25-64). Up to week 47, the most affected age group was 25 to 64 years-old (Table).

One hundred and eleven patients were hospitalised and no fatalities were recorded. Of the total number of cases (n=1,891), the majority had mild symptoms, with fever, myalgia, headaches and arthralgia.

Current geographic distribution of cumulative dengue cases by parish of residence is presented in Figure 2. The highest proportion of patients is recorded in residents of Santa Luzia parish.

As of 25 November, 29 cases of dengue fever were reported among travellers returning from Madeira. Twelve cases of dengue fever were recorded in travellers to mainland Portugal. In addition, 17 cases were detected in travellers returning to other European countries: United Kingdom (n=6), Germany (n=7), Sweden (n=1), France (n=2) and Finland (n=1).

Laboratory investigation
Sequence analyses of viral genomes (600 nucleotides), performed at the National Institute of Health Dr. Ricardo Jorge, Lisbon, Portugal, presented high sequence similarity with DEN-1 viruses circulating in Venezuela and Colombia, suggesting Latin America as the possible origin for the circulating dengue virus [12].

Entomological findings
In 2006, a mosquito surveillance system based on ovitraps, traps that collect eggs by mimicking breeding sites, was established to monitor the presence and the abundance of the vector on the island [13]. By 2009, Aedes aegypti was detected outside Funchal city in the neighbouring localities of Câmara dos Lobos and Caniço. In September 2011, cross-sectional entomological surveys were carried out using a total of 273 ovitraps dispersed throughout Madeira (253 ovitraps) and Porto Santo islands (20 ovitraps). Results showed that the mosquito population had expanded to the west (Ponta do Sol and Ribeira Brava counties) and east of Funchal city (Santa Cruz county). The vector was also detected in Porto Moniz, but its presence was not confirmed in subsequent monitoring activities (unpublished data).

After the onset of the outbreak in October 2012, the surveillance network was reinforced and, currently, mosquito activity is monitored by 141 ovitraps and 16 BG-sentinel traps, traps baited with attractants to biting females. These traps are distributed mainly along the south coast of the island and in selected areas such as harbours, the airport, healthcare facilities and potential transmission zones. Up to date the vector has not been found on Porto Santo.

Control measures
In response to the outbreak, several control measures have been undertaken by the health authorities of Madeira. In order to prevent exportation, enhanced vector control measures have been implemented at Madeira’s single airport. Currently, all aircrafts departing from the island undergo disinestation procedures. Mosquito surveillance systems at the airport and at

<table>
<thead>
<tr>
<th>Age group (years)</th>
<th>Number of cases</th>
<th>Percentage (%)</th>
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<tbody>
<tr>
<td>0-14</td>
<td>240</td>
<td>12.7</td>
</tr>
<tr>
<td>15-24</td>
<td>299</td>
<td>15.8</td>
</tr>
<tr>
<td>25-64</td>
<td>1,132</td>
<td>59.9</td>
</tr>
<tr>
<td>65+</td>
<td>220</td>
<td>11.6</td>
</tr>
<tr>
<td>Total</td>
<td>1,891</td>
<td>100</td>
</tr>
</tbody>
</table>
passenger and cargo harbours have been boosted. Particular care is being given to the passenger ship port, located in the most affected area, Funchal. Information, mainly focused on the use of personal protective measures and on the elimination of mosquito breeding sites [14], is being disseminated by several mechanisms: (i) individual information to residents through a door-to-door campaign carried-out by environmental technicians of the IASAUDE, RAM; (ii) postings on IASAUDE website and in traditional and social media, and; (iii) community-based activities involving the educational, religious and health sectors. Advice for travellers to Madeira has been issued, containing information about methods to reduce the risk of being bitten by mosquitoes and symptoms of the disease [15].

Guidelines for diagnostic and management of dengue fever were provided to healthcare professionals. Measures to ensure blood safety and the safety of substances of human origin have been implemented and published since the detection of the first cases [16].

Nationwide measures to ensure blood safety in mainland Portugal included deferral of potential blood donors who visited Madeira island for a period of 28 days after leaving the affected area, deferral of potential blood donors with fever or influenza-like symptoms for 28 days after recovery, deferral of confirmed cases of dengue infection for 120 days after recovery and post transfusion haemovigilance.

Local measures applying to the blood services of the affected area included: (i) quarantine of all red cell concentrates resulting from blood collected during the previous 28 days and retrospective testing of relevant archived samples; (ii) deferral of blood donors with fever or influenza-like symptoms up to 28 days after recovery; (iii) deferral of confirmed dengue cases for 120 days after diagnosis; (iv) laboratory screening of all blood donations by RT-PCR for dengue virus; (v) information to blood donors to report any symptoms in the 15 days after donation; (vi) stop of local platelet production and supply of platelets from the Portuguese Institute for Blood and Transplants (Instituto Português de Sangue e Transplantes – IPST). Special attention is paid to the optimal use of blood, appropriate management of the blood supply and haemovigilance.

Discussion
The 2012 dengue outbreak on Madeira is the first of its kind in Europe since 1928. Given its magnitude and...
its potential public health impact in a European context, the European Centre for Disease Prevention and Control (ECDC) in Stockholm, Sweden, has, on request of the Portuguese authorities, deployed experts to Madeira to assist the regional and national authorities in investigating and controlling the outbreak and prevent further expansion within Europe.

While *Ae. aegypti* is not considered an established mosquito species in continental Europe, its presence has been sporadically reported in restricted areas in the Netherlands and Russia [17]. The present outbreak however, draws attention to the need to better assess the possibility of dengue epidemics mediated by *Ae. albopictus*, that is present in several European countries. Besides the need to reinforce strategies aimed at averting the introduction of *Ae. aegypti* in Europe [18], it is also important to detect and monitor all imported human cases that might introduce the virus among a very competent mosquito species, *Ae. albopictus*.

At the time of publishing this report, the outbreak on Madeira is still ongoing. However, in week 48, the number of cases reported has decreased by around 54% (from 219 cases to 102 cases) when compared with the previous week. According to the weather forecast for the coming winter weeks, Madeira expects a decrease in temperature. This is assumed to have an impact on the vector, reducing its densities and, consequently, the number of dengue cases might further decline. Nevertheless, even in the absence of new reported human cases the virus may persist on the island either through vertical transmission or maintained by the very competent mosquito species, *Ae. albopictus*.

Given the challenges posed by the dengue outbreak on Madeira, continued vigilance and a coordinated regional, national and international approach are needed.

References