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OFICINA REGIONAL PARA LAS Américas

Estrategia de Gestión Integrada para la prevención y control del Dengue en las Américas

Programa Regional de Dengue OPS/OMS



dengue



Otras tecnologías en el control del *Aedes aegypti*

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Asesor Regional de Entomología

Enfermedades Desatendidas, Tropicales y Transmitidas por Vectores (VT)

OPS/OMS, Washington DC

29 de mayo, 2014



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Para enfermedades tales como Dengue el control de vectores es la única forma de prevenir la transmisión.

Se ha demostrado que las intervenciones recomendadas por la OPS/OMS de control de transmisión por *Aedes aegypti*, son efectivas cuando se implementan adecuadamente.



Como ha funcionado el control de *Ae. aegypti* en las Américas

- Muchos países tienen programas de prevención y control de enfermedades transmitidas por vectores que son remanentes de los programas de erradicación de la malaria o fiebre amarilla;
- Muchos países usan el Control químico como intervención prioritaria (transmisión/densidad) ;
- Las aplicaciones espaciales y uso de larvicidas no son suficientemente evaluadas.





Retos para lograr efectividad en el control de la transmisión

- El incremento de la resistencia a los insecticidas y el número limitado de nuevos productos ponen en peligro la efectividad del control regular o ante brotes/epidemia;

No hay nuevos insecticidas adulticidas en salud pública en uso en los últimos 20 años

1940-45	DDT				
1945-50	Lindane				
1950-55	malathion				
1955-60					
1960-65	fenitrothion	propoxur			
1965-70	chlorpyrifos-methyl				
1970-75	pirimiphos-methyl	bendiocarb	permethrin		
1975-80	cypermethrin				
1980-85	alpha-cypermethrin	cyfluthrin	lambda-cyhalothrin	deltamethrin	bifenthrin
1985-90	etofenprox				
1990-95					
1995-00					
2000-05					

Source WHO

- Organochlorines
- Organophosphates
- Carbamates
- Pyrethroids

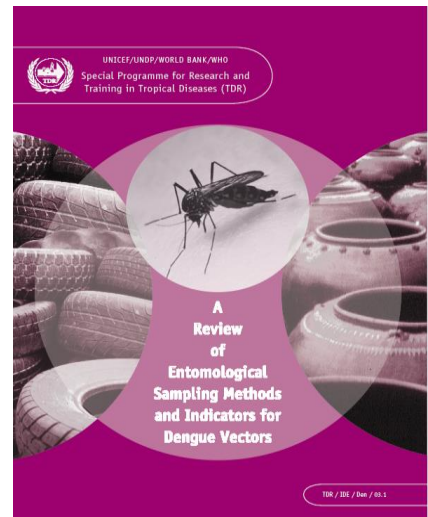
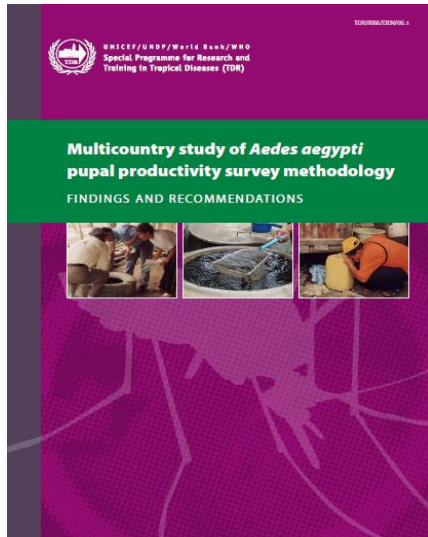


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Vigilancia del *Aedes aegypti*



Características

- identifican los criaderos predominantes y obtiene los índices tradicionales (índice de viviendas, índice de Breteau e índice de recipientes, pupas);
- Realizada en un promedio de 60 días o más, una semana (LIRAa);
- generalmente proporciona índices para grandes áreas (barrios) o estratos (LIRAa);
- dirige las actividades de control;

Levantamiento rápido para *Aedes aegypti* (LIRAa)



dengue

Estrategia de Gestión Integrada para la prevención y control del Dengue en las Américas

Nuevas tecnologías para la vigilancia del *Aedes aegypti*



MosquiTRAP



Trampa BGS



AdulTRAP



Ovitrapa



D-vac



Autocida Gravid Ovitrap (AGO)



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Trampas letales

Componentes:

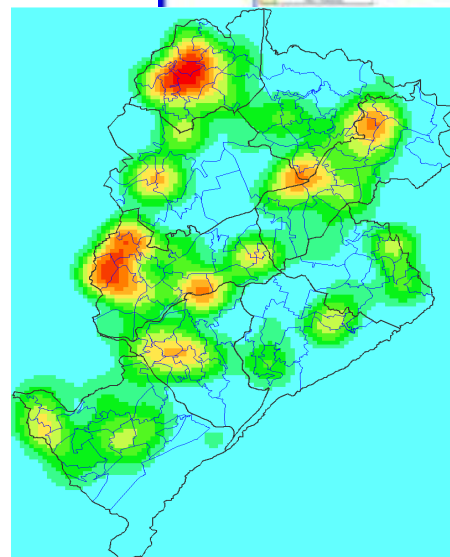
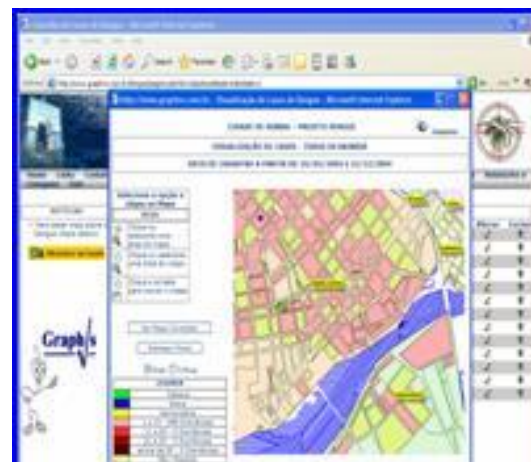
- Colores de cuerpo de la trampa y el tamaño son visualmente atractivos para los mosquitos
- La trampa contiene agua con una mezcla de 4 especies de bacterias que emiten un olor atractivo para las hembras
- El depósito contiene también un larvicida, spinosad, que ha sido aprobado para uso en agua potable de la OMS
- Recubre el interior de la trampa es un tejido de malla tratado con un adulticida piretroides, Alfa Cipermetrina (hecho de LLIN DuraNet ®)





Estrategia de Gestión Integrada para la prevención y control del Dengue en las Américas

Nuevos dispositivos para recolección/análisis de la información en terreno para la vigilancia y control del *Aedes aegypti*



Parámetros do Kernel
Tema de Eventos : Pneus
Valor do raio : 2.000000e+003
Cálculo:Densidade
Função:Quártico

Valores de Kernel

-0.0000000000000001	~ 0.000000214213100
0.000000214213100	~ 0.000000428426200
0.000000428426200	~ 0.000000642639300
0.000000642639300	~ 0.000000856852400
0.000000856852400	~ 0.000001071065500
0.000001071065500	~ 0.000001285278600
0.000001285278600	~ 0.000001499491700
0.000001499491700	~ 0.000001713704800
0.000001713704800	~ 0.000001927917900
0.000001927917900	~ 0.000002142131001

0 5000 10000 15000
Meters



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REVIEW ARTICLE

Effect of dengue vector control interventions on entomological parameters in developing countries: a systematic review and meta-analysis

T. E. ERLANGER¹, J. KEISER² and J. UTZINGER¹

¹Department of Public Health and Epidemiology and ²Department of Medical Parasitology and Infection Biology, Swiss Tropical Institute, Basel, Switzerland

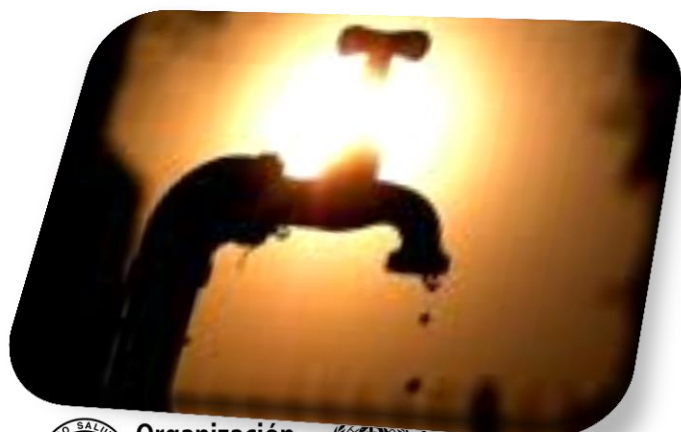
- Manejo integrado de vectores fue encontrado como el método más eficaz para reducir la IR, IV y IB, resultando en valores relativa efectividad combinada de 0.12 (c.i. 95 0.02 – 0,62), 0,17 (c.i. 95 0.02 – 1,28) y 0.33 (c.i. 95 0.22 – 0,48), respectivamente
- Manejo ambiental mostró una efectividad relativamente baja de 0,71 (c.i. 95 0.55 – 0,90) para el IB, 0.49 (c.i. 95 0.30 – 0,79 para la IR) y 0.43 (c.i. 95 0.31 – 0,59) para el IV.
- Control biológico (efectividad relativa para la IR: 0.18) generalmente dirigidos a un pequeño número de personas (tamaño mediana de la población: 200; 20 – 2500), mientras que el manejo integrado de vectores se centró en las poblaciones más grandes (mediana: 12 450; rango: 210 – 9 600 000).





Recipientes a prueba de
mosquitos

3 millones de capas
Brasil



**Avaliação da efetividade da utilização da
metodologia Communication and Marketing
Integrated for Behaviour Impact - COMBI
para controle da dengue em duas
localidades do Brasil**

by Braga, Ivenise Leal, 2011



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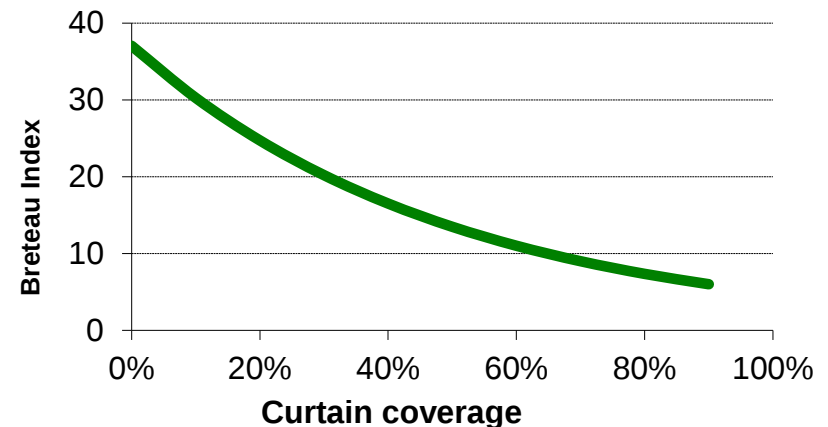


Material impregnados con insecticida



The effectiveness of insecticide treated curtains for *Aedes* control in Valera, Venezuela (Vanlerberghe et al., 2009)

Modeled effect of ITC coverage



“Cobertura inicial alta, descenso rápido a largo plazo en escenarios sin actividades de promoción o comunicación”

“Las cortinas pueden reducir los índices, pero el efecto depende de la cobertura poblacional, y si la cobertura esta muy baja el efecto desaparece”

Control Biológico



Betta splendens

EVALUATION OF THE IMPACT ON *AEDES AEGYPTI* INFESTATION IN CEMENT TANKS OF THE MUNICIPAL DISTRICT OF CANINDÉ, CEARÁ, BRAZIL AFTER USING THE *BETTA SPLENDENS* FISH AS AN ALTERNATIVE BIOLOGICAL CONTROL. Rev. Soc. Bras. Med. Trop. vol.37 no.5 Uberaba Sept./Oct. 2004

CONTROL OF *AEDES AEGYPTI* LARVAE IN HOUSEHOLD WATER CONTAINERS BY CHINESE CAT FISH. Bull World Health Organ. 1987; 65(4): 503–506



Mesocyclops

ELIMINATION OF DENGUE BY COMMUNITY PROGRAMS USING *MESOCYCLOPS*(COPEPODA) AGAINST *AEDES AEGYPTI* IN CENTRAL VIETNAM

VU SINH NAM, NGUYEN THI YEN, TRAN VU PHONG, TRUONG UYEN NINH, LE QUYEN MAI, LE VIET LO, LE TRUNG NGHIA, AHMET BEKTAS, ALISTAIR BRISCOMBE, JOHN G. AASKOV, PETER A. RYAN, AND BRIAN H. KAY
Am. J. Trop. Med. Hyg., 72(1), 2005, pp. 67-73



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Am. J. Trop. Med. Hyg., 82(6), 2010, pp. 1053–1059

doi:10.4269/ajtmh.2010.09-0603

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Bacillus thuringiensis var. *israelensis* (Bti) Provides Residual Control of *Aedes aegypti* in Small Containers

Scott A. Ritchie,* Luke P. Rapley, and Seleena Benjamin

School of Public Health, Tropical Medicine and Rehabilitation Sciences, James Cook University, Cairns, Queensland, 4870 Australia; Public Health, Valent BioSciences Corporation, Singapore

Abstract. We examined the use of megadoses of VectoBac WG for residual control of *Aedes aegypti* in 2-L plastic buckets. Doses of 10×, 20×, and 50× the recommended rate of 8 mg/L provided ≥ 90% control for 8, 8, and 23 weeks, respectively. There was no significant difference in mortality between dry (neat) or aqueous mixture of VectoBac WG. Pretreatment of dry containers up to 8 weeks before flooding did not significantly decrease efficacy through 11 success weeks. Thus, megadoses of dry formulations of Bti can be used for residual control of *Ae. aegypti* in small containers. Furthermore, these doses use small amounts of product (0.08–0.4 g/L) that is more practical to measure than the minute amounts (0.008 g/L) required by the recommended rate, and cost US\$2.18 to treat 50 Cairns yards containing an average total of 80 containers. This method could also be used to control *Aedes albopictus*.

Journal of the American Mosquito Control Association, 26(1):67–87, 2010

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A REVIEW OF SPINOSAD AS A NATURAL PRODUCT FOR LARVAL MOSQUITO CONTROL

MARK B. HERTLEIN,¹ COSTAS MAVROTAS,² CHRISTIAN JOUSSEAU,³ MICHAEL LYSANDROU,² GARY D. THOMPSON,¹ WILLIAM JANY⁴ AND SCOTT A. RITCHIE⁵

African Entomology 21(2):349–353. 2013

doi: <http://dx.doi.org/10.4001/003.021.0206>

Effectiveness of Slow-Release Tablet Formulations of the IGR Diflubenzuron and the Bioinsecticide Spinosad Against Larvae of *Aedes aegypti* (L.)

M.S. Saleh *, O.A. Abuzinadah , Kh.M. Al-Ghamdi , A.A. Alsagaf & J.A. Mahyoub

Department of Biological Science, Faculty of Science, King Abdul-Aziz University, Jeddah, Saudi Arabia

* Author for correspondence. E-mail: diptera2012@gmail.com

The larvicidal effectiveness of slow-release tablet formulations of the chitin synthesis inhibitor Dudim® (diflubenzuron) and the bioinsecticide Natular™ (spinosad) against mosquito larvae of *Aedes aegypti* (L.) has been evaluated. The results showed that the test formulations provided long-term residual control against the larvae. Effective control giving 90–100 % inhibition of adult emergence was achieved for 10 weeks post-treatment for diflubenzuron and 7 weeks for spinosad. In addition, larval treatments with slow-release diflubenzuron formulations led to a marked prolongation in the time needed for blood meal digestion and a reduction in the reproductive potential of adult survivors. On the other hand, larval treatments with slow-release formulations of spinosad tablets affect neither the time of blood meal digestion nor the reproductive capacity of mosquito adults that emerged from surviving larvae.

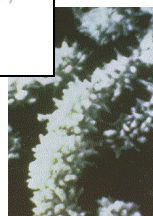


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EFFICACY OF AQUATAIN, A MONOMOLECULAR FILM, FOR THE CONTROL OF MALARIA VECTORS IN RICE PADDIES (2011)

Tullu Bukhari, Willem Takken, Andrew K. Githeko, Constantianus J. M. Koenraadt
PLoS ONE 6(6): e21713. doi:10.1371/journal.pone.0021713



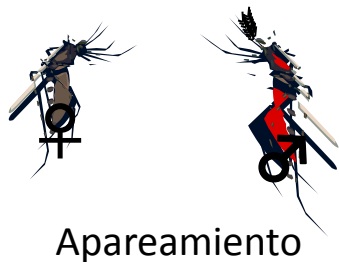
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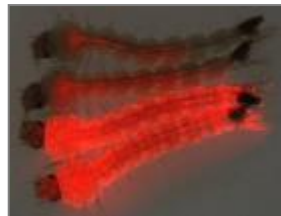
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Mosquitos genéticamente modificados



Apareamiento



Larvas mueren
en estadio temprano



No existe producción de
adultos



No existen picaduras de
hembras adultas

Reducción en la
transmisión de
enfermedad



LETTERS

193423

nature
biotechnology

Field performance of engineered male mosquitoes

Angela F Harris^{1,2}, Derric Nimmo³, Andrew R McKemey³, Nick Kelly¹, Sarah Scaife¹, Christl A Donnelly⁴, Camilla Beech¹, William D Petrie¹ & Luke Alphey^{1,3}

Dengue is the most medically important arthropod-borne viral disease, with 50–100 million cases reported annually worldwide¹. As no licensed vaccine or dedicated therapy exists for dengue, the most promising strategies to control the disease involve targeting the predominant mosquito vector, *Aedes aegypti*. However, the current methods to do this are inadequate. Various approaches involving genetically engineered mosquitoes have been proposed^{2–15}, including the release of transgenic sterile males^{16–18}. However, the ability of laboratory-reared, engineered male mosquitoes to effectively compete with wild males in terms of finding and mating with wild females, which is critical to the success of these strategies, has remained untested. We report data from the first open-field trial involving a strain of engineered mosquito. We demonstrated that genetically modified male mosquitoes, released across 10 hectares for a 4-week period, mated successfully with wild females and fertilized their eggs. These findings suggest the feasibility of this technology to control dengue by suppressing field populations of *A. aegypti*.

We therefore set out to examine this in the field, using as the RIDL system COSS3A, an engineered strain of *A. aegypti*, the principal vector of dengue. COSS3A carries a reproducible lethal genetic system to ensure that mosquitoes can be reared to maturity in the presence of tetracycline in the laboratory, although their offspring will not survive to adulthood in the absence of the repressant. The strain also expresses a fluorescent marker gene to allow detection^{19,20–22}. We selected a field site on Grand Cayman—the largest of the Cayman Islands, a British Overseas Territory in the Caribbean—on the basis of several criteria outlined in Online Methods. Dengue is rare on Grand Cayman but common in the region and an ongoing threat as long as the mosquito vector is present. High levels of insecticide resistance in *A. aegypti* in Grand Cayman²³ and neighboring regions have highlighted the need for alternative methods of control.

The success of releasing sterile male insect pests relies critically on, first, the ability of RIDL males to compete successfully with wild males for mating with wild females in the field, and second, the ability of the inherited RIDL construct to kill some or all of the progeny of such matings. In principle, field strains might vary in their pro-

Review

Genetic control of *Aedes* mosquitoes

Luke Alphey^{1,2}, Andrew McKemey¹, Derric Nimmo¹, Marco Neira Oviedo^{1,3}, Renaud Lacroix¹, Kelly Matzen¹, Camilla Beech¹

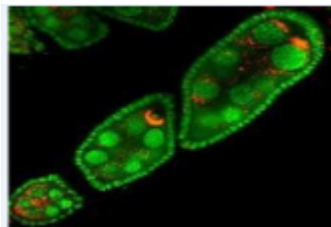
¹Oxitec Limited, 71 Milton Park, Oxford OX14 4RX, UK, ²Department of Zoology, University of Oxford, South Parks Road, Oxford OX1 3PS, UK, ³Center for Infectious Disease Research, Pontificia Universidad Católica del Ecuador, Quito, Ecuador

Aedes mosquitoes include important vector species such as *Aedes aegypti*, the major vector of dengue. Genetic control methods are being developed for several of these species, stimulated by an urgent need owing to the poor effectiveness of current methods combined with an increase in chemical pesticide resistance. In this review we discuss the various genetic strategies that have been proposed, their present status, and future prospects. We focus particularly on those methods that are already being tested in the field, including RIDL and *Wolbachia*-based approaches.

Keywords: *Aedes*, Genetic engineering, RIDL, SIT, IIT, *Wolbachia*, Dengue



Aedes & Wolbachia



FÊMEA ♀		MACHO ♂		PROLE
	+		=	
	+		=	
	+		=	

- Mosquito com Wolbachia
- Mosquito sem Wolbachia
- Ovos nascem, mas não dão origem a larvas



ELIMINAR A
DENGUE
da vida

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Stable Introduction of a Life-Shortening *Wolbachia* Infection into the Mosquito *Aedes aegypti*

Conor J. McMeniman,¹ Roxanna V. Lane,¹ Bodil N. Cass,¹ Amy W.C. Fong,¹ Manpreet Sidhu,¹ Yu-Feng Wang,^{1,2} Scott L. O'Neill^{1*}

Most pathogens require a relatively long period of development in their mosquito vector before they can be transmitted to a new human host; hence, only older insects are of epidemiological importance. The successful transfer of a life-shortening strain of the inherited bacterial symbiont, *Wolbachia*, into the major mosquito vector of dengue, *Aedes aegypti*, halved adult life span under laboratory conditions. The association is stable, and the *Wolbachia* strain is maternally inherited at high frequency. It is capable of inducing complete cytoplasmic incompatibility, which should facilitate its invasion into natural field populations and its persistence over time. Our data suggest that targeting mosquito age with inherited *Wolbachia* infections may be a viable strategy to reduce the transmission of pathogens such as dengue viruses.

The control of dengue primarily targets *Aedes aegypti*, a domesticated mosquito that prefers to live in and around human habitation (1). With few exceptions, dengue management strategies have been complicated by the inability to completely eradicate *A. aegypti* from urban settings and the ineffective application of long-lasting vector-control programs (2). This has led

to a worldwide resurgence of dengue and has highlighted the urgent need for novel and sustainable disease-control strategies.

Most pathogens that are transmitted by mosquitoes share a common property; they have to undergo a significant period of development in their insect vector before they can be transmitted to a new host. After a female mosquito ingests an

ncemag.org SCIENCE VOL 323 2 JANUARY 2009

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OPEN ACCESS Freely available online

PLOS PATHOGENS

The Endosymbiotic Bacterium *Wolbachia* Induces Resistance to Dengue Virus in *Aedes aegypti*

Guowu Bian¹, Yao Xu¹, Peng Lu¹, Yan Xie², Zhiyong Xi^{1*}

¹Department of Entomology and Genetics Program, Michigan State University, East Lansing, Michigan, United States of America, ²Center for Statistical Training & Consulting, Michigan State University, East Lansing, Michigan, United States of America

Abstract

Genetic strategies that reduce or block pathogen transmission by mosquitoes have been proposed as a means of augmenting current control measures to reduce the growing burden of vector-borne diseases. The endosymbiotic bacterium *Wolbachia* has long been promoted as a potential vehicle for introducing disease-resistance genes into mosquitoes, thereby making them refractory to the human pathogens they transmit. Given the large overlap in tissue distribution and intracellular localization between *Wolbachia* and dengue virus in mosquitoes, we conducted experiments to characterize their interactions. Our results show that *Wolbachia* inhibits viral replication and dissemination in the main dengue vector, *Aedes aegypti*. Moreover, the virus transmission potential of *Wolbachia*-infected *Ae. aegypti* was significantly diminished when compared to wild-type mosquitoes that did not harbor *Wolbachia*. At 14 days post-infection, *Wolbachia* completely blocked dengue transmission in at least 37.5% of *Ae. aegypti* mosquitoes. We also observed that this *Wolbachia*-mediated viral interference was associated with an elevated basal immunity and increased longevity in the mosquitoes. These results underscore the potential usefulness of *Wolbachia*-based control strategies for population replacement.



VECTOR CONTROL ADVISORY GROUP

Operational procedures

The World Health Organization (WHO) has established a Vector Control Advisory Group (VCAG) on New Tools to serve as an advisory body to WHO on new forms of vector control for malaria and other vector-borne diseases.

Functions

The VCAG has the following functions:

1. To review and assess the public health value, "proof of principle" (epidemiological impact) of new tools, paradigms, approaches and technologies; and
2. To make recommendations on their use for vector control within the context of integrated vector management in a disease or multi-disease settings.

When is a paradigm or category "new"?

A product, technology or approach is considered a new paradigm if one or more of the following conditions are met:

- An intervention or combination of interventions that protects humans against a different group of vectors or in different transmission settings (or contexts), different human populations or via a different mechanism from existing methods, e.g. an odour-baited trap or a genetically-modified insect.
- An intervention where one or more of its characteristics is sufficiently changed that entomological effect alone is not sufficient to imply epidemiological effect, e.g. a resistance-breaking bednet.
- When a product is not adequately described by an existing target product profile (TPP) and validation will result in the development of a new TPP, and hence a new product class.



WHO Pesticide Evaluation Scheme (WHOPES)

WHOPES home

Safety and effectiveness

Quality control

Guidelines for testing

Insecticide resistance

Equipment and application

Partnership

Links and resources

WHO Pesticide Evaluation Scheme: "WHOPES"

50 YEARS OF GLOBAL LEADERSHIP



History – 50 Years of global leadership

In 2010, WHO commemorated 50 years of its Pesticide Evaluation Scheme (WHOPES), established with the approval of the World Health Assembly in 1960. For more than half a century, the programme has been serving as a reference for setting norms and standards for public health pesticides and their life-cycle management.

DG's speech

50 YEARS OF GLOBAL LEADERSHIP

What is WHOPES and what are WHOPES' objectives?

Latest news

WHOPES recommended

- insecticides for IRS
- insecticides for ITNs
- insecticides for space spraying
- long-lasting insecticidal nets (LNs)
Updated 06 February 2014
- Mosquito larvicides

Management of public health pesticides





Recomendaciones

- El control químico adecuadamente implementado sigue siendo necesario para el control de la transmisión, pero dentro del marco de la EGI dengue;
- Lo levantamiento de índices larvarios y de pupas siguen siendo los métodos principales de vigilancia;
- Sistematizar y documentar las nuevas alternativas ya aprobadas para su uso;
- Las nuevas alternativas de vigilancia y control deben seguirse estudiando para generar mayores evidencias (costo/efectividad/sostenibilidad) que suporten su introducción como alternativas a los programas.





Estrategia de Gestión Integrada para la prevención y control del Dengue en las Américas

Obrigado
Thank you
Gracias

**Reunión sobre el
Estado del Arte para la
prevención
y control del dengue
en las Américas**

28 - 29 de mayo, 2014 • Washington DC, USA



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