



# The Japanese Encephalitis Virus Evolution and Epidemiology

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## تطور فيروس التهاب الدماغ الياباني وعلم الأوبئة

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### **ABSTRACT**

The virus that causes Japanese encephalitis (JEV) is the cause of Japanese encephalitis (JE), a potentially fatal brain infection contracted via mosquito bites. The Japanese encephalitis virus, which is endemic in 24 countries and poses a threat to nearly 3 billion people, is the primary Asia's viral encephalitis cause. An estimated one hundred thousand clinical cases and 25,000 fatalities occur worldwide each year. Travelers to endemic areas must be vaccinated, even though Arab nations are still outside the transmission zones. Prevention: Inactivated (IXIARO), live attenuated (IMOJEV, CD. JEVAX), and Indian (JENVAC) vaccine formulations are among the WHO-qualified vaccinations. Dosage, target populations, and contraindications vary for each. Although the effectiveness of current vaccines may be threatened by developing genotype V, sustained immunizations has reduced endemic countries by 73–100%. To rule out flavivirus cross-reactivity, primary diagnosis is made by IgM capture ELISA in serum/CSF (WHO-recommended), followed by confirmatory PRNT at reference laboratories. Because of temporary viraemia, RT-PCR is restricted in its ability to diagnose humans, although it is beneficial for surveillance. Socio-environmental Factors, historically recorded during the Korean War (402 US cases, 1950), social instability (war, displacement, and health system collapse) increases burden by disrupting immunization, weakening surveillance, and increasing vector exposure.

**Keywords:** Japanese encephalitis virus, vaccination, epidemiology, Culex, flavivirus

## INTRODUCTION

It is the viral encephalitis brought on by the Japanese encephalitis virus (JEV). most often in Asia [1]. Japanese encephalitis is occurred by a flavivirus which is spread via mosquitoes [2]. The JEV was identified in the brain of a JE patient in 1935; the patient subsequently succumbed to the disease. This isolate, commonly referred to as believed that the JEV prototype strain is believed to be the Nakayama strain [3]. Compared to adults, infants and kids till they turn fifteen are higher susceptible to JE and face a more risk of neurological complications [4]. The threat of JE is ever-present for over 2 billion people who live in endemic countries, and a rise in the likelihood that the disease may spread is increased by mosquito numbers to previously uninfected areas. A female *Culex tritaeniorhynchus* mosquito carrying the RNA virus known as the JEV will often bite a victim. Additionally, *Culex annulirostris*, *Culex vishnui*, *Culex pseudovishnui*, *Culex gelidus*, *Culex sitiens*, and *Culex fuscocephala* are mosquito species that have been linked to the spread of JEV. A number of mosquito species, including *Anopheles peditaeniatus*, *Anopheles hyrcanus*, and *Anopheles subpictus*, [5]. A family of birds called Ardeidae is the main source of JEV, which includes egrets and herons. Pigs can infect naive mosquitoes because they are very susceptible to JEV, which results in the virus multiplying to ideal levels and developing into an amplified host with a high circulating viral titer [6]. Pigs are known to normally release the virus orally during this period, which may facilitate JEV's horizontal dissemination [7]. Cattle and the reason horses are regarded as "dead-end" hosts is that they don't generate large virus titers, in contrast to pigs and birds (Figure 1)

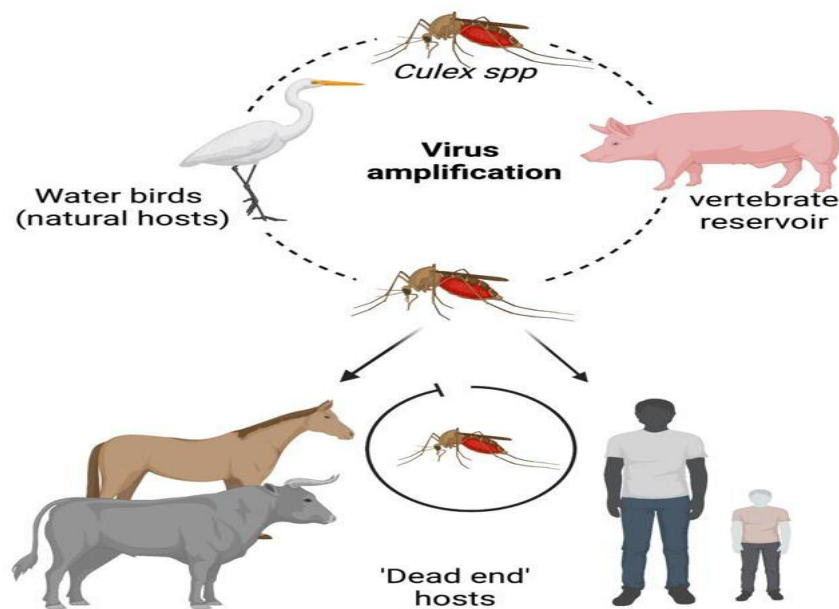


Fig.1 The model of infection with and amplification of the JEV.





however, have shown that the virus has persisted in spreading in this area. The first incidence of Japanese encephalitis (JE) was recorded in the Australian Northern Territory in 2021 [16], prompting worries about potential outbreaks in the area in the future. Furthermore, the 2016 discovery of an autochthonous case in Angola raises questions regarding the potential that JEV infection is spreading throughout Africa [17]. In most of Southeast Asia, surveillance for JE illness is still insufficient despite its high prevalence. Because many nations lack the diagnostic tools necessary to establish JEV infection, JE symptoms are frequently mislabeled as other viral illnesses. Because of this, evaluating JEV's situation in the Asia-Pacific area is challenging. Owing to the high viral seroprevalences in people, wild animals, and domestic animals [18]. In many nations without the diagnostic tools necessary to assess the JEV burden, health information is inadequate. JEV infection and transmission rates are underestimated in the Asia-Pacific area. To determine the full cost of JE, there has to be more JEV infection reporting and surveillance.

### **SYMPTOMS AND SIGNS**

Most cases of JE infections are often mild and manifest as subclinical fever or no symptoms at all. But the rate of case fatalities among individuals with severe clinical disease (encephalitis) might reach up to 30%. Approximately 30 to 50 percent of patients who survive a severe illness experience neurological or behavioral consequences, as paralysis, frequent seizures, and/or difficulty speaking [19]. In individuals who develop neurological disease, there are three phases in the course of JE. During the prodromal phase, which is the initial stage, symptoms like chills, muscle soreness, meningitis, vomiting, and diarrhea are common. Reduced consciousness, convulsions, and Parkinsonian syndrome are symptoms of the second stage or acute phase. This condition progresses to viral encephalitis, which can cause cogwheel rigidity, generalized hypertonia, tremors, and other anomalies, possibly leading to a coma. Sometimes the illness advances quickly and has the potential to be fatal. The patient either recovers or experiences prolonged neurological aftereffects during the third stage, sometimes known as the late phase [20]. The most damaged brain region is discovered to be the hippocampus, although there is also damage to the brain stem, the spinal cord's anterior horn cells are called substantia nigra. thalamus, the basal ganglia, the parenchyma, the cerebral cortex, and the midbrain. There have been several histological abnormalities observed, such as cerebral edema, perivascular cuffing, glial mesenchymal nodules, vascular leakage, crowded leptomeninges, development of microglial nodules, and necrotic lesions [21]. The World Health Organization recommends that the primary diagnostic method for Japanese encephalitis be serological detection utilising IgM antibody capture ELISA in serum or cerebrospinal fluid. To rule out cross-reactivity with other flaviviruses, confirmatory diagnosis necessitates extra specialist testing, such as plaque reduction neutralisation testing (PRNT) in reference laboratories (Table 1)



Table (1). Diagnostic Methods for Japanese Encephalitis Virus: Comparison of Available Laboratory Techniques in Terms of Accuracy and Accessibility

Diagnostic method	Advantage	Disadvantage	Source
IgM Antibody Capture ELISA	primary diagnostic tool that the CDC and WHO endorse. • After 3–8 days of symptoms, there is a high sensitivity in serum and CSF. • It can be utilised to increase accuracy in differential testing (JE vs. Dengue). • Economical in situations where resources are scarce	False positives may result from cross-reactivity with other flaviviruses, such as dengue and West Nile. • IgM may last for 30 to 90 days, possibly indicating a previous immunisation or illness. • Convalescent sample retesting is necessary if the results are negative during the first ten days. • Needs qualified staff and laboratory infrastructure.	Siddalingaiah, N., et al. (2025). Enhancing Japanese Encephalitis diagnosis with differential ELISA testing in India <i>Journal of Vector Borne Diseases</i> . Doi.org/10.4103/jvbd.jvbd_43_25.
Hemagglutinations Inhibition (HI) Assay	Easy to use, quick, and economical. • Suitable for screening large quantities of samples at high throughput.	In order to eliminate non-specific inhibitors, sera must be pretreated. • Utilizes goose erythrocytes, necessitating the production of new blood. • Compared to other viruses, JEV is less frequently standardized.	Zhao, J., et al. (2025)., Comparative Analysis of Hemagglutination -Inhibition and Plaque Reduction Neutralization Tests for Japanese Encephalitis Virus, <i>Antibody Detection. Viruses</i> , 17(1), 104. Doi.org/10.3390/v17010104.
Plaque Reduction Neutralization Test (PRNT)	Highest specificity among all JE tests. • Can distinguish between JEV and cross-reactive flavivirus antibodies. • Essential for confirmatory testing after positive ELISA. • Measures functional antibody response.	intricate and time-consuming (takes more than five days). • Needs BSL-3 facilities and live viruses. • Technically complex; needs skilled staff. • Unsuitable for high-throughput screening or routine diagnostics.	
Reverse Transcription-Polymerase Chain Reaction (RT-PCR)	high viral RNA sensitivity and specificity. • The most economical molecular surveillance technique. • Able to	Low/transient viraemia at the time symptoms occur makes it insensitive for human diagnosis. • Clinical human specimens frequently result in false negatives. •	Crispell, G., et al., (2022)., Method comparison for Japanese encephalitis virus detection in samples collected from the IndoPacific region., <i>Frontiers in Public Health</i> ., 10,1051754. Doi.org/10.3389/fpubh.2022.1051754



	distinguish between genotypes. • Beneficial for animal samples and mosquito vectors.	Needs laboratory infrastructure and skilled workers. • Contamination risk.	
Biosensor-based Detection (Electrochemical, Nanomaterial-based)	Results come quickly (in minutes to hours). • Possibility of field deployment and point-of-care. • With nanomaterial augmentation, it may be extremely sensitive. • It's compact and portable.	Not available for regular use on the market. • Needs further standardization and validation. • Limited information on clinical evaluations. • Technology is still developing.	Kumar, A., et al., (2022)., A review on Japanese Encephalitis virus emergence., pathogenesis and detection., International Journal of Biological Macromolecules, 217, 1045-1058.

### **PATHOGENESIS OF JEV**

The virus requires passage through several physical and physiological obstacles before it may begin to multiply. Once it has drawn blood from an infected host the virus releases its DNA into the cytoplasm after merging its viral envelope with the intestinal epithelial cells' plasma membrane upon entering the midgut. The virus then begins to replicate, releasing virions into the hemocoel. Once these virions pass via the trachea, they eventually arrive at the salivary glands. The virus releases virions into the saliva when it eventually penetrates the barrier protecting the salivary glands from infection and infects the acinar cells. Currently, the carrier mosquito is ready to inject the virus into the next victim. When a mosquito bites a healthy individual, the virus enters the skin and spreads to other parts of the body, where it might cause either a persistent infection that penetrates the nervous system or a latent infection that just affects mononuclear cells [22]. Pathogen-associated molecular patterns (PAMP) trigger genes by connecting to pathogen recognition receptors (PRR) via the JNK pathway. The virus can alter these interferons, changing the antiviral state of both nearby uninfected host cells and infected host cells. The skin's innate immune cells are the main targets of the virus [23]. Tissue macrophages and blood monocytes are where the virus multiplies. If humoral IgM is produced throughout the initial five days of the virus's incubation phase, the immune system might be able to eradicate the virus before it enters the central nervous system(CNS) [24].

In the event that this is not the case, the virus start to access the CNS by overcoming the blood-brain barrier, thereby initiating the clinical phase that poses the greatest risk (BBB). The cerebral blood barrier (BBB) can be compromised in a number of ways: (a) the virus can replicate directly through endothelial cells; (b) the virus can colonize in inflammatory



cells or blood lymphocytes and enter the CSF through a Trojan horse mechanism; (c) Chymase, which relaxes the tight connection between endothelial cells, can be released by metalloproteases (MMP 2/MMP 9) or mast cells or (d) by receptor-mediated endocytosis [25].

Neuronal bystander death or direct neuronal cell death may arise from any of these processes. The virus causes an excess of cytokines that promote inflammation, including IL-6, iNOS, and TNF- $\alpha$ , after it enters the CSF. This finally causes inflammation in particular brain regions (neuroencephalitis) by activating astrocytes, pericytes, and microglial cells [26].

Research indicates that protection against JEV is provided by IFN- $\alpha$  (anti-inflammatory cytokine), This prevents assembly, release, and replication at various phases. Although its effectiveness has been investigated in vitro, no comparable result was seen in humans [27]. JEV further uses GKN3 and PLVAP receptors to enter neuronal cells. It then causes the death of neuronal cells and a decrease in the levels of IL-10 and IL-4, two anti-inflammatory cytokines. Further neuronal death is brought caused by the synthesis of inflammatory mediators by microglial and astrocyte cells, which is triggered by the destruction of nerve cells [28]. By enabling blood lymphocyte entrance into the CSF, chemokines (IP-10/ RANTES/ IL-8/ and MCP-1) secreted by microglial cells and astrocytes also contribute to an increase in infection. Neuron progenitor stem cells (NPSCs), which regulate the immune response to CSF, are prevented from proliferating by viral infection. For healthy brain cells, Interferon-induced long-term reduction of protein synthesis can be lethal. Research has shown that people with greater CSF levels of anti-JEV IgM have a higher chance of surviving without experiencing significant adverse impact on the central nervous system [29].

Created at the surface of the endoplasmic reticulum through the interplay of host factors, non-structural virus proteins, and LC3-1, EDEM 1, and SEI1L are examples of proteins known as endoplasmic reticulum-associated protein degradation (ERAD) [30]. In order to create double-stranded replicative forms, single-stranded RNA (ssRNA) must multiply. These forms serve as templates for the creation of positive sense mRNA. Next, host and viral proteases produce and break down viral proteins. Once the host furin at the trans-Golgi cleaves it to M, the virion matures by the release of prM, which forms in the lumen of the ER (figure 2) [ 31].





sequence availability could be the cause of these relationships. It will need more research to see whether genotype-defining substitutions may shed any light on the molecular mechanisms behind this phenomenon [35].

Long-lasting dry spells may be linked to fewer mosquito breeding grounds and the species that serve as their reservoirs, which will lessen the spread of JEV. Lower river flows can result in less irrigation water available, and significantly fewer rice fields can lead to much lower transmission. There will always be a replacement for amplifying species, like pigs. Pig vaccination is an uncommon practice, and in situations where there is little to no transmission, there may be less motivation for immunizing humans. Both human and pig populations that are vulnerable may rise significantly as a result of this scenario. JEV can spread over great distances by aquatic birds and is greatly amplified by pigs that lack immunity following a severe rain event that causes flooding [36], leading to a cascade into the vulnerable human populace.

Wind can carry mosquitoes a considerable distance [37]. Despite the well-established role that cyclones and other weather phenomena contribute to the spread of bluetongue and infected *Culicoides* outbreaks, there is little proof that wind-borne mosquitoes can disseminate the Japanese encephalitis virus (JEV) to new locations. Global warming-induced increases in the frequency and severity of these occurrences could enhance JEV's dispersion or bring it into regions where outbreaks are not yet occurring. In the future, there's a significant likelihood that endemic and epidemic zones with obvious seasonal fluctuations may disperse throughout Asia [38].

According to the most recent data, the following table shows the incidence, fatalities, and geographic distribution of Japanese encephalitis worldwide (table 2)



Table (2) incidence, fatalities, and geographic distribution of Japanese encephalitis worldwide.

statistic	value	source
Annual Deaths	Approximately 25,000 deaths per year	Gavi (2025)., Japanese encephalitis. Gavi., the Vaccine Alliance. Retrieved from <a href="https://www.gavi.org/types-support/vaccine-support/japanese-encephalitis">https://www.gavi.org/types-support/vaccine-support/japanese-encephalitis</a>
Annual Cases (Estimated Primary)	Every year, about 100,000 clinical cases	WHO (2024)., Japanese encephalitis., World Health Organization. Fact sheet., Retrieved from <a href="https://www.who.int/news-room/fact-sheets/detail/japanese-encephalitis">https://www.who.int/news-room/fact-sheets/detail/japanese-encephalitis</a>
Case Fatality Rate (CFR)	Up to 30% of people with symptoms of acute encephalitis	
Endemic Countries	24 nations in the Western Pacific and South-East Asia regions	
The Population at Risk	More than 3 billion individuals	

There are no local epidemiological statistics available for Arab countries because, according to all sources, they are beyond the virus's conventional transmission zone.

### **Vaccination against JEV**

Several JEV-endemic nations are now using a number of JEV vaccines that have been developed. Across Southeast Asia, mouse brain or cell culture-derived inactivated vaccinations have shown astounding efficacy. Early research, carried out in Taiwan in 1965, showed an 80% effectiveness rate. Acceptance of the vaccine has been hindered due to its poor long-term immunity, expensive production costs, and reports of unpleasant effects, despite the fact that it has been found to be 80–100% efficacious in adults in the US, Thailand, Japan, India, and other countries. Vaccinations that are inactivated have gradually been replaced by live attenuated vaccinations. Children receive two or three doses of these vaccinations [39].

The immunogenic live-attenuated vaccine takes one or two doses throughout childhood and produces long-term immunity. 97.5% and 98% of recipients in China and Nepal, respectively, experienced seroconversion after two dosage regimens, and strong antibody levels persisted for up to five years following immunization [40].



Although it was once exclusively available in China, it is currently the advised vaccination for numerous Asian endemic nations. In addition, a recombinant, live attenuated form JEV vaccine was created; it is both safe and efficacious, with 97% of children showing seroprotection five years following the booster dose. The efficiency of immunizations has come under scrutiny due to the emergence of JEV GI. Though at lower degrees of neutralization, serum from GIII-inactivated vaccine users could cross-neutralize GI viruses. Comparable research shows little cross-protection against local GI isolates when using animal antisera from GIII vaccinations [41]. Consequently, more investigation is needed to assess the efficacy of GIII JEV vaccines (table 2).

Campaigns, recurring immunization schedules, or a combination of the two are examples of JE vaccination tactics. A successful immunization approach is a one-time campaign aimed at young individuals, with the vaccine then being incorporated into routine immunization programs. 94% of the target population in Nepal was covered by widespread immunization efforts conducted between 2006 and 2009, aimed at children aged 1 to 15 [42]. In order to immunize children between the ages of 5 and 15, a JEV vaccine program was started in Myanmar in 2017. 2018 saw the implementation of routine immunization [43]. With around 12.6 million youngsters immunized, the campaign achieved 92.5% coverage nationally. As of yet, no information about the impact of this campaign on JEV circulation in Myanmar has been released. According to an extensive study conducted on 14 endemic countries between 2007 and 2021, campaign and routine immunization programs reduced cases of juvenile epilepsy (JE), deaths, and cases with sequelae. This decreased the number of disability-adjusted life years (DALYs) by 6,622,923 and saved USD19 million in cases of adult male epilepsy [44]. Therefore, it is anticipated that the introduction of JE vaccination programs in endemic nations will be extremely advantageous.

JEV is becoming easier to control thanks to vaccination campaigns, but more surveillance is required to raise the accuracy of JEV burden estimates. One of the main issues in determining the local and worldwide burden of JE and improving the identification of disease-risk areas is underreporting. The WHO has surveillance guidelines accessible [45]. Nevertheless, the ability to diagnose JEV is restricted in many JE-endemic nations. A WHO JEV laboratory network has been created to address this problem; it is modeled after laboratory networks for measles and polio [46]. Through growth and training, the network hopes to increase capacity for surveillance and diagnosis. With the network, JE surveillance is currently conducted in 92% of endemic Asian nations, up from 75% in the previous year [47]. These reports are most likely the result of earlier reporting irregularities combined with improvements in surveillance techniques.



Table (3) Decline in Japanese Encephalitis cases and the role of vaccination

Country	Evidence of decline	Role of Vaccination	Source
Global Estimates	Annual global burden estimated at 100000 cases and 25,000 deaths	JE is a vaccine-preventable disease, widespread vaccination has significantly reduced incidence globally over the past two decades	WHO (2024)., Japanese encephalitis., World Health Organization. Fact sheet., Retrieved from <a href="https://www.who.int/news-room/fact-sheets/detail/japanese-encephalitis">https://www.who.int/news-room/fact-sheets/detail/japanese-encephalitis</a>
Vaccine-introducing counties prior to 2006 (5 countries + Chinese Taipei)	73–100% decrease in the incidence of JE in all age groups between 5 and 20 years after introduction	Significant reductions were attained by long-term national immunization campaigns, but it required about 20 years and significant funding.	Letson, G. W., et al., the JE Vaccine Global Impact Assessment Team (2024)., Impact of vaccination against Japanese encephalitis in endemic countries. PLoS Neglected Tropical Diseases, 18(9), e0012390. Doi.org/10.1371/journal.pntd.0012390
Six countries have been introducing vaccines since 2006.	Between 2015 and 2021, the incidence of JE among children under the age of fifteen decreased by 14–79%.	Impact was shown, but after the first introduction, coverage needs to be maintained by ongoing delivery system upgrades.	
Japan	Before the 1960s, there was a sharp drop from over 1,000 cases annually to less than 10 instances annually; however, new reports reveal that this number may have been underestimated.	The JE vaccine was first developed in 1954, and its widespread use in children starting in 1967 significantly reduced the burden of disease.	Yoshizawa, K., et al., (2025)., Emergence of Japanese encephalitis in a previously non-reported area: Three consecutive annual cases from a tertiary center in Narita, Chiba, Japan. Journal of Infection and Chemotherapy., 31(6), 102706. Doi.org/10.1016/j.jiac.2025.102706.
Guizhou Province, China	A notable decrease in the prevalence of JE was observed between 2005 and 2021; the average incidence was 0.83 per 100,000, and it peaked in 2020 during the COVID-19 pandemic.	Participation in expanded immunisation programmes and catch-up efforts had a notable effect on the incidence rate's decline.	Zhang, W.X. Zhao, S., Pan, C., Zhou, Y., Wang, C., Rui, L., Du, J., Wei, T. T., Liu, Y. Q., Liu, M., Lu, Q. B., & Cui, F. (2024). Mass immunisation to eradicate Japanese encephalitis: Real-world evidence from Guizhou Province in 2005–2021. Journal of Virus Eradication, 10(1), 100366. <a href="https://doi.org/10.1016/j.jve.2024.100366">https://doi.org/10.1016/j.jve.2024.100366</a>



Gavi supported countries (Laos, Nepal, Myanmar, Cambodia, Indonesia)	By the end of 2023, more than 7.8 million children had received vaccinations through regular programs, and more than 17.7 million through catch-up initiatives.	The Gavi programme started in 2015, and Bangladesh is expected to launch a subnational campaign in 2026.	Gavi (2025). Japanese encephalitis. Gavi, the Vaccine Alliance. Retrieved from <a href="https://www.gavi.org/types-support/vaccine-support/japanese-encephalitis">https://www.gavi.org/types-support/vaccine-support/japanese-encephalitis</a>
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Table (3) Types of Japanese Encephalitis Vaccines Available in Arab Countries: Advantages and Disadvantages

Vaccine type	Advantages	Disadvantages	Source
Inactivated Vero Cell Vaccine IXIARO® (Valneva, Austria) / JESPECT	Most extensively accessible abroad and in travel clinics. All children ≥2 months are approved (paediatric data available). • Safe for pregnant women and those with impaired immune systems (where risk exceeds risk). 95% seroprotection at 1 month after the first series and 92% at 5–6 months indicate high immunogenicity. • Shown to be interchangeable with booster doses from other vaccination classes.	requires a two-dose main series spaced 28 days apart to provide complete protection. • Series completion is required at least one week prior to departure. • Seniors have lower seroprotection (65% in those aged ≥64 years). Among the local reactions are redness, discomfort, and soreness (50 per cent in adults and children). Fatigue (11%), fever (8%), and muscle discomfort (31%) are examples of systemic responses.	Furuya-Kanamori, L., et al., (2021). Comparison of immunogenicity and safety of licensed Japanese encephalitis vaccines, A systematic review and network meta-analysis. Vaccine., 39(32), 4429-4436. Doi.org/10.1016/j.vaccine.2021.06.023.
Live chimeric vaccine (yellow fever 17D backbone) IMOJEV® (Sanofi Pasteur) / JE-CV / ChimeriVax™-JE	just one dosage. • Of all the JE vaccines, the most immunogenic: OR = 5.93 for seroconversion compared to previous mouse-brain vaccines. • Long-term protection: Following	Immunocompromised and pregnant women should not use this medication. • Has the backbone of the yellow fever virus (rare hypersensitivity potential). • Inadequate data on senior citizens. •	



	a single dose, models anticipate protection for up to 25 years. • Shown to be interchangeable with CD. JEVAX® enhancer	Typical response: children's fever and agitation	
Inactivated Vero Cell Vaccine (Indian Strain) JENVAC® (Bharat Biotech, India)	Unlike other inactivated vaccines that need two doses, a single-dose primary series is feasible. • At 1 year, the highest long-term immunogenicity was GMT 33.7, compared to 12.2 for live attenuated. • One-year seroprotection: 81.7% (compared to 47.9% for live attenuated). • Seroprotected at 2 years: 88.5%. • Showed that live attenuated SA 14-14-2 could be interchanged.	primarily accessible in South Asia and India at the moment. • International availability is lower than that of IXIARO. • In non-endemic traveller populations, there is less data.	Infectious Disease Advisor (2020). Long-Term Immune Response With Single-Dose, Inactivated Japanese Encephalitis Vaccine. Based on Vadrevu, K. M., et al. (2020). Persistence of immune responses, with an inactivated Japanese encephalitis, single-dose vaccine., JENVAC and interchangeability with a live-attenuated vaccine. Journal of Infectious Diseases., 222(9), 1478-1487. Doi.org/10.1093/infdis/jiz672.
Live Attenuated Vaccine (SA14-14-2 strain) CD.JEVAX® (Chengdu Institute, China)	Mass campaigns are made simpler with a single-dosage primary series. • WHO prequalified, extensively utilised in Asia. • Long-term protection: five years after the booster, 97.5% seroprotection was observed. • GMT 2092.4 at 30 days after booster; highly immunogenic. • Proven compatibility with IMOJEV® for booster dosages	Not recommended for immunocompromised people (live virus) or pregnant women. • Within seven days, 20% of recipients experience a common reaction: fever. • Lower GMT (12.2 vs. 33.7) in comparison to inactivated JENVAC at 1 year. • Only 47.9% of patients had 1-year seroprotection (compared to 81.7% for JENVAC). • In certain schedules, a booster is advised after a year	Porntharukchareon, T., et al. (2024), Long-term immunogenicity of the SA14-14-2 Japanese encephalitis (JE), vaccine (CD.JEVAX) booster following chimeric JE (IMOJEV) vaccine priming in Thai children. Human Vaccines and Immunotherapeutics, 20(1), 2407663.



Inactivated Vaccine (Indian Pediatric) JEEV® (Biological E. Limited, India)	WHO was prequalified. Offered in formulations for adults (6 µg) and children (3 µg). Frequently utilized in Indian vaccination campaigns	<ul style="list-style-type: none"> <li>Limited availability outside of South Asia and India.</li> <li>Needs two doses</li> </ul>	The Tribune (2025). JE vaccine interchangeability, ICMR to study mixed regimen impact. Retrieved from <a href="https://www.tribuneindia.com/news/je-vaccine-interchangeability-icmr-to-study-mixed-regimen-impact/">https://www.tribuneindia.com/news/je-vaccine-interchangeability-icmr-to-study-mixed-regimen-impact/</a>
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## CONCLUSIONS

In many Asian countries, despite the availability of safe and effective vaccinations, vaccination rates are still low in the majority of places where JEV is most needed, making it a persistent public health risk. As mosquito species that have adapted to human habitats spread regionally and climate change may have an effect, there are worries that the Japanese encephalitis virus may continue to spread to new places because there are insufficient vector control measures in place.

## Conflict of interests.

There are non-conflicts of interest.

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## الخلاصة

فيروس التهاب الدماغ الياباني (JEV) هو سبب التهاب الدماغ الياباني (JE)، وهو عدوى دماغية قد تكون قاتلة تنتقل عبر لدغات البعوض. يُعدّ فيروس التهاب الدماغ الياباني (JEV)، المتوطن في 24 دولة ويُشكّل خطرًا على ما يقرب من 3 مليارات شخص، السبب الرئيسي لالتهاب الدماغ الفيروسي في آسيا. ويُقدّر عدد الحالات السريرية بنحو 100,000 حالة، وعدد الوفيات بنحو 25,000 حالة سنويًا في جميع أنحاء العالم. يجب على المسافرين إلى المناطق الموبوءة التطعيم، حتى وإن كانت الدول العربية لا تزال خارج مناطق انتقال العدوى. الوقاية: تشمل اللقاحات المعتمدة من منظمة الصحة العالمية لقاحات معطلة (IXIARO)، ولقاحات حية مُضعفة (IMOJEV، JEVAX، CD)، ولقاحات هندية (JENVAC). تختلف الجرعة والفئات المستهدفة وموانع الاستخدام لكل لقاح. على الرغم من أن فعالية اللقاحات الحالية قد تتأثر بظهور النمط الجيني الخامس، إلا أن حملات التطعيم المستمرة قد خفضت عدد الدول الموبوءة بنسبة تتراوح بين 73% و 100%. لاستبعاد التفاعل المتبادل مع فيروسات الفلافيفيروس، يُجرى التشخيص الأولي باستخدام اختبار ELISA لالتقاط الأجسام المضادة Igm في المصل/السائل النخاعي (موصى به من منظمة الصحة العالمية)، يليه اختبار PRNT التأكيدي في المختبرات المرجعية. ونظرًا لوجود الفيروس في الدم بشكل مؤقت، فإن قدرة اختبار RT-PCR على تشخيص البشر محدودة، على الرغم من فائدته في المراقبة. وتزيد العوامل الاجتماعية والبيئية، التي سُجلت تاريخيًا خلال الحرب الكورية (402 حالة في الولايات المتحدة، 1950)، من العبء بسبب عدم الاستقرار الاجتماعي (الحرب، والنزوح، وانهيار النظام الصحي) من خلال تعطيل التحصين، وإضعاف المراقبة، وزيادة التعرض للنواقل.

**الكلمات المفتاحية:** فيروس التهاب الدماغ الياباني، التطعيم، علم الأوبئة، الكيولكس، الفيروس المصفر .