# ation ZIKA VIRUS MICROCEPHALY AND GUILLAIN-BARRÉ SYNDROME

SITUATION REPORT 26 FEBRUARY 2016

### **SUMMARY**

- Between 1 January 2007 and 25 February 2016, a total of 52 countries and territories have reported autochthonous (local) transmission of Zika virus, including those where the outbreak is now over and countries and territories that provided indirect evidence of local transmission. Among the 52 countries and territories, Marshall Islands, Saint Vincent and the Grenadines, and Trinidad and Tobago are the latest to report autochthonous transmission of Zika virus.
- The geographical distribution of Zika virus has steadily widened since the virus was first detected in the Americas in 2015. Autochthonous Zika virus transmission has been reported in 31 countries and territories of this region. Zika virus is likely to be transmitted and detected in other countries within the geographical range of competent mosquito vectors, especially *Aedes aegypti*.
- So far an increase in microcephaly cases and other neonatal malformations have only been reported in Brazil and French Polynesia, although two cases linked to a stay in Brazil were detected in two other countries.
- During 2015 and 2016, eight countries and territories have reported an increased incidence of Guillain-Barré syndrome (GBS) and/or laboratory confirmation of a Zika virus infection among GBS cases.
- Evidence that neurological disorders, including microcephaly and GBS, are linked to Zika virus infection remains circumstantial, but a growing body of clinical and epidemiological data points towards a causal role for Zika virus.
- The global prevention and control strategy launched by WHO as a Strategic Response Framework<sup>1</sup> encompasses surveillance, response activities and research, and this situation report is organized under those headings. Following consultation with partners and taking changes in caseload into account, the framework will be updated at the end of March 2016 to reflect epidemiological evidence coming to light and the evolving division of roles and responsibilities for tackling this emergency.

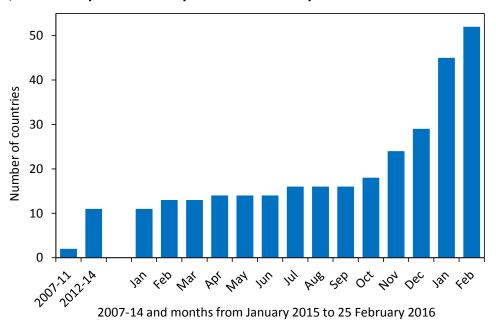
<sup>&</sup>lt;sup>1</sup> Zika Strategic Response Framework & Joint Operations Plan: <a href="http://apps.who.int/iris/bitstream/10665/204420/1/ZikaResponseFramework\_JanJun16\_eng.pdf?ua=1">http://apps.who.int/iris/bitstream/10665/204420/1/ZikaResponseFramework\_JanJun16\_eng.pdf?ua=1</a>

### I. SURVEILLANCE

## **Incidence of Zika virus**

■ From 1 January 2007 to 25 February 2016, Zika virus transmission was documented in a total of 52 countries and territories (Fig. 1 and Fig. 2). This includes 40 countries that reported autochthonous transmission between 2015 and 2016, six countries with indirect evidence of viral circulation, five countries with reported terminated outbreaks and one country with a locally acquired infection in the absence of any known mosquito vectors (United States of America; Table 1).

Figure 1: Cumulative number of countries, territories and areas reporting Zika virus transmission, 2007-2014, and monthly from 1 January 2015 to 25 February 2016.



- Towards the end of 2014, Brazil detected a cluster of cases of febrile rash in the Northeast Region of the country. The diagnosis of Zika virus infection was confirmed (RT-PCR test for viral RNA²) in May 2015. The Brazilian Ministry of Health estimates that there were 0.4-1.3 million cases of Zika virus infection in 2015, many more than were reported or confirmed.³
- Recently the virus has spread rapidly across the region. By 25 February 2016, 31 countries and territories in the Americas had reported local transmission of the virus. The reported rate of its spread across South and Central America accelerated from October 2015 onwards (Fig. 1, Table 1).

<sup>3</sup> The full report is available in Portuguese at: <a href="http://portalsaude.saude.gov.br/images/pdf/2016/janeiro/22/microcefalia-protocolo-de-vigilancia-e-resposta-v1-3-22jan2016.pdf">http://portalsaude.saude.gov.br/images/pdf/2016/janeiro/22/microcefalia-protocolo-de-vigilancia-e-resposta-v1-3-22jan2016.pdf</a>

<sup>&</sup>lt;sup>2</sup> Reverse transcriptase polymerase chain reaction (RT-PCR).

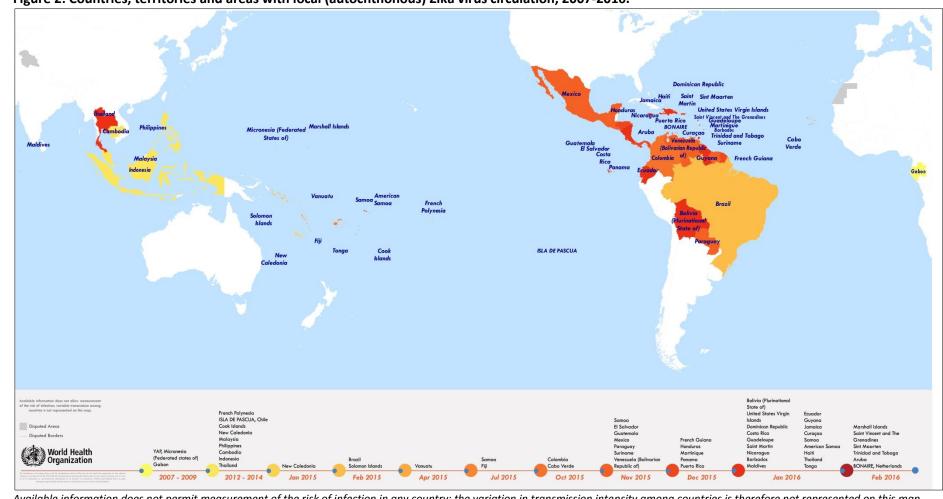


Figure 2: Countries, territories and areas with local (autochthonous) Zika virus circulation, 2007-2016.<sup>4</sup>

Available information does not permit measurement of the risk of infection in any country; the variation in transmission intensity among countries is therefore not represented on this map. Zika virus is not necessarily present throughout the countries/territories shaded in this map.

<sup>&</sup>lt;sup>4</sup> http://www.who.int/emergencies/zika-virus/situation-report/19-february-2016/en/

Figure 3. Countries, territories and areas reporting Zika virus, microcephaly and Guillain-Barré syndrome, 2013-2016. Puerto Rico El Salvador Martinique Venezuela (Bolivarian Republic of) HAWAII Suriname French Polynesia Disputed Areas Disputed Borders Reported Zika virus with increased incidence of microcephaly (no increased incidence of GBS) World Health Organization Reported Zika virus with increased incidence of GBS and microcephaly Reported Zika virus with increased incidence of GBS (no increased incidence of microcephaly) Reported Zika virus (no increased incidence of GBS or microcephaly)

Table 1. Countries, territories and areas with autochthonous transmission of Zika virus, 2007–2016.\*

	WHO Regional Office	Country or territory or area <sup>#</sup>
	AFRO (n=1)	Cabo Verde
Reported autochthonous transmission <sup>a</sup> (n=40)	AMRO/PAHO (n=31)	Aruba, Barbados, Bolivia, Bonaire, Brazil, Colombia, Costa Rica, Curaçao, Dominican Republic, Ecuador, El Salvador, French Guiana, Guadeloupe, Guatemala, Guyana, Haiti, Honduras, Jamaica, Martinique, Mexico, Nicaragua, Panama, Paraguay, Puerto Rico, Saint Martin, Sint Maarten, Saint Vincent and the Grenadines, Suriname, Trinidad and Tobago, United States Virgin Islands, Venezuela
	SEARO (n=2)	Maldives, Thailand
	WPRO (n=6)	American Samoa, Marshall Islands, Samoa, Solomon Islands, Tonga, Vanuatu
Indication of viral circulation <sup>b</sup> (n=6)	AFRO (n=1)	Gabon
	SEARO (n=1)	Indonesia
	WPRO (n=4)	Cambodia, Fiji, Philippines, Malaysia
Countries/territories/areas with outbreaks terminated (n=5)	AMRO/PAHO (n=1)	Isla de Pascua – Chile
	WPRO (n=4)	Cook Islands, French Polynesia, New Caledonia, Yap – Micronesia (Federated States of)
Locally acquired without vector borne transmission <sup>d</sup> (n=1)	AMRO/PAHO (n=1)	Texas – United States of America

<sup>\*</sup>Available information does not permit qualification of the intensity of viral circulation and therefore the risk of infection; the situation is extremely variable according to countries, and this information should be used with caution.

- From 1 October 2015 to 13 February 2016, Colombia reported 37 011 cases, including 1612 laboratory confirmed cases of Zika virus infection.<sup>5</sup>
- From 2007, locally acquired Zika cases have been reported in 14 countries and territories in the Western Pacific Region. Four Pacific Island countries and areas (American Samoa, Marshall Islands, Samoa and Tonga) have reported Zika infections in 2016. Nauru has

 $<sup>^{\#}</sup>$  For overseas territories/countries/provinces or islands, the affected area rather than the country is reported.

<sup>&</sup>lt;sup>a</sup> Reported autochthonous transmission: Formal notification through IHR, of at least one (1) case of autochthonous transmission by the affected Member State or the Member State where the diagnosis has been performed (for travellers). Autochthonous infection is considered to be any infection acquired in the country i.e. among patients with no history of travel during the incubation period or travels exclusively to non-affected areas.

b Indication of viral circulation: Indirect information of at least one Zika biologically confirmed case (by RT-PCR or sero-neutralisation) either diagnosed domestically or exported and diagnosed abroad.

<sup>&</sup>lt;sup>c</sup> Countries, territories or areas with outbreaks terminated: Countries or territories where the interruption of the viral circulation has be documented can through the surveillance data (including syndromic surveillance, laboratory confirmation of suspected cases, etc.) and/or where no suspect case has been reported since 31 December 2014.

<sup>&</sup>lt;sup>d</sup> Locally acquired without vector-borne transmission: Autochthonous infection but through another mode of transmission than vector borne (including sexual, blood-borne, or organ transplant) and for where vector population is unlikely to allow sustained vector borne transmission.

<sup>&</sup>lt;sup>5</sup> Full report available at: <a href="http://www.ins.gov.co/boletin-epidemiologico/Boletn%20Epidemiologico/2016%20Boletin%20epidemiologico%20semana%206.pdf">http://www.ins.gov.co/boletin-epidemiologico/Boletn%20Epidemiologico/2016%20Boletin%20epidemiologico%20semana%206.pdf</a>

- declared Zika virus as a national emergency of concern for purposes of preparedness, but to date no ZIKV cases have been reported.
- From 1 October 2015 to 7 February 2016, Cabo Verde (African region), reported 7325 suspected cases of Zika virus disease. The outbreak peaked during the last week of November 2015 and has been on a steady decline since then; 67 cases were reported in the week up to 7 February 2016. Preliminary information, subject to confirmation, indicates that this outbreak has been caused by an African strain of Zika virus. No neurological abnormalities have been reported.

## **Incidence of microcephaly**

Between 22 October 2015 and 20 February 2016 a total of 5640 cases of microcephaly and/or central nervous system (CNS) malformation have been reported by Brazil including 120 deaths. This contrasts with the period from 2001 to 2014, when an average of 163 microcephaly cases was recorded nationwide per year

Table 2. Countries, territories and areas reporting microcephaly cases potentially associated with Zika virus infection.

Reporting country	Number of reported microcephaly cases potentially related to a Zika virus infection	Probable location of infection
French Polynesia	9	French Polynesia
Brazil	583	Brazil
Hawaii (United States of America) <sup>6</sup>	1	Brazil
Slovenia <sup>7</sup>	1	Brazil

- The reported increase in microcephaly incidence in Brazil is concentrated in the Northeast Region (Fig. 3).
- Of the 5640 suspected cases of microcephaly reported in Brazil, investigations have been concluded for 1533 cases. Among these cases, 950 were discarded (i.e. not fulfilling the operational case definition of microcephaly and/or CNS malformation associated with congenital infection), 583 were confirmed and 4107 remain under investigation (Table 2).
- Among the 5640 suspected cases of microcephaly and/or CNS malformation, 120 child deaths occurred after birth or during pregnancy (miscarriage or stillbirth); 30 of these were confirmed as having microcephaly and/or CNS malformation potentially linked to congenital Zika virus infection, 80 remain under investigation and 10 were discarded.

8 http://portalsaude.saude.gov.br/images/pdf/2016/fevereiro/23/coes-microcefalia-informe-epidemiologico14-se07-2016-fev2016-14.pdf

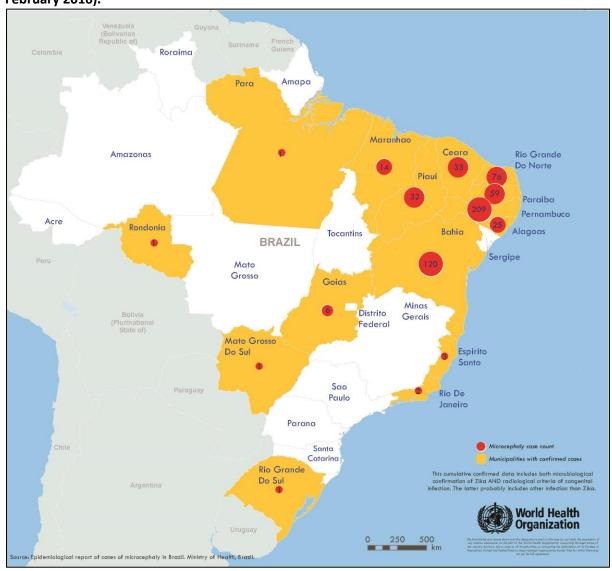
<sup>&</sup>lt;sup>6</sup> <a href="http://governor.hawaii.gov/newsroom/doh-news-release-hawaii-department-of-health-receives-confirmation-of-zika-infection-in-baby-born-with-microcephaly/">http://governor.hawaii.gov/newsroom/doh-news-release-hawaii-department-of-health-receives-confirmation-of-zika-infection-in-baby-born-with-microcephaly/</a>

<sup>&</sup>lt;sup>7</sup> http://www.nejm.org/doi/pdf/10.1056/NEJMoa1600651

http://portalsaude.saude.gov.br/index.php/cidadao/principal/agencia-saude/22295-casos-suspeitos-chegam-a-4-107-emtodo-o-pais

- An outbreak of Zika virus in French Polynesia was followed by an increase in the number of CNS malformations in children born between March 2014 and May 2015. 18 cases were reported including nine microcephaly cases compared to the national average of 0-2 cases per year.
- Zika virus is not proven to be a cause of the increased incidence of microcephaly in Brazil. However, given the temporal and geographical associations between Zika virus infections and microcephaly, and in the absence of a compelling alternative hypothesis, a causal role for Zika virus is a strong possibility that is under investigation.

Figure 4: Confirmed cases of microcephaly in states of Brazil (583 cases reported up to 20 February 2016).<sup>10</sup>



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http://portalsaude.saude.gov.br/index.php/cidadao/principal/agencia-saude/22295-casos-suspeitos-chegam-a-4-107-em-todo-o-pais

## Incidence of Guillain-Barré syndrome (GBS)

In the context of Zika virus circulation eight countries or territories have reported increased GBS incidence and/or laboratory confirmation of a Zika virus infection among GBS cases (Table 3, Fig. 3).

Table 3. Countries, territories or areas reporting GBS potentially related to Zika virus infection.

Increased incidence of GBS cases (without biological confirmation of the association with Zika)	Increased GBS incidence and biological confirmation of Zika infection in at least some of the cases	Reporting GBS with laboratory confirmed Zika virus infection (without increase of GBS incidence)
Brazil	French Polynesia	Martinique
El Salvador	Suriname	Puerto Rico
Colombia	Venezuela	

- In 2015, 42 GBS cases were reported, among which 26 (62%) had a history of symptoms consistent with Zika virus infection in the state of Bahia in Brazil. A total of 1708 cases of GBS were registered nationwide, representing a 19% increase from the previous year (1439 cases of GBS in 2014), though not all states reported an increase in incidence.
- Colombia reported an increase in the incidence of GBS as 201 GBS cases with history of suspected Zika virus infection were reported in the nine weeks to 14 February. Most of the cases are from Norte de Santander and Barranquilla areas where many of the Zika virus cases have been registered. To date, none of these cases have been laboratory confirmed for Zika virus infection, or other possible causes.
- El Salvador recorded 118 GBS cases from 1 December 2015 to 8 January 2016, including five deaths, while the annual average number of GBS cases is 169. To date, none of those reported GBS cases have been laboratory confirmed for Zika virus infection or other causes.
- On 29 January 2016, Suriname reported an increased incidence of GBS: 10 GBS cases reported in 2015 and three GBS cases were reported during the first three weeks of 2016, while Suriname registers on average approximately four cases GBS per year. For two of the GBS cases reported in 2015 a Zika virus infection was laboratory confirmed by RT-PCR.
- Venezuela has also reported an increased incidence of GBS. From 1 January to 31 January 2016, 252 cases of GBS suspected to be related to Zika virus were reported. The largest number of cases (66) was reported from six municipalities of Zulia state, mostly concentrated in the Maracaibo municipality. Zika virus was confirmed in a three of the GBS cases by RT-PCR.
- GBS cases with laboratory confirmed Zika virus infections were reported from Martinique (two cases) and Puerto Rico (one case).
- In French Polynesia, 42 GBS cases were identified during the 2013 2014 Zika outbreak,
   88% of those reported an illness compatible with Zika infection. Retrospective analysis

- (seroneutralisation test) demonstrated that all 42 cases were positive for dengue and Zika virus infection.
- As with microcephaly, Zika virus is not proven to be a cause of increased GBS incidence in Brazil, Colombia, El Salvador, Suriname or Venezuela. However, a causal role for Zika virus is a strong possibility. Confounding factors include the contemporary circulation of dengue and chikungunya in the Americas, which are transmitted by the same species of mosquito. Further investigations are needed to identify the potential role of other factors (including infections) known to be associated, or potentially associated, with GBS.

### **II. RESPONSE**

- The principal activities being undertaken jointly by WHO and international, regional and national partners in response to this public health emergency are laid out in Table 4.
- WHO and partners are working together to develop and maintain the Joint Operations Plan that combines activities against the six main areas of work; coordination, surveillance, care, vector control, risk communication and community engagement, and research at the global, regional and country level.
- WHO and partners are appealing for the sum of US\$ 56 million for an inter-agency, international response to the spread of Zika virus disease and subsequent spikes in cases of microcephaly and neurological disorders. The request represents the consolidated requirements of 23 partner organizations to address this emergency over the next six months. US\$ 25 million is required to fund the WHO and PAHO emergency response and US\$ 31 million to fund partners' activities. Approximately 45 donors attended a meeting to discuss the Strategic Response Framework. Donors are reviewing stated needs and requirements.
- On 18 February 2016 the World Bank Group announced that it had made US\$ 150 million immediately available to support countries in Latin America and the Caribbean affected by the Zika virus outbreak. This amount follows the WHO declaration of a Public Health Emergency of International Concern (PHEIC) on 1 February 2016 for the recent cluster of microcephaly cases and other neurological disorders reported in the Americas amid the growing Zika virus outbreak. The World Bank Group has engaged with governments across the region, including sending technical experts to affected countries. If additional financing is needed, the World Bank Group stands ready to increase its support. These initial estimates assume that the most significant health risks are for pregnant women.
- WHO has developed new advice and information on Zika case definitions; prevention of sexual transmission of Zika virus; vector control; blood safety; identification and case management of Guillain-Barrè syndrome; breast feeding in the context of Zika; and case definition, and assessment of infants with microcephaly in the context of Zika.<sup>11</sup>

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<sup>&</sup>lt;sup>11</sup> See resources listed at Annex 1

■ These materials are being transformed into many formats to support risk communication, community engagement and for the use of key stakeholders including health workers.

Table 4. Strategic Response Framework and Joint Operational Response Plan: response activities.

activities.	
Public health	<ul> <li>Activate networks of social science experts to advise on community</li> </ul>
risk	engagement.
communication	<ul> <li>Coordinate and collaborate with partners on risk communication messaging</li> </ul>
and community	and community engagement for Zika.
engagement	<ul> <li>Develop communication and knowledge packs and associated training on</li> </ul>
activities	Zika and all related and evolving issues for communication experts.
	<ul> <li>Engage communities to communicate risks associated with Zika virus</li> </ul>
	disease and promote vector control, personal protection measures, reduce
	anxiety, address stigma, and dispel rumours and cultural misperceptions.
	<ul> <li>Disseminate material on Zika and potentially associated complications for</li> </ul>
	key audience such as women of reproductive age, pregnant women, health
	workers, clinicians, and travel and transport sector stakeholders.
	<ul> <li>Conduct social science research to understand perceptions, attitudes,</li> </ul>
	expectations and behaviours regarding fertility decisions, contraception,
	abortion, pregnancy care, and care of infants with microcephaly and
	persons with GBS.
	Support countries to monitor impact of risk communications.
Vector control	Regularly update and disseminate guidelines/recommendations on
and personal	emergency <i>Aedes</i> mosquito control and surveillance.
protection	Support insecticide resistance monitoring activities.
against 	Support countries in vector surveillance and control, including provision of
mosquitoes	equipment, insecticides, personal protection equipment (PPE) and training.
Care for those	Assess and support existing capacity and needs for health system
affected and	strengthening, particularly around antenatal, birth and postnatal care,
advice for their carers	neurological and mental health services, and contraception and safe abortion.
Carers	<ul> <li>Map access barriers limiting women's capacity to protect themselves</li> </ul>
	against unintended pregnancy.
	<ul> <li>Develop guidance for: families affected by microcephaly, GBS or other</li> </ul>
	neurological conditions; women suspected or confirmed to have Zika virus
	infection, including women wanting to get pregnant, pregnant women, and
	women who are breastfeeding; health workers on Zika virus health care,
	blood transfusion services, tools for triage of suspected Zika, chikungunya
	and dengue cases; and for health services management following a Zika
	virus outbreak.
	<ul> <li>Provide technical support to countries on health service delivery</li> </ul>
	refinements and national level planning to support anticipated increases in
	service needs.
	<ul> <li>Procure and provide equipment and supplies for prioritized countries and</li> </ul>
	territories to prepare their healthcare facilities in provision of specialized
	care for complications of Zika virus.

## III. RESEARCH

- Public health research is critical for establishing the causal link between Zika virus infection in pregnant women and microcephaly in their babies and for understanding the pathogenesis of Zika virus infection. Technical assistance is being coordinated with various partner agencies globally and in affected countries to identify and answer critical questions (Table 5).
- A meeting on defining the public health research agenda is being organized by PAHO in Washington, D.C. from 1 to 2 March 2016 which will include a workshop with Global Outbreak Alert and Response Network (GOARN) technical partners
- A global consultation on research related to Zika virus infection will be held from 7 to 9
   March 2016 to assess the research landscapes and plan for additional research.

Table 5. Strategic Response Framework and Joint Operational Response Plan: research objectives and activities.

research	Investigate reported increase in incidence of microcephaly and neurological syndromes and their possible association with Zika virus infection.	•	Conduct research studies to assess link between Zika virus and microcephaly. Conduct research to assess potential sexual transmission and mother-to-child transmission. Research women's and health workers' perceptions of pregnancy risk and consequent decisions on contraceptive use, safe abortion and post abortion care in context of Zika virus.
development	Fast-track research and development of new products including diagnostics, vaccines and therapeutics.		Identify research gaps and prioritize needs for products. Support the conduct of research related to Zika virus diagnostics, therapeutics, vaccines and novel vector control approaches Convene research actors and stakeholders. Coordinate introduction of products after assessment and evaluation. Coordinate supportive research activities including regulatory support and data sharing mechanisms.

#### Annex 1: Additional information

#### **Zika Virus**

- Zika virus disease is caused by a virus transmitted by Aedes mosquitoes. Other transmission modes are still under investigation.
- People with Zika virus disease usually have a mild fever, skin rash (exanthema), and conjunctivitis. These symptoms normally last for 2-7 days.
- At present there is no specific treatment or vaccine currently available. The best form of prevention is protection against mosquito bites.
- Zika virus is known to circulate in Africa, the Americas, Asia, and the Pacific region. Zika virus had only been known to cause sporadic infections in humans until 2007, when an outbreak in Micronesia infected 31 people.

## Microcephaly

Microcephaly is an uncommon condition where a baby's head circumference is less than expected based on the average for their age and sex. The condition is usually a result of the failure of the brain to develop properly, and can be caused by genetic or environmental factors such as exposure to toxicins, radiation, or infection during development in the womb. Microcephaly can be present as an isolated condition or may be associated with other symptoms such as convulsions, developmental delays, or feeding difficulties.

## **Guillain-Barré syndrome**

• Guillain-Barré syndrome in its typical form is an acute illness of the nerves that produces a lower, bilateral, and symmetrical sensorimotor development deficit. In many cases there is a history of infection prior to the development of the Guillain-Barré syndrome. The annual incidence of GBS is estimated to be between 0.4 and 4.0 cases per 100,000 inhabitants per year. In North America and Europe GBS is more common in adults and increases steadily with age. Several studies indicate that men tend to be more affected than women.

## **Resources from WHO**

- Zika virus www.who.int/mediacentre/factsheets/zika
- Microcephaly www.who.int/emergencies/zika-virus/microcephaly/en/
- Guillain-Barré syndrome <u>www.who.int/mediacentre/factsheets/guillain-barre-</u> syndrome/en/
- Infants with microcephaly <u>www.who.int/csr/resources/publications/zika/assessment-infants/en/</u>
- Guillain-Barré syndrome www.who.int/csr/resources/publications/zika/guillain-barresyndrome/en/
- Breastfeeding <u>www.who.int/csr/resources/publications/zika/breastfeeding/en/</u>
- Sexual transmission <a href="http://who.int/csr/resources/publications/zika/sexual-transmission-prevention/en/">http://who.int/csr/resources/publications/zika/sexual-transmission-prevention/en/</a>
- Vector control <a href="http://www.who.int/emergencies/zika-virus/articles/mosquito-control/en/">http://www.who.int/emergencies/zika-virus/articles/mosquito-control/en/</a>
- Blood safety <a href="http://who.int/csr/resources/publications/zika/safe-blood/en/index.html">http://who.int/csr/resources/publications/zika/safe-blood/en/index.html</a>