



Review Article

Transforming Dengue Control: Scientific Challenges, Operational Gaps, and Elimination Prospects

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DOI: <https://doi.org/10.24321/0019.5138.2026109>

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How to cite this article:

Narayanasamy S, Sivagnaname Y, Baskar V, Velan L. Transforming Dengue Control: Scientific Challenges, Operational Gaps, and Elimination Prospects. *J Commun Dis.* 2025;57(4):153-158.

Date of Submission: 2025-07-31

Date of Acceptance: 2025-12-15

ABSTRACT

Dengue has evolved into the most widespread arboviral disease, now threatening nearly half the world's population. Despite the availability of diverse control tools—from insecticides and Wolbachia-based methods to improved surveillance and vaccines—the global burden of dengue continues to rise. This review dissects the scientific and operational roadblocks undermining control efforts, including silent transmission by asymptomatic carriers, serotype co-circulation, antibody-dependent enhancement (ADE), and growing insecticide resistance. These challenges are compounded by rapid urbanisation, climate change, fragmented surveillance, and weak intersectoral coordination. The authors advocate for a paradigm shift: integrating real-time entomological surveillance, climate-informed predictive models, and next-generation diagnostics within a strengthened Integrated Vector-borne Disease Management (IVDM) framework. Emphasizing community engagement, cross-sectoral governance, and transdisciplinary research, this paper outlines a forward-looking strategy that aligns innovation with implementation. Transforming dengue control from a reactive response into a proactive public health mission is critical to achieving the long-standing goal of dengue elimination.

Keywords: Dengue, asymptomatic conundrum, serotype co-circulation, climate change, Urbanisation, ecological expansion, fragmented surveillance, delayed response.

Introduction

Dengue, a mosquito-borne viral infection caused by four antigenically distinct serotypes (DENV-1 to DENV-4), remains a formidable and escalating global health challenge. With an estimated 390 million infections annually—of which nearly 96 million presents with clinical symptoms—dengue has significantly outpaced existing public health interventions.

¹ Once endemic to select tropical regions, dengue now threatens nearly half of the world's population and has established transmission in over 125 countries, making it the most widespread arboviral disease. ^{2,3}The World Health Organization (WHO) classifies dengue as a high-priority disease with epidemic potential. ⁴This designation, however, contrasts sharply with the persistent rise in disease



burden despite the availability of key control tools, including insecticide-based interventions⁵, Wolbachia-infected mosquito releases⁶, improved surveillance systems⁷, and even a licensed vaccine⁸. This paradox highlights a disconnect between scientific advancements and field-level impact.

The reasons for this failure are multifactorial—fragmented surveillance, reactive interventions, inadequate community engagement, weak infrastructure, and insufficient political commitment.⁹ These systemic weaknesses are further exacerbated by climate change¹⁰, rapid urbanisation, insecticide resistance¹¹, and the silent transmission driven by asymptomatic carriers.¹²

As the global health community aspires toward dengue elimination, it is imperative to critically examine the unresolved scientific and operational challenges that keep dengue control at a crossroads—where the promise of elimination remains both urgent and elusive.

The Asymptomatic Conundrum

A major impediment to dengue control is the high prevalence of asymptomatic and subclinical infections, estimated to constitute 50–80% of all DENV cases.^{12,13} Although clinically silent, many of these individuals remain viraemic for several days, enabling *Aedes* mosquitoes to acquire and transmit the virus unknowingly. This cryptic reservoir fuels sustained transmission, particularly in hyperendemic settings where multiple serotypes co-circulate. Traditional surveillance systems—largely dependent on passive reporting of febrile illnesses—fail to capture this hidden burden, leading to underestimation of transmission intensity and misinformed vector control responses. Population-wide serological or molecular screening, while ideal, remains logistically daunting and cost-prohibitive in resource-limited settings. Additionally, the lack of rapid, highly sensitive, and field-deployable diagnostic tools limits the feasibility of early detection and containment strategies targeting asymptomatic carriers. Without integrating this silent fraction into surveillance frameworks, dengue control efforts will continue to fall short of breaking the chain of transmission and achieving elimination.

Serotype Co-Circulation and Antibody-Dependent Enhancement

The concurrent circulation of all four-dengue virus (DENV) serotypes create a complex immunological landscape that heightens the risk of severe disease and challenges public health strategies. Although a primary DENV infection provides durable immunity to that specific serotype, the transient cross-protective immunity to heterologous serotypes wanes within months, rendering individuals vulnerable to secondary infections. Antibody-dependent enhancement (ADE) is central to this phenomenon,

whereby sub-neutralising antibodies from a prior infection bind but do not neutralise a different serotype, instead promoting viral uptake via Fc_Y receptors on monocytes and macrophages. This leads to higher viraemia, exaggerated immune responses, and severe manifestations such as dengue haemorrhagic fever and shock syndrome.¹⁴ ADE also complicates vaccine development, as demonstrated by the Dengvaxia rollout in the Philippines. The vaccine's poor safety profile in seronegative recipients led to severe outcomes and sparked intense public backlash, regulatory restrictions, and broader hesitancy toward dengue vaccination efforts.⁸ Insecticide Resistance: A Vector Control Crisis

Aedes aegypti's escalating resistance to pyrethroids and other commonly used insecticides poses a significant barrier to effective dengue control. This resistance stems largely from the indiscriminate and prolonged use of insecticides, particularly in fogging and ultra-low volume spraying campaigns that are often deployed without adequate entomological surveillance or efficacy assessments.¹⁵ Such reactive and non-targeted applications frequently fail to suppress mosquito densities below epidemic transmission thresholds. While larvicide is conceptually more targeted and efficient, its operational success is constrained by the widespread presence of cryptic breeding habitats and the inconsistent involvement of communities in source reduction. Compounding the issue is the limited pipeline of novel insecticides with new modes of action, undermining the sustainability of chemical control interventions.^{5,15} In this context, insecticide resistance is not just a technical challenge but a critical vulnerability in current vector control programmes that urgently demands innovation, strategic rotation, and integration with non-chemical approaches.

Urbanisation and Habitat Explosion

The rapid pace of urbanisation in low- and middle-income countries has created ideal conditions for the proliferation of *Aedes aegypti* mosquitoes, the primary dengue vector. Urban sprawl, especially in informal settlements and peri-urban slums, leads to dense human populations living amid poorly managed environments. Discarded containers, rooftop water tanks, and flowerpots become abundant larval habitats. Compounding this, inadequate municipal infrastructure, unreliable water supply systems, poor waste disposal, and lack of regulatory enforcement undermine source reduction efforts. Community participation in vector control remains minimal, further diminishing the effectiveness of interventions aimed at eliminating breeding sites and controlling dengue transmission.¹⁶ Climate Change and Ecological Expansion

Climate change is fundamentally reshaping the ecological landscape for dengue transmission, enabling the expansion of *Aedes* mosquitoes into previously non-endemic,

temperate regions.^{10,17} Rising ambient temperatures accelerate mosquito development, prolong adult survivability, and enhance virus replication and transmission efficiency. Additionally, unpredictable rainfall patterns and urban flooding generate abundant and persistent breeding habitats, often in densely populated areas. These dynamic environmental changes render traditional static vector control strategies insufficient. Despite the growing recognition of these risks, the integration of real-time entomological, virological, and meteorological data into predictive models remains limited, curtailing the operational utility of climate-informed early warning systems.¹⁸

Fragmented Surveillance and Delayed Response
 Surveillance systems in many dengue-endemic countries remain passive, delayed, and poorly integrated. Entomological, clinical, and laboratory data are often collected in silos and rarely synthesised, resulting in fragmented responses that lack precision and timeliness. Fogging operations and other vector control measures are typically reactive rather than anticipatory, commencing only after outbreaks are already in progress.^{7,9} Modern tools such as GIS mapping, real-time case-based reporting, and syndromic surveillance remain underutilised due to persistent capacity constraints and the lack of sustained financial and institutional investment. Additionally, cross-border data sharing is negligible, despite the growing role of human mobility and transnational transmission in dengue epidemiology.¹⁹ A major limitation in entomological surveillance is the absence of efficient adult mosquito traps^{20,21}, which are essential for accurately estimating vector density thresholds and calculating the minimum infection rate (MIR)—critical metrics for early outbreak detection and evidence-based intervention planning.

Operational and Research Gaps in Novel Strategies
 Although novel vector control tools like Wolbachia-infected mosquitoes and genetically modified vectors offer considerable promise, their real-world implementation faces several operational and research barriers.^{6,23} Scaling up these interventions requires significant logistical coordination, sustained funding, and regulatory approval frameworks that are often underdeveloped in endemic regions. Concerns about ecological impacts and unintended consequences, such as species replacement or altered transmission dynamics, persist. Furthermore, public perception and community acceptance remain fragile, especially in areas with limited scientific literacy or distrust in authorities. On the therapeutic front, antiviral drug development remains stagnant due to the challenges posed by antibody-dependent enhancement (ADE) and the need for pan-serotype efficacy. These scientific barriers are compounded by weak market incentives and limited pharmaceutical investment, given dengue's status as a neglected tropical disease.²⁰ Addressing these gaps demands transdisciplinary re-

search, long-term financial commitment, and inclusive public engagement to translate innovation into impactful and sustainable public health outcomes.

The Socio-Political Dimension

Dengue control is not merely a biomedical challenge but a socio-political issue rooted in fragmented governance, short political cycles, and siloed bureaucracies. Ministries of health, urban development, and sanitation frequently operate without synergy, resulting in disjointed vector control and reactive outbreak responses.⁹ Political short-termism often prioritises visible but ineffective measures like fogging over sustained surveillance and prevention. Community participation—a cornerstone of integrated vector management—is weakened by misinformation, intervention fatigue, and the perception that dengue is non-fatal.⁹ Despite this, the socioeconomic cost is immense, driven by healthcare expenses, loss of income, and strain on public health systems.⁹ The controversy surrounding Dengvaxia—stemming from inadequate risk communication and early deployment without full safety clarity—has significantly damaged vaccine confidence, complicating future immunization campaigns.¹⁰ Overcoming these socio-political barriers requires strong leadership, intersectoral governance, transparent risk communication, and long-term investments that prioritise

The way forward

Dengue control must evolve from reactive, mosquito-centric interventions to forward-looking, evidence-based strategies rooted in Integrated Vector Management (IVM).^{24,25} Integrated Vector Management (IVM) can be evolved into Integrated Vector-Borne Disease Management (IVBDM), emphasising the parallel control of both vectors and diseases. This approach offers a sustainable framework by harmonising diverse strategies for vector-borne disease control through intersectoral collaboration, policy integration, and community empowerment. Crucially, vector control should be embedded within a broader public health system that prioritises early-warning surveillance using real-time entomological and epidemiological data, remote sensing, and climate-informed predictive modelling.²⁶⁻²⁸ Investment in innovative, affordable point-of-care diagnostics and pathogen detection in vectors can bridge the gap between surveillance and response.²⁹ Furthermore, strategies must be hyper-localised-tailored to reflect spatial heterogeneity in transmission dynamics, socio-economic vulnerabilities, urban infrastructure, and cultural behaviors.^{30,31} Transdisciplinary research linking virology, vector biology, human behaviour, and systems thinking will be pivotal to designing adaptive interventions.^{32,33} By shifting from short-term control to long-term prevention and resilience, dengue management can transform from a fragmented response into an integrated, context-aware public health paradigm.^{34,35}

Summary

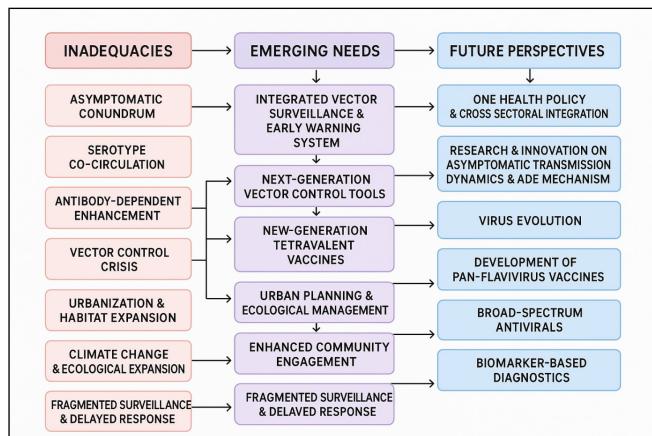


Figure 1. Summary

Conclusion

The transformation of dengue control stands at a critical crossroads, shaped by persistent scientific challenges, entrenched operational gaps, and a complex ecological and socio-political landscape. The high burden of asymptomatic infections, co-circulating serotypes, and the immunological paradox of antibody-dependent enhancement (ADE) continue to obscure disease dynamics and thwart conventional control strategies. Meanwhile, rapid urbanisation, expanding mosquito habitats, and climate-driven ecological shifts exacerbate vector proliferation and virus transmission. These are further compounded by fragmented surveillance systems, delayed outbreak responses, and gaps in both implementation and research of novel interventions.

To overcome these multifaceted barriers, a paradigm shift is essential—one that transitions from reactive containment to anticipatory, integrated, and adaptive control. Future success will depend on the establishment of robust, integrated vector surveillance coupled with real-time early warning systems, deployment of next-generation vector control tools, and investment in tetravalent vaccines and pan-flavivirus candidates. Urban planning must align with ecological management, supported by sustained community engagement and cross-sectoral governance under a One Health framework. Additionally, deeper research into asymptomatic transmission, ADE mechanisms, viral evolution, and host-pathogen interactions must guide the development of broad-spectrum antivirals and biomarker-based diagnostics. Only through such a comprehensive, scientifically grounded, and systemically integrated approach can the vision of dengue elimination shift from a distant aspiration to a tangible public health milestone.

Conflict of Interest: None

Source of Funding: None

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