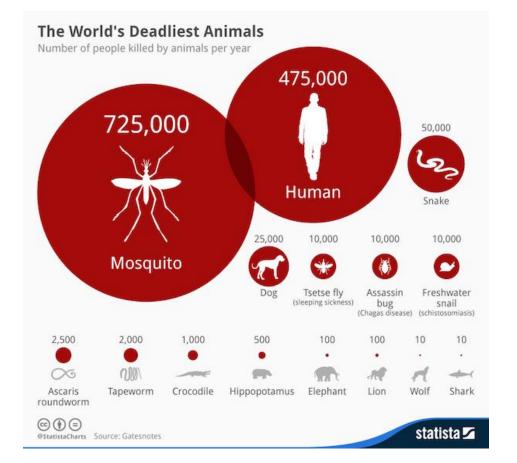


IVAN DARIO VELEZ MD, MSc, PhD Gerente WMP Colombia Director PECET, U. de Antioquia









Populations at risk from dengue, zika, chikungunya and yellow fever



Global Burden Of Dengue And Zika



Costo del dengue en Colombia





A multi-country study of the economic burden of dengue fever: Vietnam, Thailand, and Colombia

Jung-Seok Lee^{1,2}*, Vittal Mogasale²*, Jacqueline K. Lim²*, Mabel Carabali²*, Kang-Sung Lee²*, Chukiat Sirivichayakul³*, Duc Anh Dang⁴*, Diana Cristina Palencia-Florez⁵*, Thi Hien Anh Nguyen⁴*, Arthorn Riewpaiboon⁶*, Pornthep Chanthavanich³*, Luis Villar⁵*, Brian A. Maskery⁵*, Andrew Farlow¹*

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These authors contributed equally to this work.
 islee @ivi.int



Citation: Lee J-S, Mogasale V, Lim JK, Carabali M, Lee K-S, Sirivichayakul C, et al. (2017) A multicountry study of the economic burden of dengue tever: Vetram, Thailand, and Colombia. PLoS Negl Trop Dis 11(10): e0008037. https://doi.org/ 10.1371/journal.pntd.0008037

Editor: Benjamin Althouse, Institute for Disease Modeling, UNITED STATES

Received: April 18, 2017

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Data Availability Statement: All relevant data are within the paper and its Supporting Information

Abstract

Dackgrou

Dengue fever is a major public health concern in many parts of the tropics and subtropics. The first dengue vaccine has already been licensed in six countries. Given the growing interests in the effective use of the vaccine, it is critical to understand the economic burden of dengue fever to guide decision-makers in settling health policy priorities.

Methods/Principal findings

A standardized cost-of-lines study was conducted in three dengue endemic countries: Vietnam, Thailand, and Colombia. In order to capture all costs during the entire perid of illness, patients were tested with rapid diagnostic tests on the first day of their clinical visits, and multiple interviews were scheduled until the patients recovered from the current illness. Various cost items were collected such as direct medical and non-medical costs, indirect costs, and non-out-of-pocket costs. In addition, socio-economic factors affecting diseases severity were also identified by adopting a logit model. We found that total cost per episode ranges from \$141 to \$385 for inpatient and from \$410 to \$158 outpatient, with Colombia having the highest and Thailand having the lowest. The percentage of the private economic burAm. J. Trop. Med. Hyg., 94(5), 2016, pp. 1065–1072 doi:10.4269/ajtmh.15-0280 Copyright © 2016 by The American Society of Tropical Medicine and Hygiene

The Burden of Dengue and the Financial Cost to Colombia, 2010–2012

Raúl Castro Rodríguez,* Gabriel Carrasquilla, Alexandra Porras, Katia Galera-Gelvez, Juan Guillermo Lopez Yescas, and Jorge A. Rueda-Gallardo

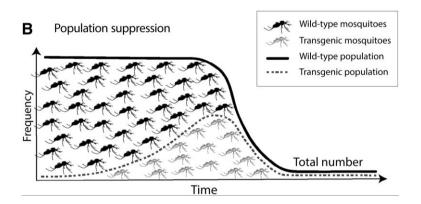
Department of Economics, Universidad de los Andes, Bogotá, Colombia; Centro de Estudios e Investigación en Salud (CEIS), Fundación Santa Fe de Bogotá, Bogotá, Colombia; Fundación Santa Fe de Bogotá, Universidad El Bosque, Bogotá, Colombia; Sanofi Pasteur Latin America, Mexico City, Mexico

Abstract. Data on the burden of dengue and its economic costs can help guide health policy decisions. However, little reliable information is available for Colombia. We therefore calculated the burden of the disease, expressed in disability-adjusted life years (DALYs), for two scenarios: endemic years (average number of cases in non-epidemic years 2011 and 2012) and an epidemic year (2010, when the highest number of dengue cases was reported in the study period). We also estimated the total economic cost of the disease (U.S. dollars at the average exchange rate for 2012), including indirect costs to households derived from expenses such as preventing entry of mosquitos into the home and costs to government arising from direct, indirect, and prevention and monitoring activities, as well as the direct medical and non-medical costs. In the epidemic year 2010, 1,198.73 DALYs were lost per million inhabitants versus 83.88 in endemic years. The total financial cost of the disease in Colombia from a societal perspective was US\$167.8 million for 2010, US\$129.9 million for 2011, and US\$131.7 million for 2012. The cost of mosquito prevention borne by households was a major cost driver (accounting for 46% of the overall cost in 2010, 62% in 2011, and 64% in 2012).





ESTRATEGIAS DE CONTROL : No vacunas, No medicamentos...













News MARZO 2015

Carcinogenicity of tetrachlorvinphos, parathion, malathion, diazinon, and glyphosate



http://dx.doi.org/10.1016/ For more on the IAPC Monographs see http:// monographs.iarc.fr June 2-9, 2015, Volume 113: insecticides and some Red meat and processed meat Monograph Working Group

genicity of the organophosphate including in pet flea collars. pesticides tetrachlorvinphos, parathion, March 20, 2015 Monographs.

Upcoming meetings studies was scarce and considered (combined),3 malignant pancreatic no increased risk of non-Hodokin chlorphenoxy herbicides malignant) in mice, renal tubule gland adenocarcinoma (after sub- cohort (AHS). Occupational use was Oct 6-13, 2015, Volume 114: tumours (beniqn or malignant) in cutaneous injection in females). 4 associated with an increased risk Members in male rats. Tetrachlorvinphos is distributed. Parathion metabolism to case-control study⁸ and in the AHS, A Blair (USA)—Meeting Chair, a reactive oxon with affinity for the bioactive metabolite, paraoxon, L Fritschi (Australia): LMcLaughlin: C.M. Sergi (Canada): GMCalaf (Chilk): FLe Curleux tetrachlorvinphos is systemically bacterial mutagenesis tests were for other pesticides. In mice, (Finland): (Baldi (France); distributed, metabolised, and negative, parathion induced DNA and malathion increased hepatocellular FForastiere (Italy); H Kromhout eliminated in urine. Although bacterial chromosomal damage in human cells adenoma or carcinoma (combined).10 (Netherlands): A 't Mannetie (New Zealand), T Rodriguez mutagenesis tests were negative, in vitro. Parathion markedly increased In rats, it increased thyroid carcinoma (unable to attend) (Nicaraqua): tetrachloryinphos induced genotoxicity rat mammary gland terminal end in males, hepatocellular adenoma or PErcenty funable to attend, GD in compactory (chromosomal damage hud density 4 Parathion use has been

In March, 2015, 17 experts from cell proliferation (hyperplasia in 11 countries met at the International rodents). Tetrachloryinphos is banned diazinon were dassified as "probably Agency for Research on Cancer (IARC; in the European Union, In the USA, carcinogenic to humans" (Group 2A). Lyon, France) to assess the carcino- it continues to be used on animals, Malathion is used in agriculture, public

For parathion, associations with It continues to be produced in (table). These assessments will be observed in occupational studies, "possibly carcinogenic to humans" in females. In rats, parathion induced Hodgkin lymphoma in the USA,

malathion, diazinon, and glyphosate cancers in several tissues were substantial volumes throughout the world. There is limited evidence in Published online published as volume 112 of the IARC but the evidence in humans remains humans for the carcinogenicity of sparse. In mice, parathion increased malathion. Case-control analyses The insecticides tetrachloryinghos bronchioloalyeolar adenoma and/or of occupational exposures reported and parathion were classified as carcinoma in males, and lymphoma positive associations with non-(Group 2B). The evidence from human adrenal cortical adenoma or carcinoma Canada,6 and Sweden,7 although inadequate, Tetrachlorvinphos induced tumours, and thyroid follicular cell lymphoma was observed in the hepatocellular tumours (benign or adenoma in males, and mammary large Agricultural Health Study male mice,2 and spleen haemangioma Parathion is rapidly absorbed and of prostate cancer in a Canadian which reported a significant trend for esterases. In experimental animals, is similar across species. Although aggressive cancers after adjustment carcinoma (combined) in females, and mammary gland adenocarcinoma after subcutaneous injection in females.4 Malathion is rapidly absorbed and distributed. Metabolism to the bioactive metabolite, malaoxon, is similar across species. Malaoxon strongly inhibits esterases; atropine reduced carcinogenesis-related effects in one study.4 Malathion induced DNA and chromosomal damage in humans, corroborated by studies in animals and in vitro. Bacterial mutagenesis tests were negative. Compelling evidence supported disruption of hormone pathways. Hormonal effects probably mediate rodent thyroid and mammary

The insecticides malathion and

health, and residential insect control.

gland proliferation. Diazinon has been applied in agriculture and for control of home and garden insects. There was limited evidence for diazinon carcinogenicity

Ja	An Some assays (Chromosomal damage bud density: Parathion use has beel shinke; CW Jameson, MT Martin, MT Kross; Rusyn; LZeise (USA) in rats and in vitro) and increased severely restricted since the 1980s.										
		Activity (current status)	Evidence in humans (cancer sites)	Evidence in animals	Mechanistic evidence	Classification*					
	Tetrachlorvinphos	Insecticide (restricted in the EU and for most uses in the USA)	Inadequate	Sufficient		2B					
	Parathion	Insecticide (restricted in the USA and EU)	Inadequate	Sufficient		2B					
	Malathion	Insecticide (currently used; high production volume chemical)	Limited (non- Hodgkin lymphoma, prostate)	Sufficient	Genotoxicity, oxidative stress, inflammation, receptor-mediated effects, and cell proliferation or death	2A†					
	Diazinon	Insecticide (restricted in the USA and EU)	Limited (non- Hodgkin lymphoma, leukaemia, lung)	Limited	Genotoxicity and oxidative stress	2A†					
	Glyphosate	Herbicide (currently used; highest global production volume herbicide)	Limited (non- Hodgkin lymphoma)	Sufficient	Genotoxicity and oxidative stress	2A†					

EU-European Union. *See the International Agency for Research on Cancer (IARC) preamble for explanation of classification system (amended January, 2006). †The 2A classification of diazinon was based on limited evidence of carcinogenicity in humans and experimental animals, and strong mechanistic evidence; for malathion and glyphosate, the mechanistic evidence provided independent support of the 2A classification based on evidence of carcinogenicity in humans and experimental animals.

Table: IARC classification of some organophosphate pesticides

	Activity (current status)	Evidence in humans (cancer sites)	Evidence in animals	Mechanistic evidence	Classification*
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EU=European Union. *See the International Agency for Research on Cancer (IARC) preamble for explanation of classification system (amended January, 2006). †The 2A classification of diazinon was based on limited evidence of carcinogenicity in humans and experimental animals, and strong mechanistic evidence; for malathion and glyphosate, the mechanistic evidence provided independent support of the 2A classification based on evidence of carcinogenicity in humans and experimental animals.

Table: IARC classification of some organophosphate pesticides



www.thelancet.com/oncology Vol 16 May 2015



Es una iniciativa sin fines de lucro que trabaja para **proteger a la comunidad mundial** de enfermedades transmitidas por mosquitos como el dengue, el Zika, el chikungunya y la fiebre amarilla.









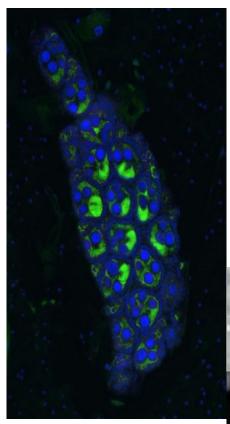
Fundación Bill y Melinda Gates, a través de la convocatoria Grandes Desafíos para la Salud Global, quien entrega la mayor parte de los dineros para su financiación

La Agencia de Cooperación Internacional de Estados Unidos **USAID**





¿Qué es Wolbachia?



- Bacteria exclusiva de insectos
- Intracelular obligatoria (simbionte)
- Incapaz de sobrevivir por fuera del insecto
- Transmisión materna
- No hay transferencia horizontal de *Wolbachia* a los predadores de mosquitos
- No hay transferencia horizontal de *Wolbachia* al ambiente (suelo, plantas...)
- Presente en 60% especies de insectos
- · Está en Colombia
- Produce efectos en el insecto que son útiles para el control de ETVs

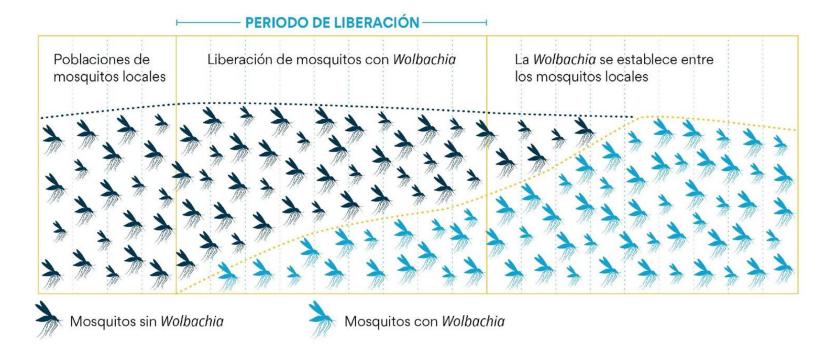






Nuestro método innovador Video

Establecimiento de la Wolbachia en las comunidades de mosquitos locales







vww.csiro.au

Risk Analysis on the Australian release of *Aedes aegypti* (L.) (Diptera: Culicidae) containing *Wolbachia*

Murphy, B, Jansen, C, Murray, J & De Barro, P

March 2010

Vietnam Eliminate Dengue Project

Risk Assessment of the Pilot Release of Aedes aegypti mosquitoes containing Wolbachia

Hanoi September 2011

Evaluation Team

Prof. Truong Quang Hoc, DSc.

Prof. Truong UyenNinh, PhD

Nguyen Van Tuat, PhD

Nguyen Viet Hung, PhD

Nguyen Dinh Cuong, MD. MPH





Cambios en el ecosistema

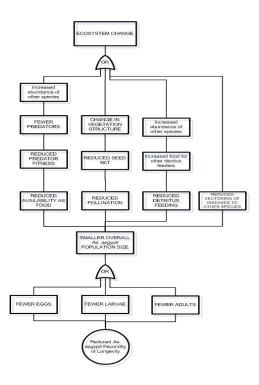


Figure 9. Expert derived fault tree describing the ecological implications of reduced Ae. aegypti life span or fecundity.

Table 6. Hazard definitions for 'Ecosystem Change' fault tree.

Name	Type		scription
Change in	Hazard		The vegetative composition of an area changes because of
vegetation Structure		Ш	changes (decline) in plant fecundity.
Ecosystem change	Hazard		The release leads to ecosystem change or removal of ecosystem services.
Fewer adults	Hazard		Fewer Ae. aegypti adults present in environment.
Fewer eggs	Hazard		Wolbachia Ae. aegypti has lower fecundity than naturally occurring Ae. aegypti.
Fewer larvae	Hazard		Fewer Ae. aegypti larvae present in environment.
Fewer predators	Hazard		Fewer predators occur in ecosystem because of reduced prey.
Increased abundance of other species (x2)	Hazard		Some species increase in numbers because of a reduction in predation or increased availability of food (e.g. detritus) and reduced competition because of smaller Ae. aegypti populations.
Increased food for other detritus feeders	Hazard		Reduced cycling of detritus as a result of lower numbers of Ae. aegypti larvae leaves more food for other detritus feeders.
Reduced Ae. aegypti fecundity or longevity	Und. Event		Wolbachia Ae. aegypti has lower fecundity or shorter life span than naturally occurring Ae. aegypti.
Reduced availability as food	Hazard		Reduced numbers of eggs, larvae and adult Ae. aegypti reduces their availability to predators.
Reduced detritus feeding	Hazard		Lower numbers of Ae. aegypti larvae provide reduced cycling of detritus.
Reduced pollination	Hazard		Plant species receive less pollination by Ae. aegypti males as a result of fewer Ae. aegypti being present to feed on flowers.
Reduced predator fitness	Hazard		Predators have reduced fitness because of reduced Ae. aegypti prey.
Reduced seed set	Hazard		Plant species have lower fecundity because of reduced pollination services.
Reduced vectoring of disease to other species	Hazard		A smaller Ae. aegypti population leads to lower transmission of diseases/arboviruses.
Smaller overall Ae. aegypti population size	Hazard		Mean density of Wolbachia Ae. aegypti is lower than that of naturally occurring Ae. aegypti.

Árbol de fallas para la evaluación de seguridad – Impacto ecológico y en la salud humana

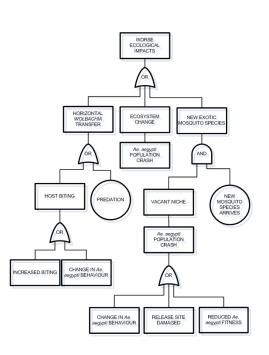


Figure 4. 'Worse ecological impacts 'component of 'Cause More Harm' fault tree.

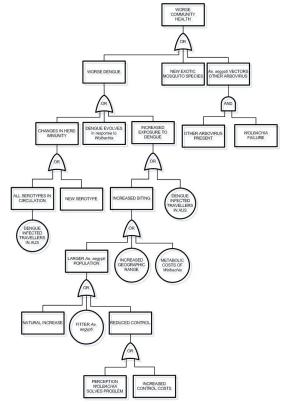


Figure 7. 'Worse community health' component of 'Cause More Harm' fault tree.



SCIENTIFIC REPORTS

OPEN The wMel strain of Wolbachia Reduces Transmission of Zika virus by Aedes aegypti

Received: 28 April 2016 Accepted: 10 June 2016 Matthew T. Aliota¹, Stephen A. Peinado¹, Ivan Dario Velez² & Jorge E. Osorio¹





The wMel Strain of Wolbachia Reduces Transmission of Chikungunya Virus in Aedes aegypti

Matthew T. Aliota¹*, Emma C. Walker¹, Alexander Uribe Yepes², Ivan Dario Velez², Bruce M. Christensen¹, Jorge E. Osorio¹

- 1 Department of Pathobiological Sciences, University of Wisconsin-Madison, Madison, Wisconsin, United States of America, 2 Programa de Estudio y Control de Enfermedades Tropicales (PECET), Universidad de Antioquia, Medellin, Colombia
- * mtaliota@wisc.edu Abstract

Background

New approaches to preventing chikungunya virus (CHIKV) are needed because current methods are limited to controlling mosquito populations, and they have not prevented the invasion of this virus into new locales, nor have they been sufficient to control the virus upon arrival. A promising candidate for arbovirus control and prevention relies on the introduction of the intracellular bacterium Wolbachia into Aedes aegypti mosquitoes. This primarily has been proposed as a tool to control dengue virus (DENV) transmission; however, evidence suggests Wolbachia infections confer protection for Ae. aegypti against CHIKV. Although this approach holds much promise for limiting virus transmission, at present our understanding of the ability of CHIKV to infect, disseminate, and be transmitted by wMel-infected Ae. aegypti currently being used at Wolbachia release sites is limited.

G OPEN ACCESS

Citation: Aliota MT, Walker EC, Uribe Yepes A. Dario Velez I, Christensen BM, Osorio JE (2016) The wifel Strain of Wolbachia Reduces Transmission of Chikungunya Virus in Aedes aegypti. PLoS Negl Trop Dis 10(4): e0004677. doi:10.1371/journal. pntd.0004677

Editor: Philip M. Armstrong, The Connecticut Agricultural Experiment Station, UNITED STATES

Received: February 25, 2016



Acciones del MAP



1. Perfil comunitario e investigación de **línea de base**



5. Activación de un sistema de PQRS



2. Creación grupo de referencia



 Evaluación del nivel de aceptación de las liberaciones



3. Campañas de información y comunicación



 7. Liberaciones de mosquitos portadores de Wolbachia



4. Actividades de participación comunitaria



8. Monitoreo del aumento de la Wolbachia



 Evaluación de la percepción de la comunidad después de terminadas las liberaciones















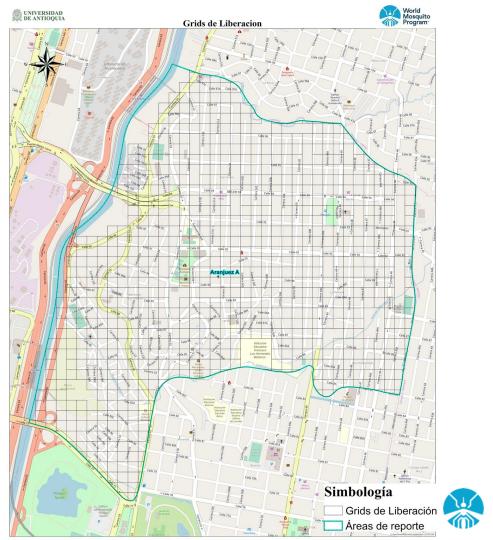




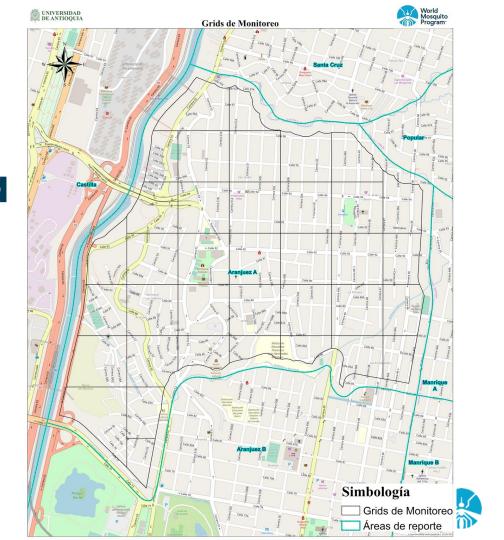




Preparación para actividades de liberación



Preparación para el despliegue del sistema de monitoreo



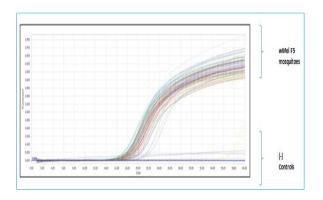


OVITRAMPAS



TRAMPAS BG-SENTINNEL

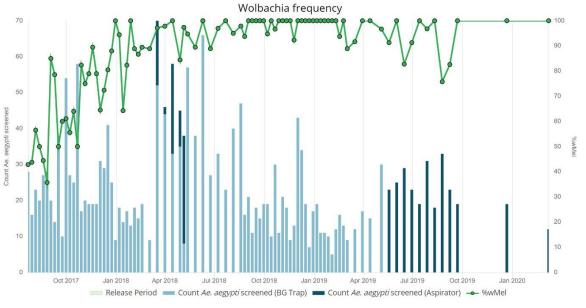


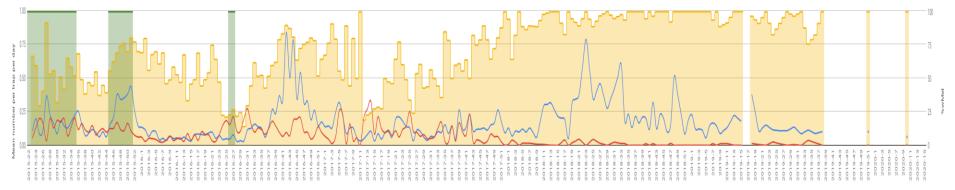


Real time PCR for Wolbachia and DENV



wMel Infection in Paris









FIRST MEETING OF THE TECHNICAL ADVISORY GROUP ON PUBLIC HEALTH ENTOMOLOGY

PAN AMERICAN HEALTH ORGANIZATION

SUMMARY

The first meeting of the Technical Advisory Group on Public Health Entomology (TAG PHE), was held from 8 to 10 March 2016 at the PAHO headquarters in Washington, DC. The topics discussed covered two scenarios: 1) vector control and prevention actions in regards to the current public health emergency of international concern (PHEIC) declared by WHO on the epidemic of Zika virus infections in the Americas, and 2) review and analysis of the actions that can be implemented in the medium and long term for the prevention and control of vectors responsible for vector-borne diseases (VBD) in the Americas.

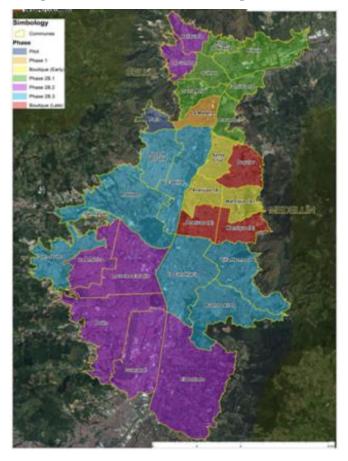
Encourage the rapid, robust and accelerated evaluation of new and supplemental tools for
 Aedes control, such as Wolbachia infections and genetically modified insect technology to
 ensure adequate technical cooperation and funding for this purpose.

MONITORING AND EVALUATION

- Develop and provide a protocol for penetration testing of insecticides, and testing of biological
 efficacy, especially in countries where the situation of insecticide resistance is unknown.
- Evaluate the actions of vector control, field operations, work equipment and operational
 procedures used in prevention and control activities; in terms of impact of actions and costeffectiveness, and the measured impact on disease transmission and incidence. The method
 and evaluation of most-productive mosquito larval containers should be included.
- The countries should support the evaluation of new tools, without losing focus on the emergency and its corresponding response to their level of scale, feasibility, cost effectiveness and acceptability by the community.



Expansión Bello y Medellín



- 2.750 000 habitantes
- 100 km²
- Población heterogénea: todos los niveles económicos y educativos
- Equipo de 150 personas

RETOS

- Permisos
- Desarrollo de Modelo Epidemiológico
- Modelo de Aceptación Pública
- Infraestructura
- Cultivo masivo
- Liberaciones
- QA/QC

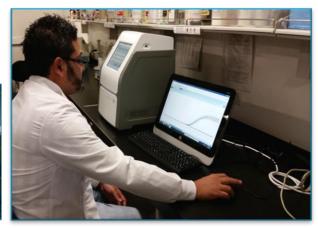


Construcción de infraestructura



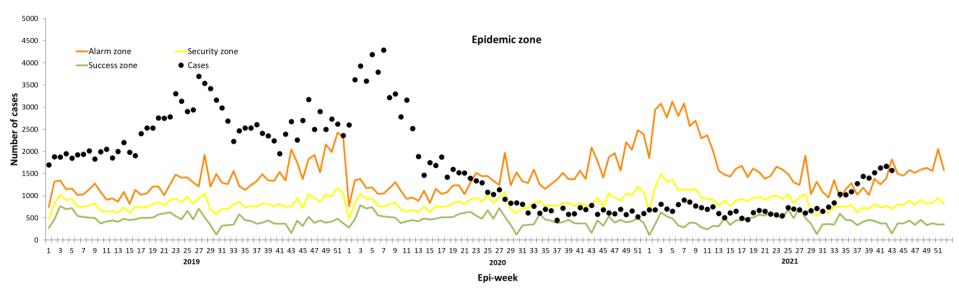








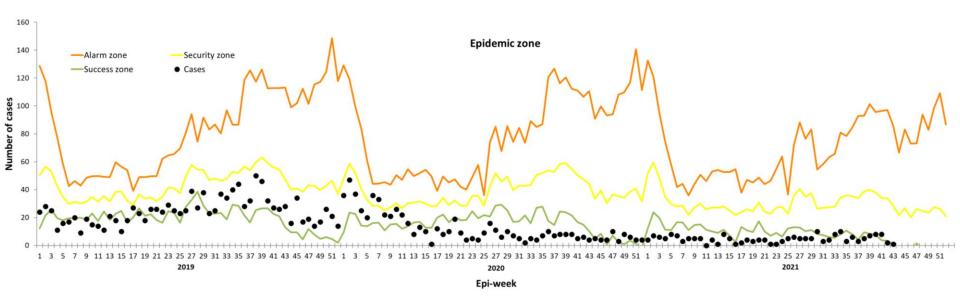
Dengue Endemic Channel, Colombia 2019-2021* Epi week 42 (17th – 23th October, 2021)



^{*}Calculation based on geometric means of incidence proportions (PI) per 100,000 inhabitants and their confidence intervals (95% CI). Bortman, Marcelo (1999). Epidemic years are excluded.

Source: Sivigila, National Health Institute, Colombia, 2021

Dengue Endemic Channel, Medellín 2019-2021* Epi week 43 (24th – 30th October, 2021)



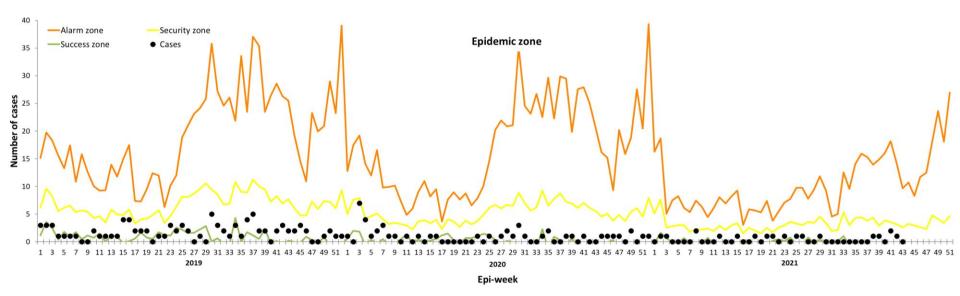
^{*}Calculation based on geometric means of incidence proportions (PI) per 100,000 inhabitants and their confidence intervals (95% CI). Bortman, Marcelo (1999). Epidemic years are excluded.

Source: Medellin local Health Department, 2021

Total notified cases = 2019 (1313), 2020 (631), 2021 (200, until week 43)



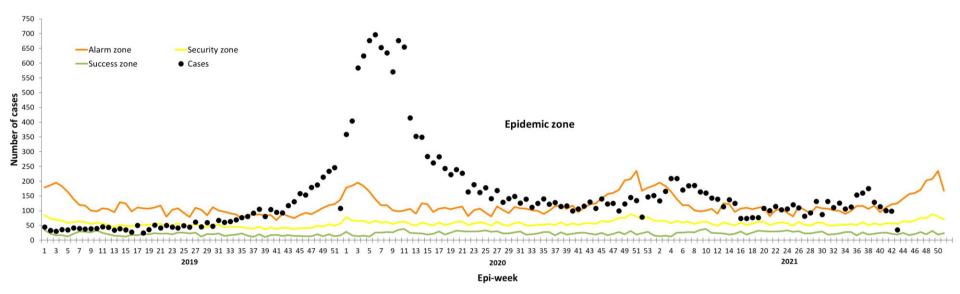
Dengue Endemic Channel, Bello 2019-2021* Epi week 43 (24th – 30th October, 2021)



^{*}Calculation based on geometric means of incidence proportions (PI) per 100,000 inhabitants and their confidence intervals (95% CI). Bortman, Marcelo (1999). Epidemic years are excluded.

Source: Bello local Health Department, 2021

Dengue Endemic Channel, Cali 2019-2021* Epi week 43 (24th – 30th October, 2021)



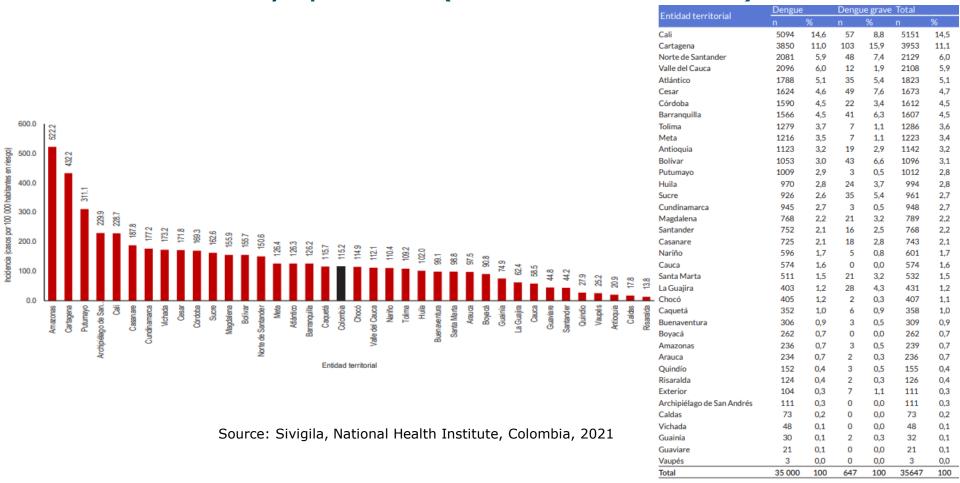
^{*}Calculation based on geometric means of incidence proportions (PI) per 100,000 inhabitants and their confidence intervals (95% CI). Bortman, Marcelo (1999). Epidemic years are excluded.

Source: Cali local Health Department, 2021

Total notified cases = 2019 (3975), 2020 (13 555), 2021 (5408, until week 43)



Incidence rate and number of dengue cases by territorial entity, Epi week 43 (24th – 30th October, 2021)



Evidencias internacionales Evaluación de impacto



Evaluación de impacto en el mundo

VIETNAM Vinh Luong LIBERACIONES COMPLETAS 2018 1 km² 2 12,000 Incidencia del dengue ↓ 86% comparado con el vecino, la ciudad de Nha Trang

INDONESIA

Yogyakarta LIBERACIONES COMPLETAS 2018

- ♦ 17 km²

 \$\text{QL} 241,000
- Incidencia del dengue ↓ 72% En área de estudio piloto.
 - 다 En paralelo, un ensayo controlado aleatorizado está en curso hasta el 2020

Llave

Área de liberación

Población

studio clínico

Vigilancia rutinaria de salud pública

AUSTRALIA

Cairns y Townsville LIBERACIONES COMPLETAS 2018

- ⊕ 240 km² QQ 275,000
- No brotes de dengue en los últimos 5 años

El WMP comenzó recientemente proyectos en Fiji, Vanuatu, Kiribati y México, con resultados esperados en 2020.

COLOMBIA

Medellín y Bello LIBERACIÓN EN CURSO

⊕ 77 km² Д№ 1,502,000

No hay brotes de dengue en las zonas alcanzadas con *Wolbachia*

Estudio de casos y controles en curso a lo largo 2019

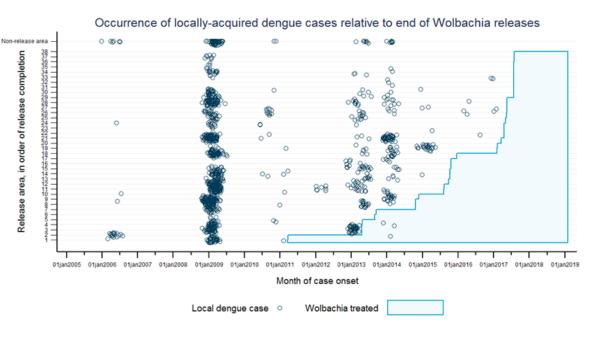
BRASIL
Río de Janeiro y Niteroi
LIBERACIÓN EN CURSO

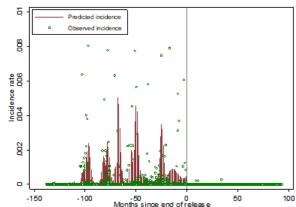
106 km² (1,154,000

Mesultados preliminares de despliegues en toda la ciudad indican ↓del 46% en incidencia del dengue



Interrupted time series analysis of dengue case notifications Example (from Cairns)

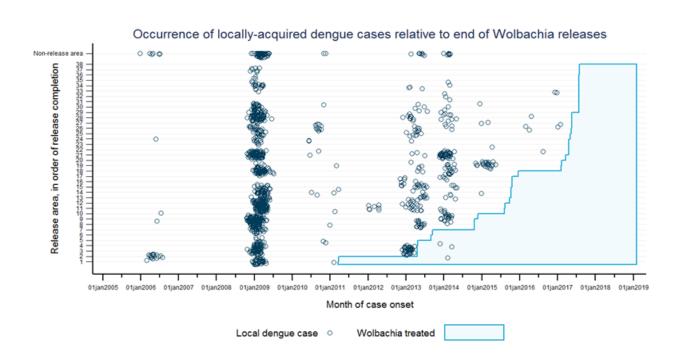




IRR = 0.028 (95%CI 0.014-0.053) corresponding to a 97.2% (94.7 - 98.6%) reduction in the incidence of locally-acquired dengue.



Interrupted time series analysis of dengue case notifications. Example (from Cairns)





Contribuimos al logro de los Objetivos de Desarrollo Sostenible

- 1. Fin de la pobreza
- 3. Salud y bienestar
- 17. Alianzas para lograr los objetivos





