

#### MESA FORUM | VIRTUAL

# RESPONDING TO THE THREAT OF ANOPHELES STEPHENSI INVASION IN AFRICA

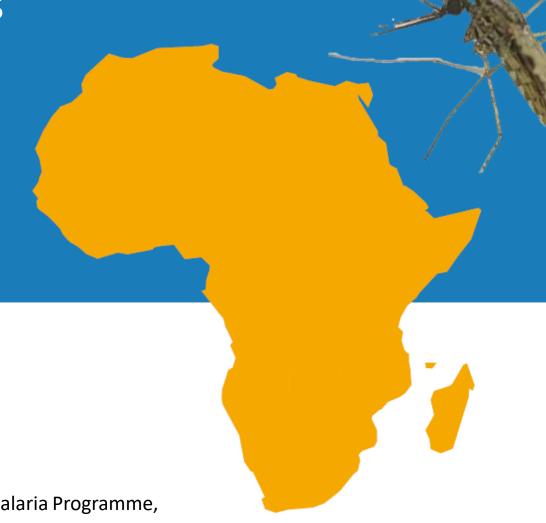


21 February, 2023

2-3:30 PM CET



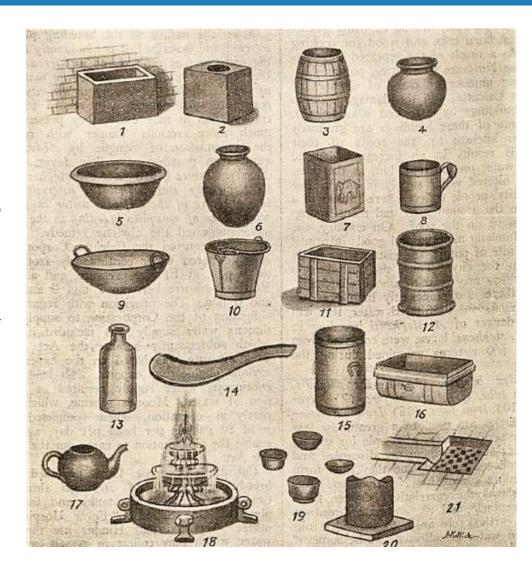
Overview, relevance, and the current status of *An. stephensi's* geographical spread: WHO Threats Map



**Seth Irish**, Technical Officer, Vector Control & Insecticide Resistance Unit, Global Malaria Programme, World Health Organization

# Anopheles stephensi

- Major malaria vector from south Asia
- First reported finding in Africa in 2012
- Flexibility in larval site choice, especially able to use urban larval sites
- Host preference for cattle/goats
- Good biological vector for P. falciparum and P. vivax
- Resistant to many insecticides used for public health



### WHO initiative



Information exchange



Increasing collaboration



Strengthening surveillance



Prioritizing research



Developing guidance

Last Updated: 8/30/2022

There are no studies available with the specific

mena

isecticide class

izecucioe ciass

secticide type

Select

Test typ

SHELL

Years

310

2015

### Malaria Threats Map

Tracking biological challenges to malaria control and elimination



#### VECTOR INSECTICIDE RESISTANCE

Resistance of malaria mosquitoes to insecticides used in core prevention tools of treated bed nets and indoor residual sprays threatens vector control effectiveness



#### PARASITE pfhrp2/3 GENE DELETIONS

Gene deletions among some malaria parasites cause false negative diagnostic test results, complicating case management and control



#### PARASITE DRUG EFFICACY AND RESISTANCE

Resistance of malaria parasites to artemisinin – the core compound of the best available antimalarial medicines – threatens antimalarial drug efficacy

> GREATER MEKONG SUBREGION



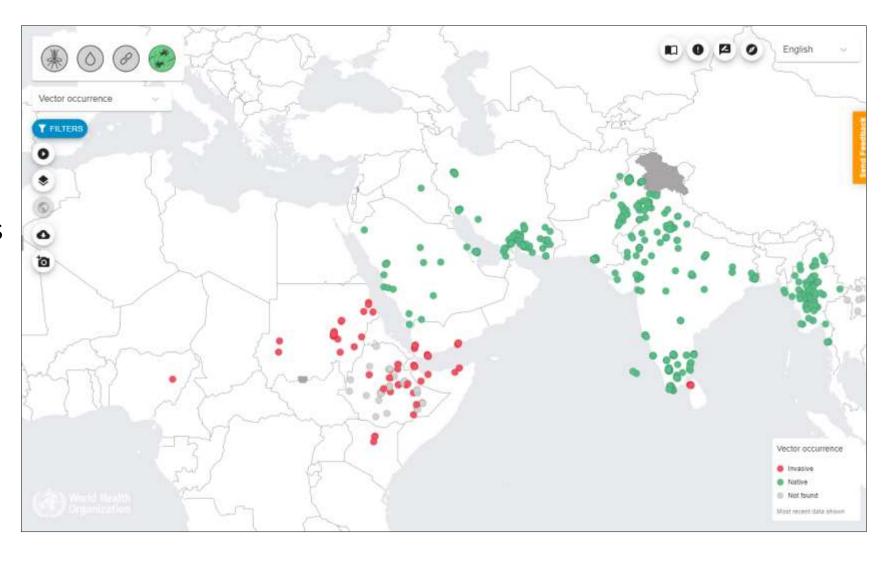
English

#### INVASIVE VECTOR SPECIES

The spread of anopheline mosquito vector species and their establishment in ecosystems to which they are not native poses a potential threat to the control and elimination of malaria

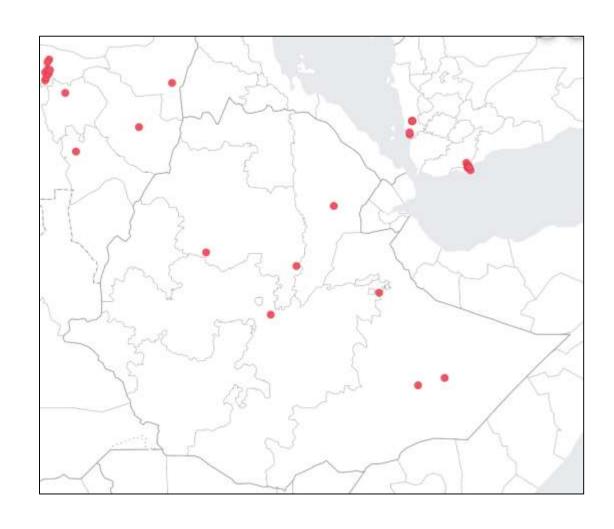
# Tracking the spread

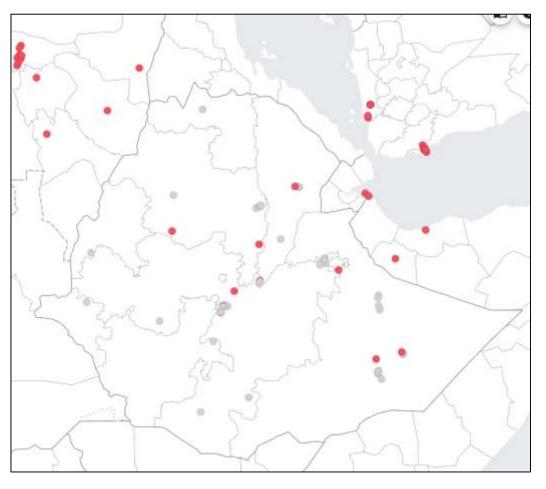
- Malaria Threats Map
  - Native occurrences
  - Invasive occurrences
  - Negative findings



# Value of negative

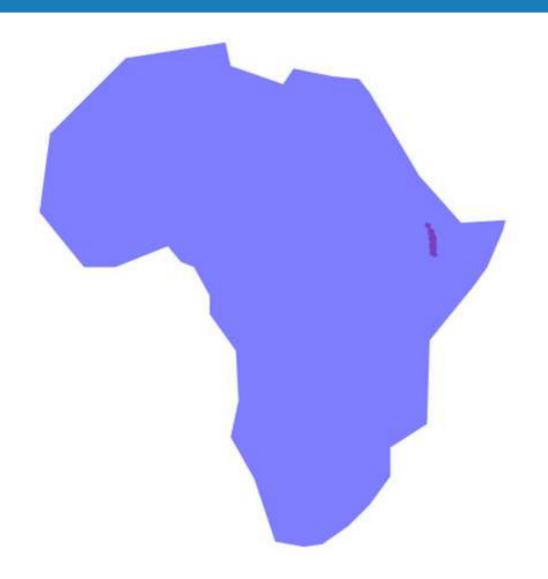
findings

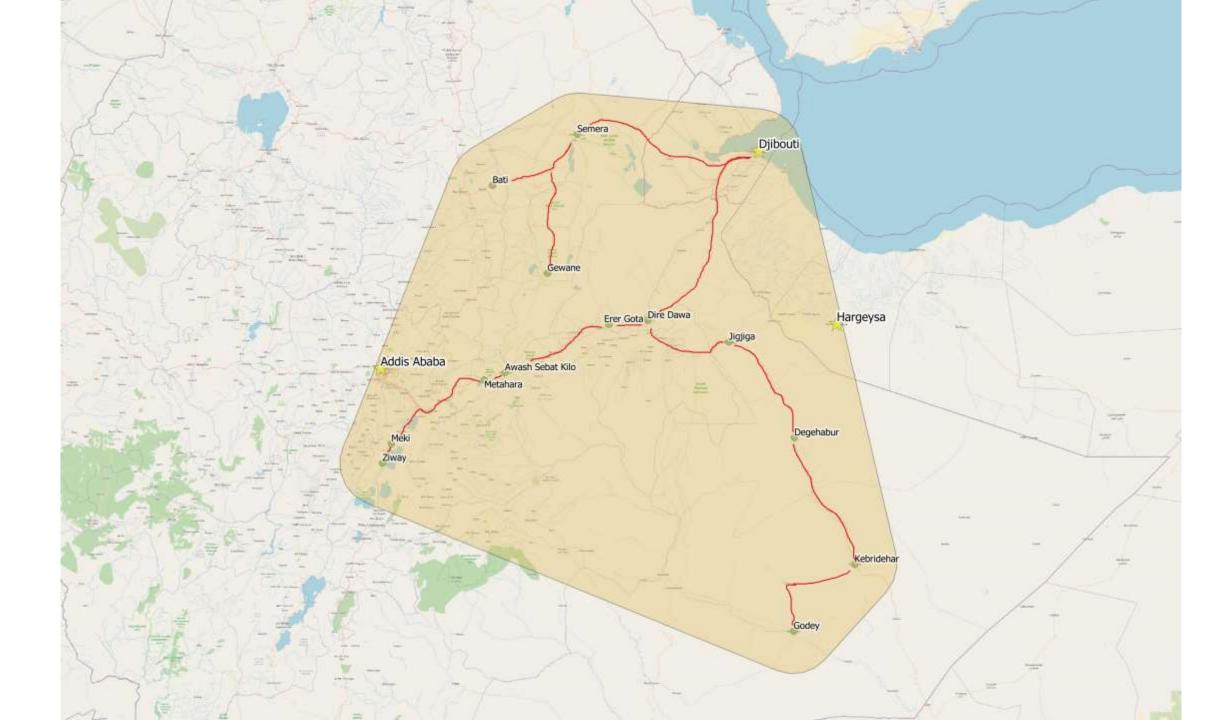




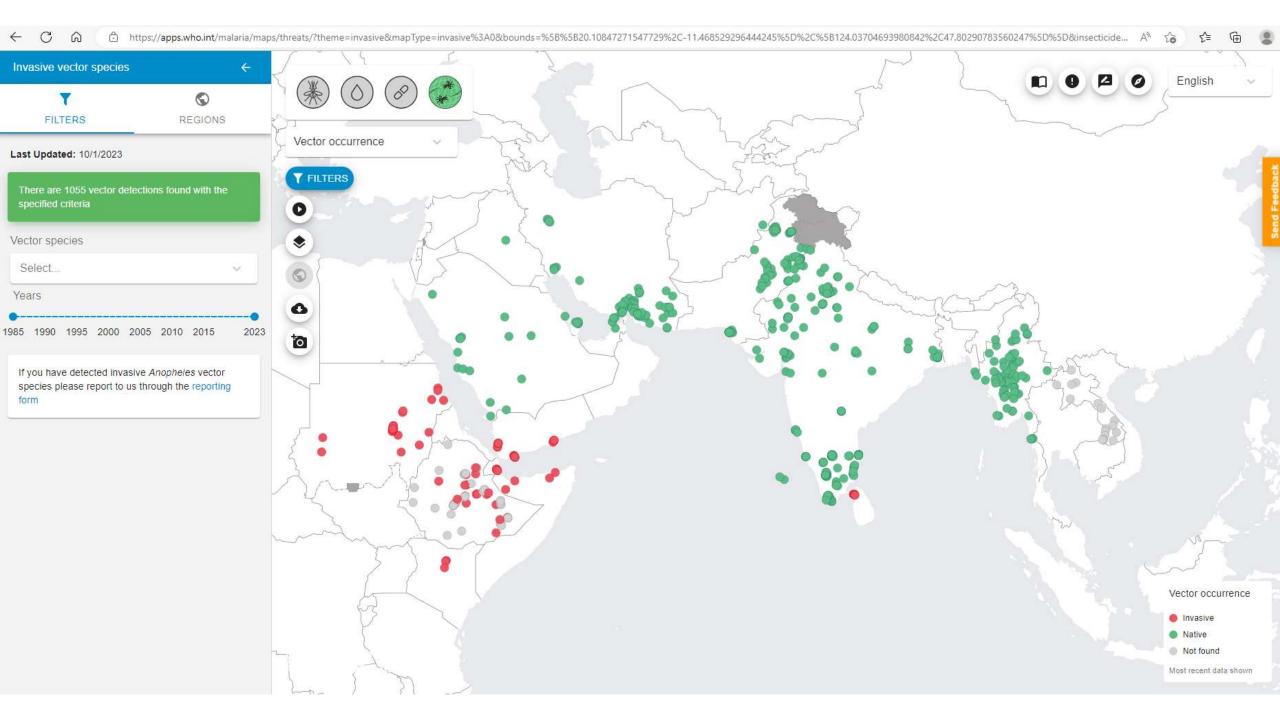
# Implications for elimination

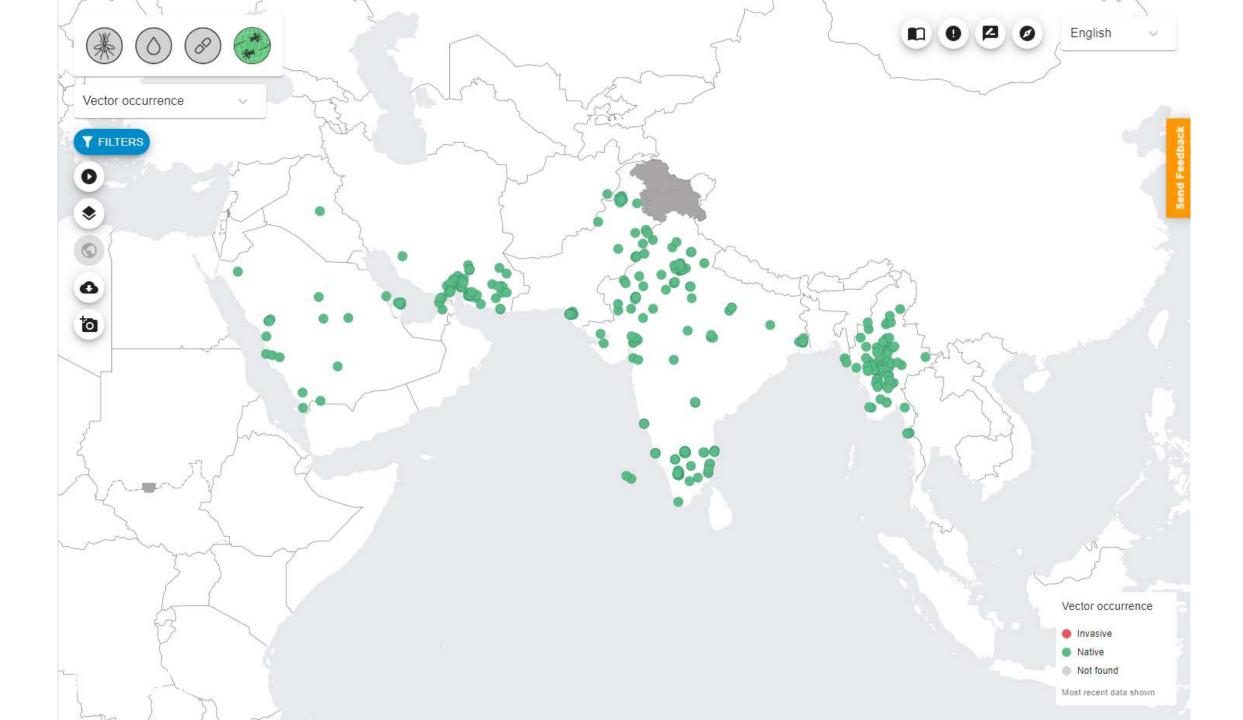
- In Brazil, An. arabiensis spread to an area of approximately 54,000km<sup>2</sup>, just smaller than the area of the country of Togo.
- What area has *An. stephensi* spread to in Africa?

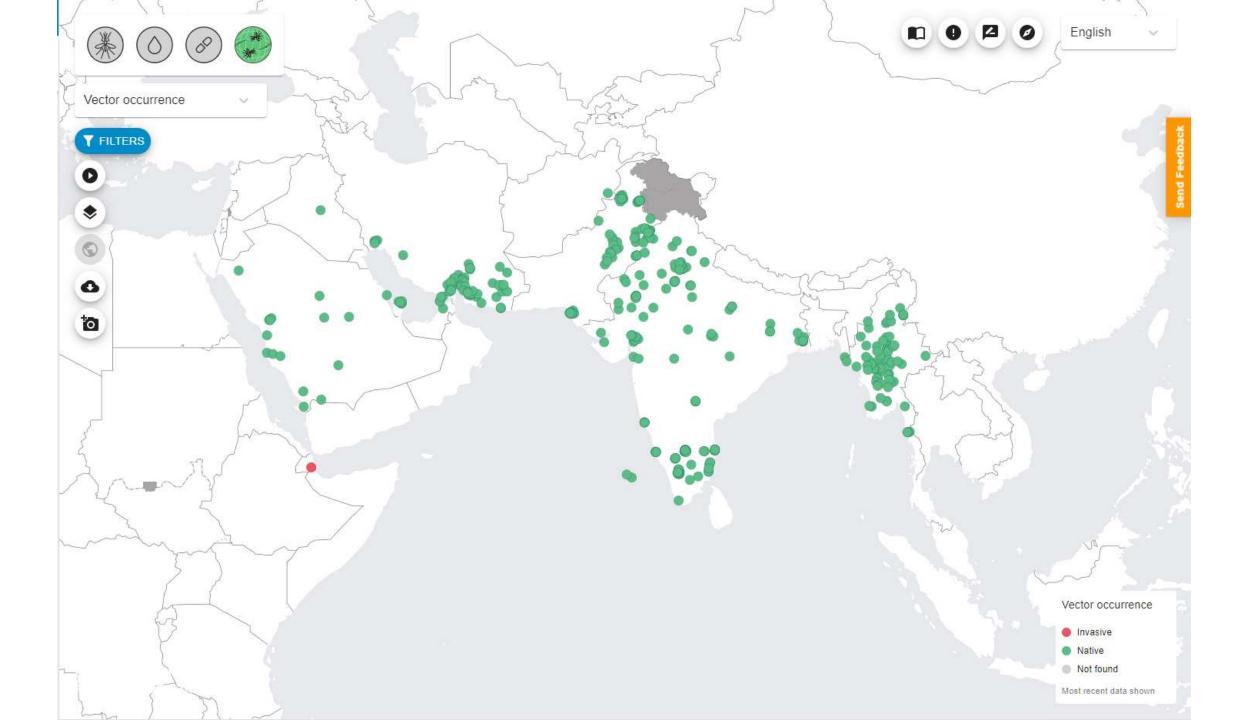


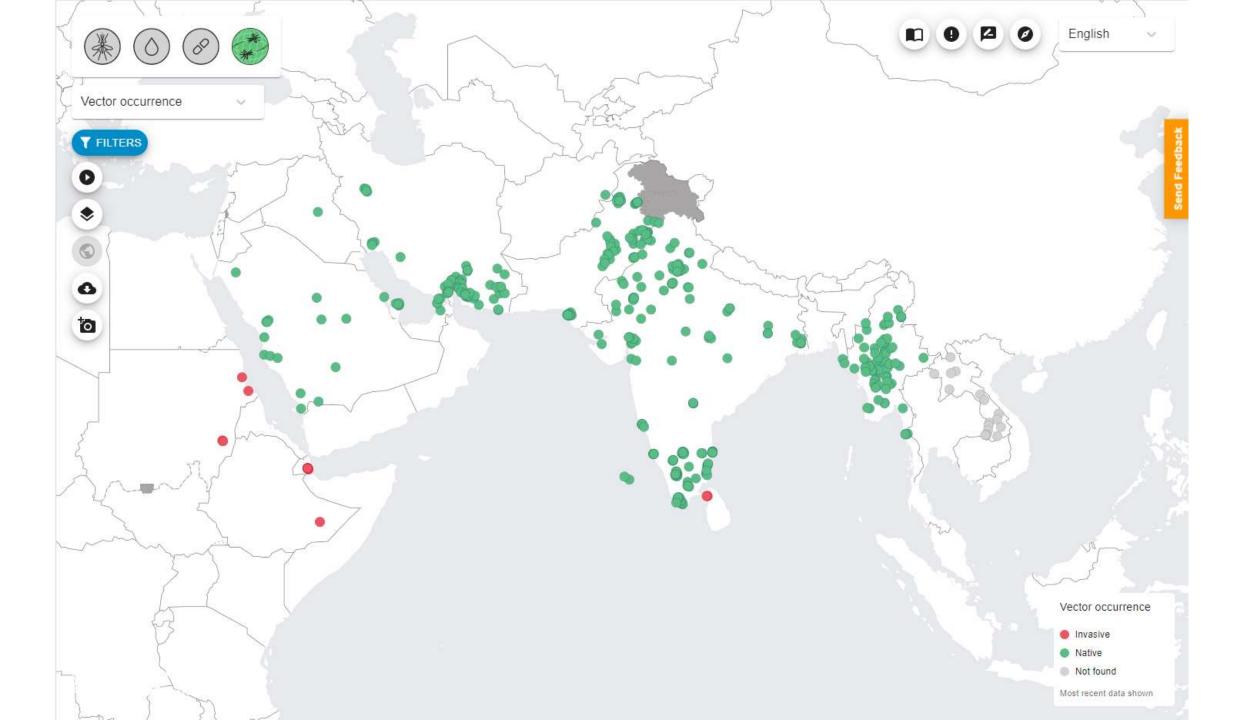


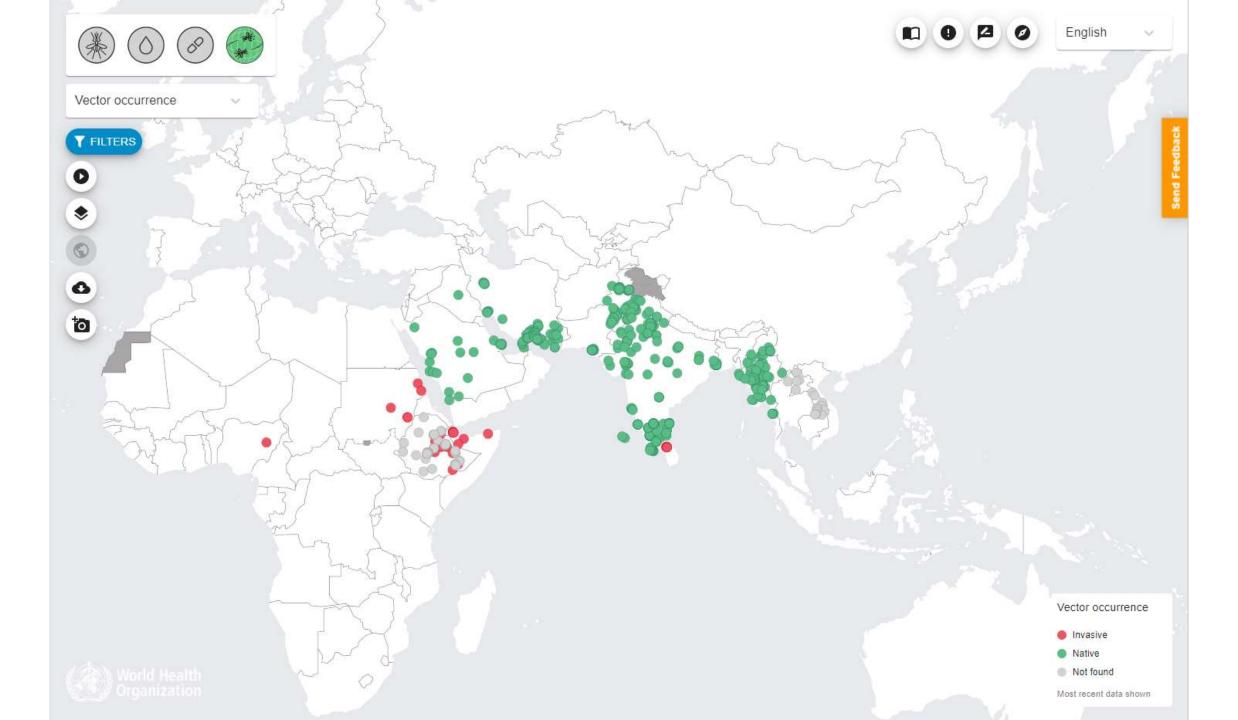


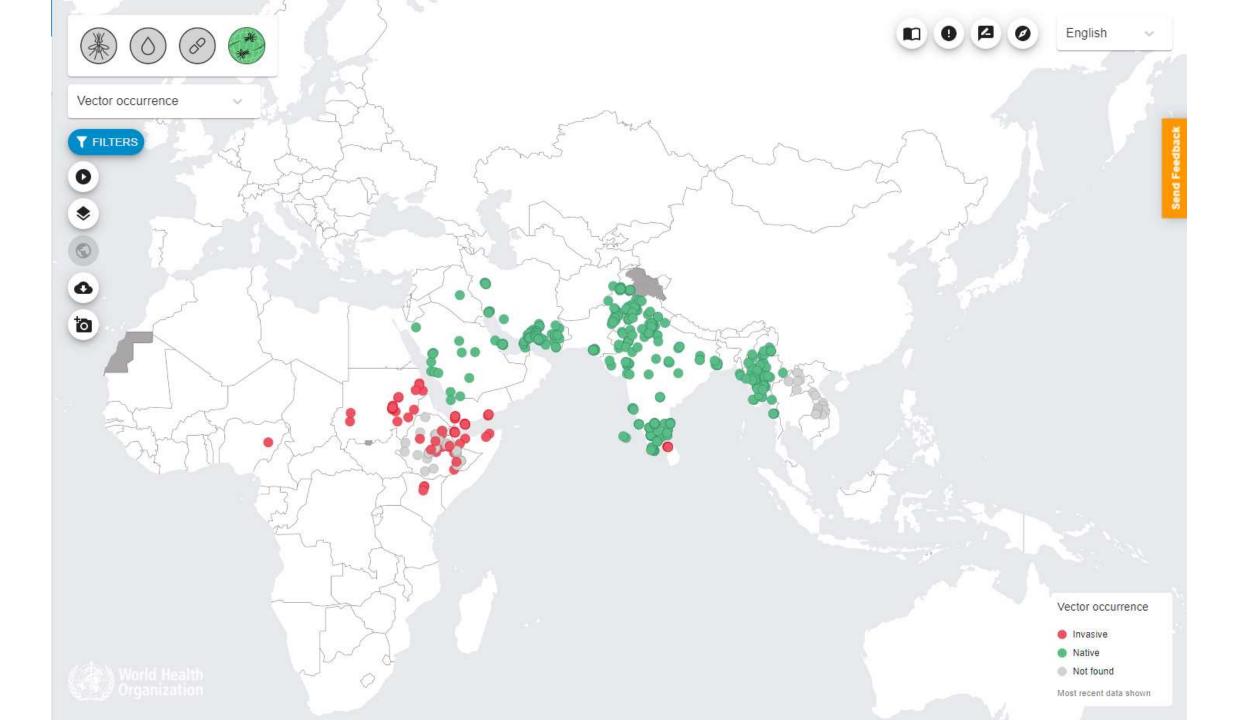


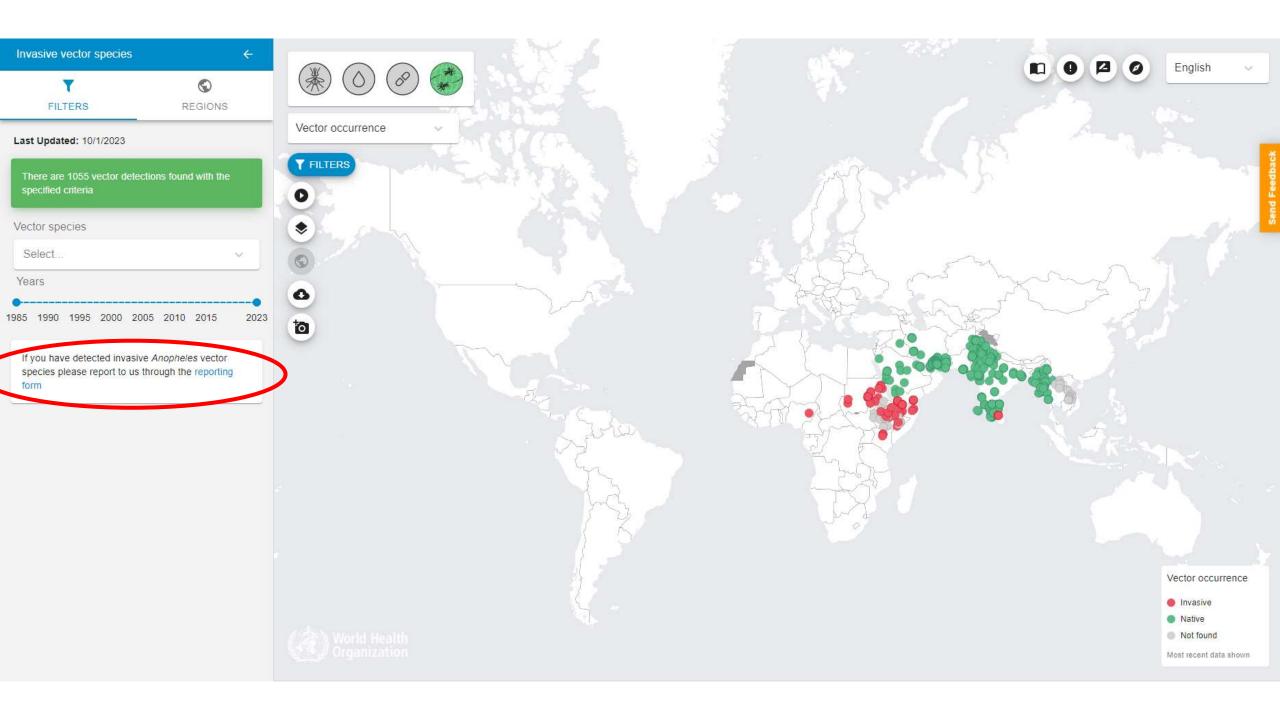


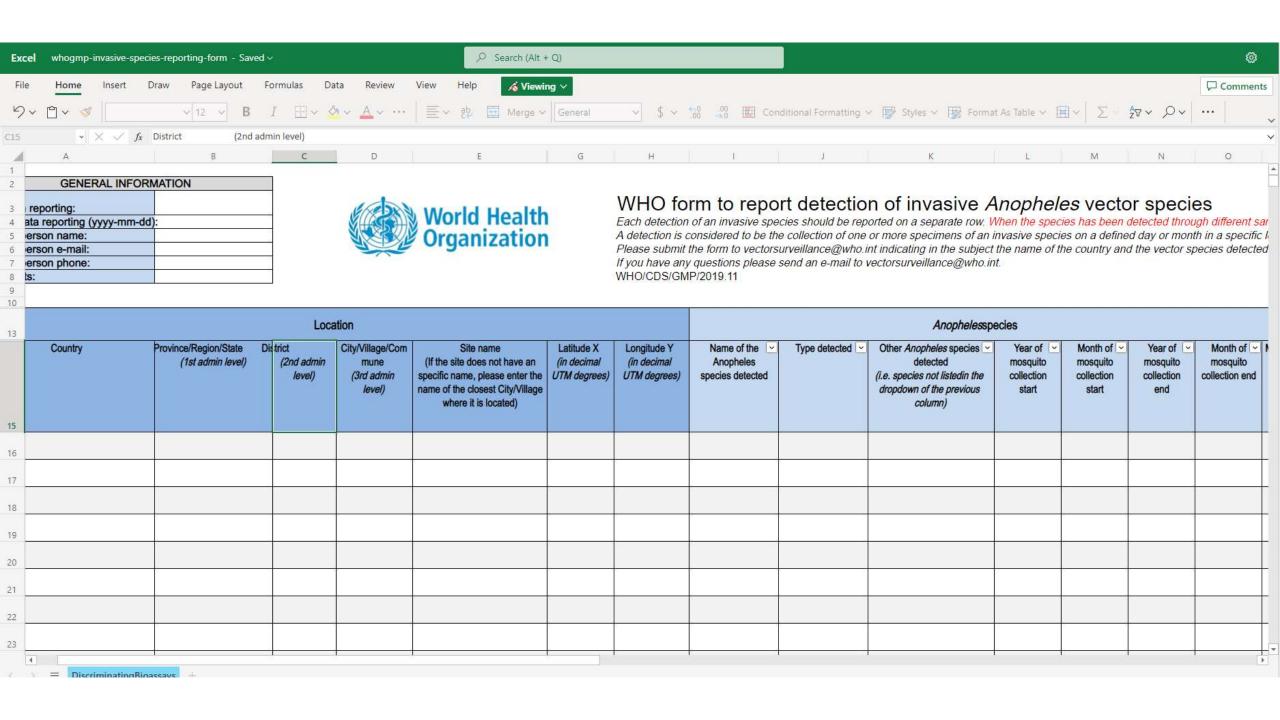












# Malaria threats map

- A start, not an end to itself
- Key questions remain:
  - Where is *An. stephensi*?
  - How is it spreading?
  - What is its role in malaria transmission?
  - How to control it?



#### RESPONDING TO THE THREAT OF ANOPHELES STEPHENSI INVASION IN AFRICA

Ethiopia's Action Plan on An. stephensi and recent updates

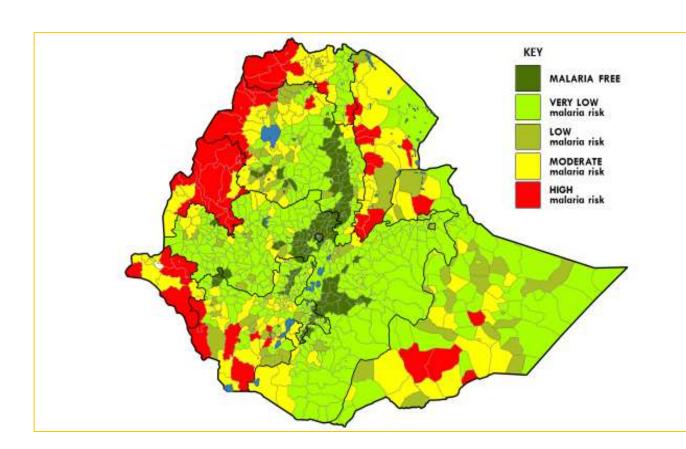
Gudissa Assefa (NMEP Manager) 21 February 2022

### Contents

- Introduction
- Planning process
- Actionable Plan (2022-2026): Summary
- Implementation arrangement
- What is next?

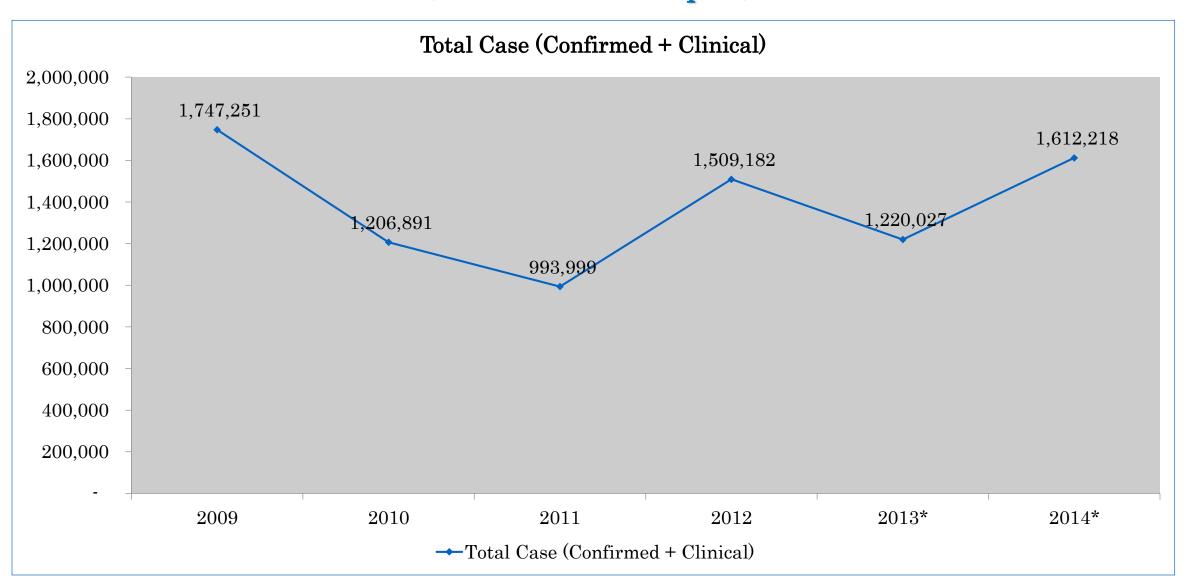
### Malaria: Burden & Epidemiology

• Malaria burden and epidemiology exhibit a marked variations in the country.

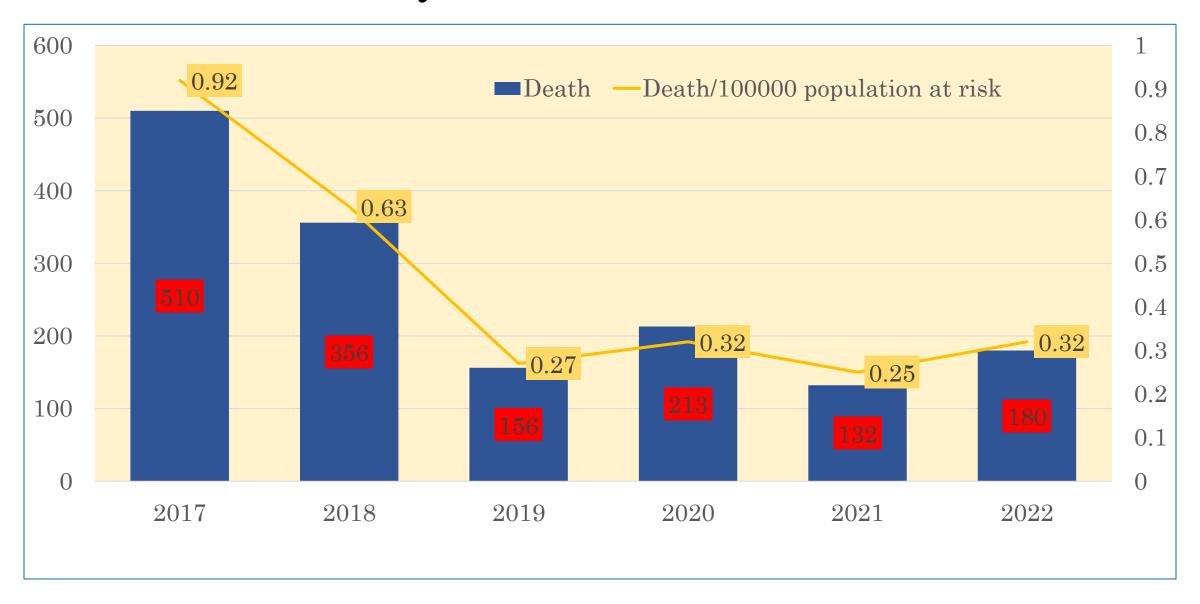


- 3/4<sup>th</sup> of the area is suitable for malaria transmission;
- Population at risk of malaria infection 52%
- **Determinants:** Altitude & climate (rainfall & temperature)
- Mainly occurs up to 2,000 masl; rare case up to 2,500 masl
- Transmission: seasonal & unstable
- Parasites: P. f (80.1%) & P. vivax (19.9%) DHIS2 (2020/21)
- Vector: An. Arabiensis & An. Stephensi

# Trend for total malaria and confirmed cases, 209-2014 EFY (DHIS-2 Disease report)

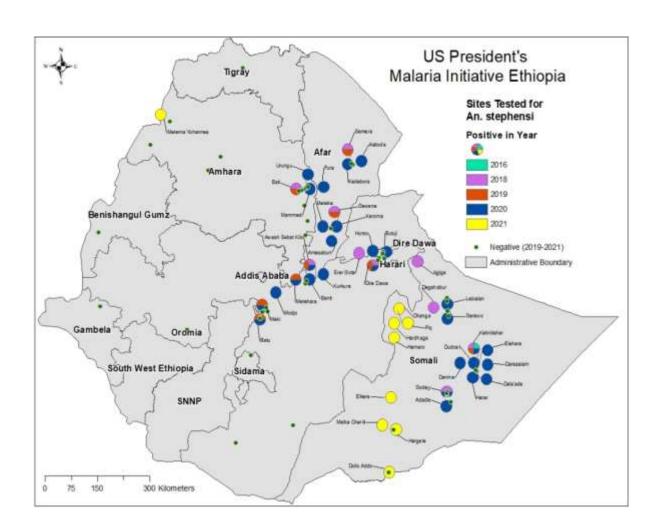


### Malaria Mortality Rate (2017-2022 DHIS 2)



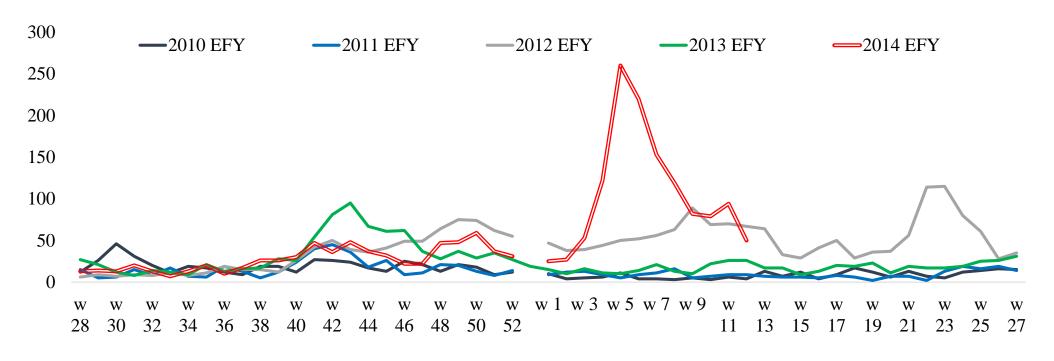
### Introduction: An. stephensi





- Detected in Ethiopia in 2016
- In Ethiopia 45 sites found positive for An. Stephensi
- Entomological surveillance
   monitoring is ongoing (CEASE
   project)

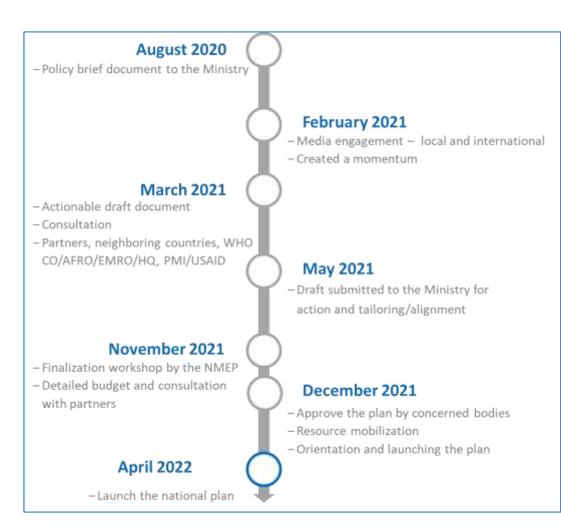
### Case Trend in Dire Dawa city, Ethiopia



Since WHO epi week 1 to 13/2022, there was <u>outbreak</u> in the city administration. Majority of cases were from Sabian location and Gore health center; and Dire Dawa University (DDU). The reason behind the outbreak are: there is larva positive **water treatment in DDU compound** and **Butuji river** which cross the city more than 9 kilometer is larva positive.



## Actionable Plan (2022-2026): Summary



#### **Planning Process**

- 1. Evidence Generation for Planning
- Evidences collected and reviewed desk review
- 2. Endorsing the evidences and Organization of Partners
- Sources for relevant evidences (PMI vector link, AHRI, EPHI, Universities and regions,....)
- Formed TWG to develop actionable plan
- Enriching the plan further and finalized
- The final plan was formally launched in the presence of all stakeholders
- Implementation is ongoing as per the plan

#### Action plan for the Integrated Surveillance and Control of Anopheles Stephensi and Aedes Aegypti,

With a targeted elimination of An. Stephensi in Ethiopia, 2022-2026

#### Main Objective:

• To eliminate An. Stephens from Ethiopia by 2026 and there by control Ae. aegypti

### Specific Objectives

- 1. To *delineate the distribution* of *An. stephensi* within Ethiopia by outlining and implementing strategies for enhanced surveillance including the evaluation and implementation of integrated surveillance with *Ae. aegypti*
- 2. To control An. stephensi and there by Ae. aegypti in the already invaded areas by implementing proven vector control measures, monitoring their impact, and evaluating potential new methods and tools
- 3. Continue control of *An. stephensi* and there by *Ae. aegypti* in areas at risk including the *use of new methods*
- 4. To establish an *Early Warning, Alert and Response System (EWARS)* to prevent An. stephensi and there by Ae. aegypti from invading any new geographical areas through the combined surveillance and control response outlined
- 5. To establish a functioning *inter-sectoral control response* by assigning clear roles and responsibilities to key partners involved in the response against An. stephensi and there by Ae. Aegypti
- 6. To provide *monitoring and evaluation approaches* to surveillance and intervention

### **Major Interventions**

#### 1.Surveillance

- a) Surveillance in areas where presence of An. stephensi is not yet confirmed
- b) Surveillance to monitor An.stephensi and *Ae. aegypti* abundance in areas where presence is confirmed
- c) Determining insecticide susceptibility of An. stephensi and Ae. aegypti

(\* already started and ongoing by some partners)

### Major Interventions

#### 2. Control of An. stephensi

- i. Interventions ready in the short-term
  - a) Environmental and Larval Source Management
  - b) Insecticide treated nets and indoor residual spraying (resistance, out door biting/resting behavior) may compromise this part of the intervention
- ii. Potential new interventions
  - a) Interventions available to be tested
  - b) Interventions being developed
- iii. Enact or introduce by-laws to regulate water storage, construction and solid waste management practices and set up system for enforcement

### LSM Launching Ceremony @ Dire Dawa



### Larva Source Management @ Batu town





### Major interventions

- 3. Cross cutting (Stakeholders, governance, coordination, and leadership of implementation)
  - Multi-sectoral engagement
  - Community Engagement
  - Human resource development (training)
  - Advocacy/Communication/SBCC
  - Monitoring and evaluation

# Cross Cutting (Partnership)



#### Budget Required for Actionable plan

Detailed Activities Budget has been set for 5 years (2022-2026)

- For surveillance = 4,965,437USD
- For Control/Intervention = 42,173,167USD
- Grand total **47,138,604 USD**
- Budget revision may be necessary in the course of time

#### What is Next?

- Resource Mobilization
- Implementation based on the endorsed plan of action with no time to waste.
- Existing potentials (Adama new training center lead by AHRI, PMI (vector link, ACIPH, PATH, S4ME, Universities...) being great potentials for the way forward
- Strong collaboration with all stakeholders

## Thank you for your Attention!

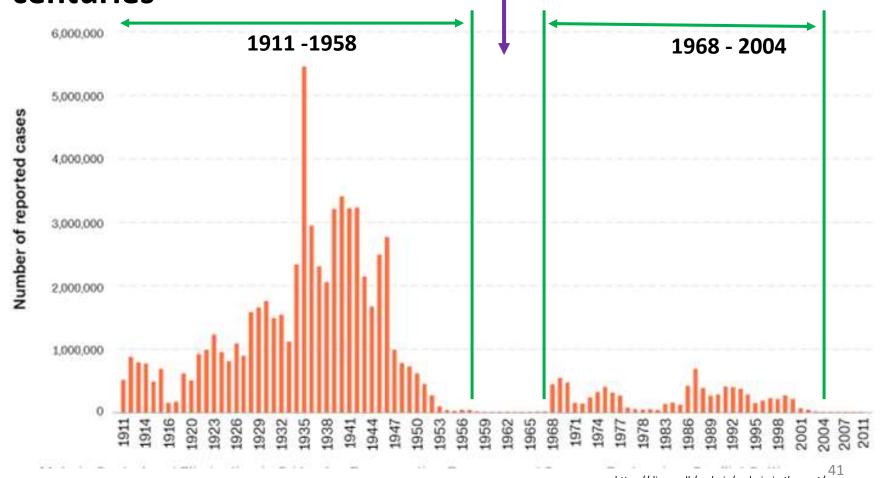
# Case Study: Recent range expansion of Anopheles stephensi to Sri Lanka and staying malaria free



S.N. Surendran Department of Zoology University of Jaffna

#### Malaria in Sri Lanka

• Sri Lanka has been endemic for rural malaria for centuries



#### Malaria in Sri Lanka......

- Due to a well-coordinated efforts of the AMC through
  - Early diagnosis
  - Treatment and
  - Vector control

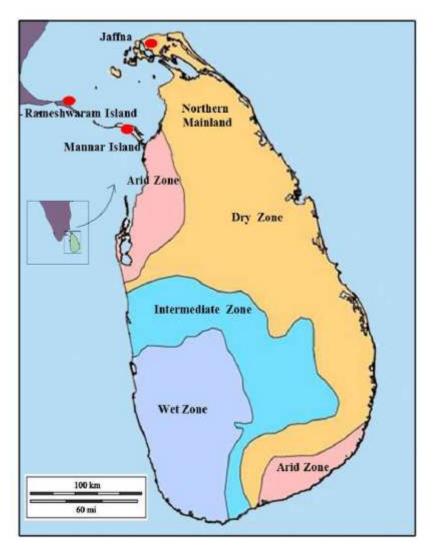
No indigenous cases reported since 2013

- Sri Lanka was declared free of any indigenous malaria in 2016
- However, imported malaria cases have been reported in the country even after 2016

Year	No. of Imported cases		
2013	95		
2014	49		
2015	36		
2016	41		
2017	56		
2018	47		
2019	53		
2021	26		
2023 (up to February)	05		

#### Malaria Vectors in Sri Lanka

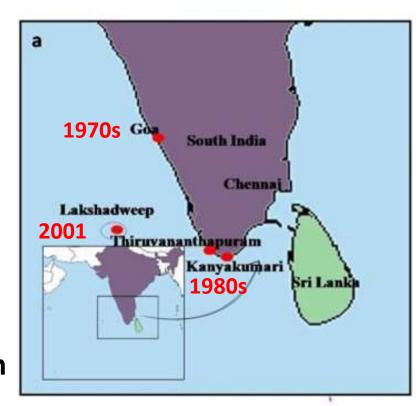
- Malaria has been endemic in the dry and intermediate climatic zones of Sri Lanka
- Out of 24 anophelines mosquitoes, Anopheles culicifacies is the major local vector
- An. subpictus is the secondary vector
- There are other anopheline species which are able to transmit malaria parasites



#### Invasive An. stephensi....

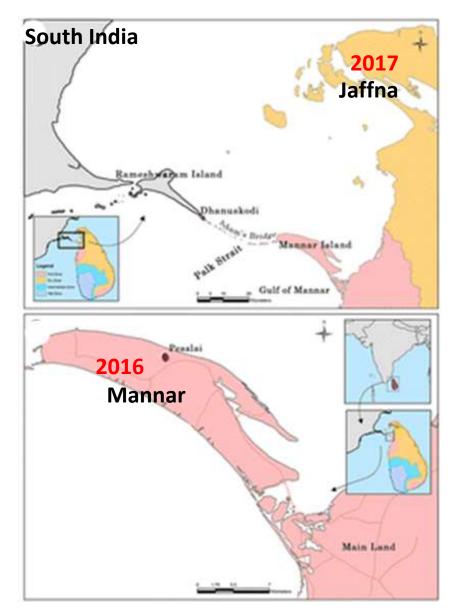
- A southward range expansion of *An. stephensi* was observed in India and attributed to urbanization and associated water storage practices.
- An. stephensi reached Goa in the 1970s
- Kanyakumari at the southernmost location of India in the 1980s, and subsequently
- Lakshadweep islands in the Arabian sea in 2001
- It was predicted in 2001 that *An.*stephensi may further expand
  southwards to invade Sri Lanka and
  the Maldives

  Suren



#### **Invasion to Sri Lanka**

- Anopheles stephensi was detected for the first time in Sri Lanka in 2016 in the island of Mannar subsequently in Jaffna in 2017.
- The adaptation of *An.*stephensi to undergo
  preimaginal development in
  domestic wells and cement
  water storage tanks has
  been considered to facilitate
  range expansion into new
  territories



#### Invasion to Sri Lanka...

#### Arrival:

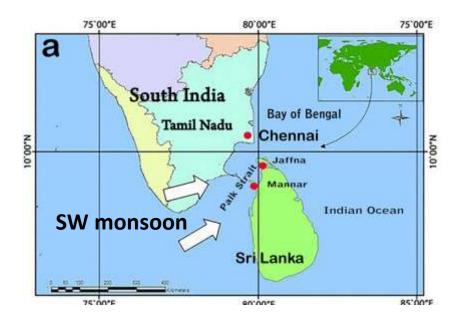
- movement of people across the 64 to 137 km-wide Palk strait that separates south India and Sri Lanka.
- Wind-borne migration due to SW monsoon???

#### • Establishment:

 the readily availability of urban domestic wells and cement water storage tanks as habitats to which it was already adapted

#### • Spread:

• it is therefore likely that An. stephensi can utilize similar anthropogenic fresh and brackish water habitats to further extend its range within the Jaffna peninsula and other coastal areas of Sri Lanka.



• Surendran et al, 2020: 30;13(1):156

#### Bionomics of *An. stephensi* in Sri Lanka

- Undergo preimaginal development in wells and cement water storage tanks
- Can lay eggs and undergo preimaginal development in brackish water of up to 3.5 g/L salt proving that *An. stephensi* has euryhaline characteristics.

<ul> <li>Highly resistant to 4% DDT, 5%</li> </ul>	
malathion and 0.05% Deltamethri	n

<ul> <li>It is therefore likely that</li> </ul>	t they can
utilize similar habitats to	o further
extend its range within	the Jaffna
peninsula and other coa	astal areas of
Sri Lanka.	Surendran

Insecticide	An. stephensi			
	% mortality	95% CI		
DDT 4%	R <sub>14</sub> ± 5 (100)	12-15		
Malathion 5%	$R_{34} \pm 9 (110)$	3-36		
Deltamethrin 0.05%	$R_{56} \pm 1 (100)$	55-57		

Surendran et al., Parasit Vectors. 2020;13(1):156.

## Bionomics of *An. stephensi* in Sri Lanka.....

- Based mode of the number of egg ridges on egg-floats type/intermediate/mysorensis forms are present in Jaffna.
- Spiracular index revealed presence of type and mysorensis
- AsteObp1 intron-1 region that identified all three biotypes in Iran and Afghanistan failed to differentiate the biotypes in Sri Lanka as in the case of Indian samples.

KJ557465.1 Type	GTGAGCTTGG	GTGTCTTCTG	GATATTGTTC	TAATGTGTTT	TCTGTCTATA	AGTTTTAAAC	ACAAA
KJ557453.1 Mysorensis			c		.TACCT	T.CTG	.G
KT587049.1 Mysorensis			c		.TACCT	T.CTG	.G
KJ557452.1 Intermediate	T	*******				TG	
KT587050.1 Intermediate	T					TG	
ASJF1				A		100.	
ASJF2	T		A			TG	
ASJF3	T		A		c	TG	
ASJF4	T					TG	
KJ557465.1 Type	TCTGGACTCT	GCATCCTAAA	GATAATGTCC	TGTCATGCAA	TGTCATCACT	TTCCAG	
KJ557453.1 Mysorensis	A.T	.AGT	TC				
KT587049.1 Mysorensis	A.T	.AG	TC				
KJ557452.1 Intermediate	**252**2**2*********	-12000000000000000000000000000000000000	T			1000 T000 T000	
Intermedade					********	*****	
ASJF1			cA				
ASJF1 ASJF2 ASJF3		.A			G		

### Bionomics of *An. stephensi* in Sri Lanka.....

- The results suggested that **numbers of egg ridges**, **spiracular indices** and **AsteObp1 intron 1** sequences were not useful for differentiating **An**. stephensi biotypes in Jaffna.
- It is proposed that the observed differences between *An. stephensi* mosquitoes in Jaffna now results from normal population variance in the context of rapidly changing bionomics in India and northern Sri Lanka

## Vector surveillance strategies of Antimalaria Campaign (AMC)

- Two categories of entomological investigations:
  - spot surveys: Spot surveys were carried out as reactive spots when a malaria patient was reported and as proactive spots when vulnerability of a certain area is increased.
  - Sentinel surveys: Sentinel monitoring has been carried out in foci where vulnerability and /or receptivity was moderate to high, on quarterly basis.
- Special entomological surveys are carried out in areas where the *Anopheles stephensi* is found.
- Special larval surveys have been conducted as pre and post intervention larval surveys in areas where invasive Anopheles stephensi has been found.

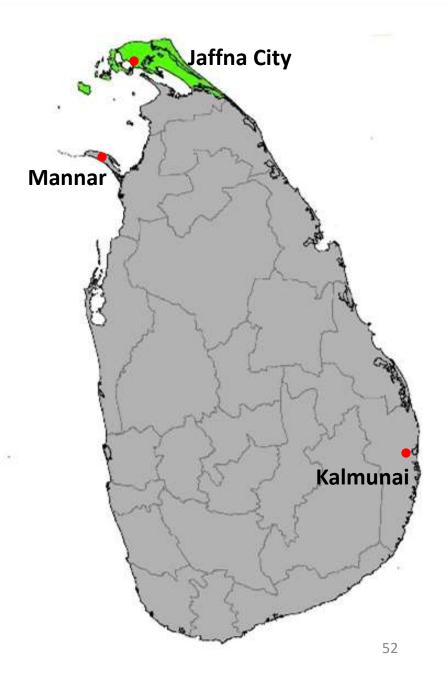
#### **Control efforts**

- The AMC continues **Reactive Spot Vector Surveys in all areas** with reported cases.
- Soon after the detection of *An. stephensi* extensive vector surveys were carried out mainly in the northern Sri Lanka to identify areas invaded by the mosquito.
- All identified preimaginal develop sites and potential sites were treated either with temephos or larvivorous fish or both and covered with mosquito proving nets wherever possible



#### **Control efforts**

- Because of the a wellcoordinated vector survey and control activities further expansion of An. stephensi to mainland of Sri Lanka has been prevented.
- Its presence is now mainly confined to three coastal areas



## Challengers in controlling *An.* stephensi

 Adaptation to undergo preimaginal development in urban environment –transmission of urban malaria

 Ability to undergo preimgainal development in brackish waters – transmission of malaria in coastal areas

 Resistant to common insecticides – difficulties in control during malaria transmission

## Implications and staying malaria free

- Posing a challenge to prevent re-emergence rural and urban malaria in Sri Lanka due to
  - Imported malaria cases and
  - Presence of local vectors along with invasive urban malaria vector *An. stephensi*
- The recent spread of *An. stephensi* to new territories are also likely to have been caused by anthropogenic and urbanization-associated factors similar to those discussed for Sri Lanka.
- Broad appreciation of the effects of anthropogenic drivers of mosquito vector-adaptation by global health decision-makers, and the development of appropriate mitigating strategies is clearly important.

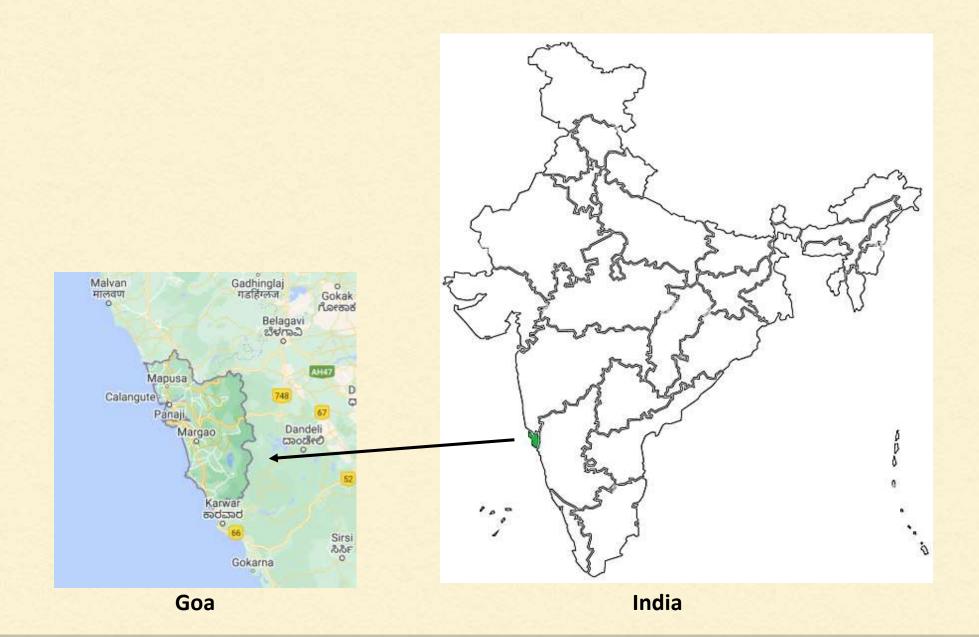
## Thank you

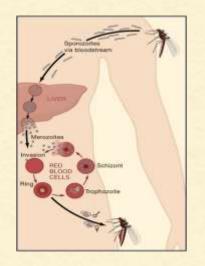


#### Anopheles stephensi Liston

#### India case study: Lessons learnt from decades fighting the vector

Ashwani Kumar Director ICMR-Vector Control Research Centre, Puducherry, India



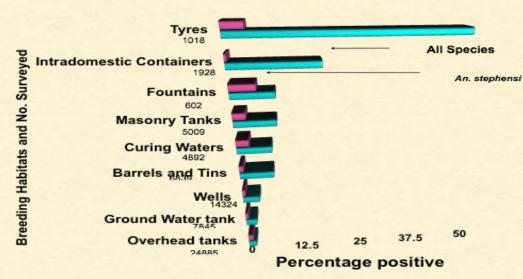


Malaria: Human Plasmodial Infections

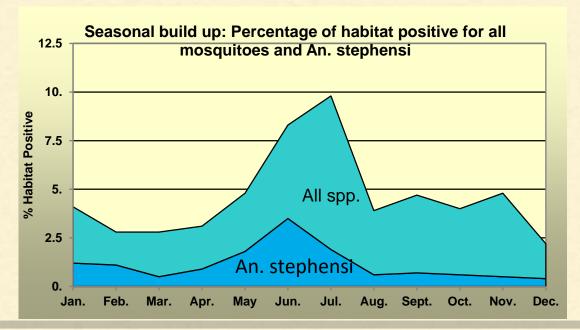
#### ANOPHELES STEPHENSI TRANSMITTED MALARIA INCIDENCE IN PANAJI, GOA, INDIA FROM 1985-1988

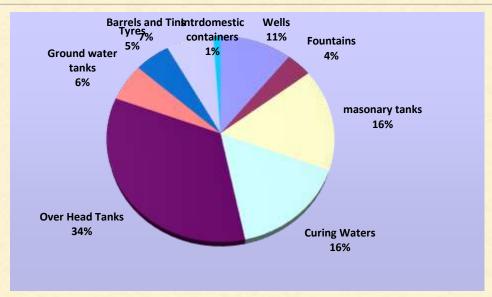
Year	BSE	Total Pos.	P. vivax	P. falciparum	Pf%	API
1985	2497	5	4	1	20	0.12
1986	6539	352	351	1	0.3	8.3
1987	21710	4416	4409	7	0.2	103.6
1988	29853	5677	5435	242	4.3	132.6

#### Malaria Problem Delineation

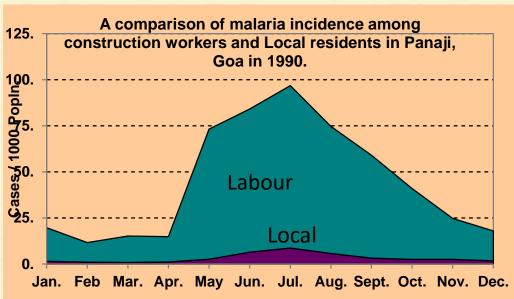


Types and number Breeding sites surveyed and their vector status

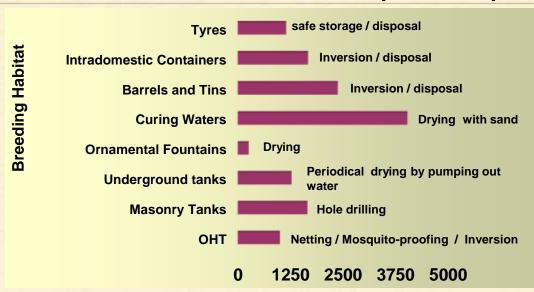


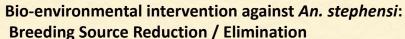


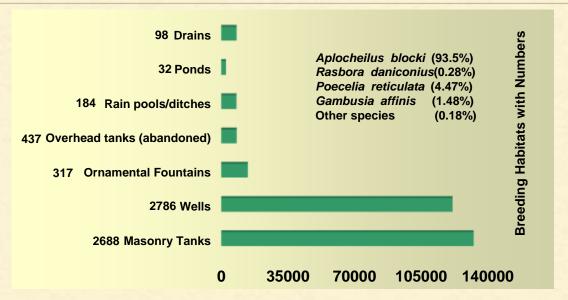
Relative contribution of various breeding habitats to *An. stephensi* breeding in 1990



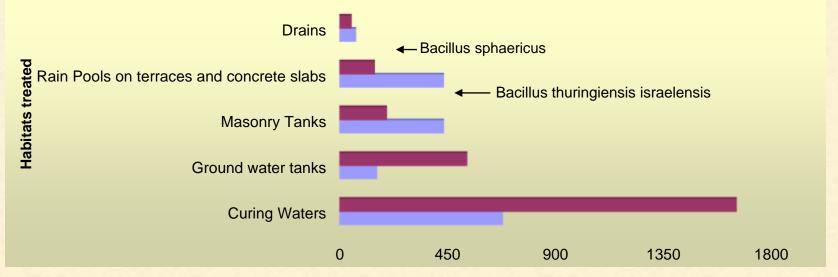
#### **Anopheles stephensi** Control Interventions







Biological control utilizing larvivorous fishes in Panaji, Goa, India (1990-92)



Biological Control of An. stephensi with biolarvicides in Panaji from 1990-92



#### Mosquito-Proofing















Construction sites

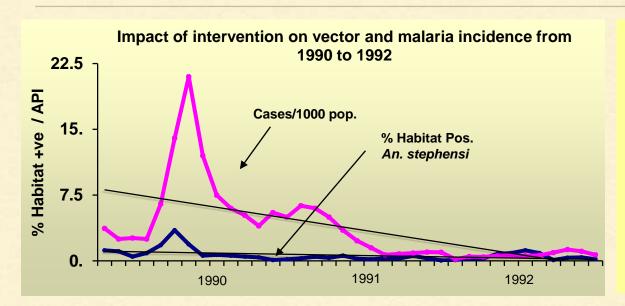


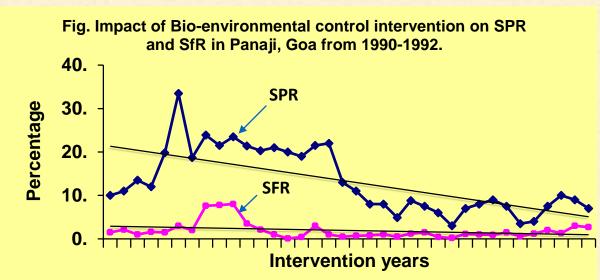
Unlined surface drains

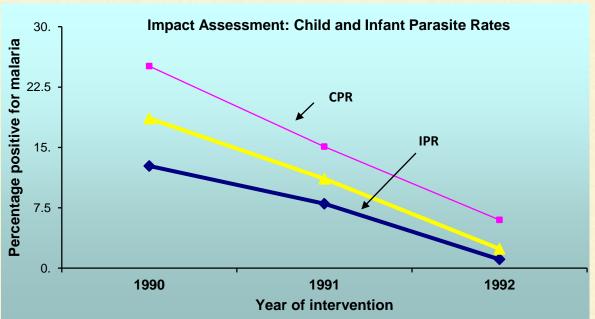


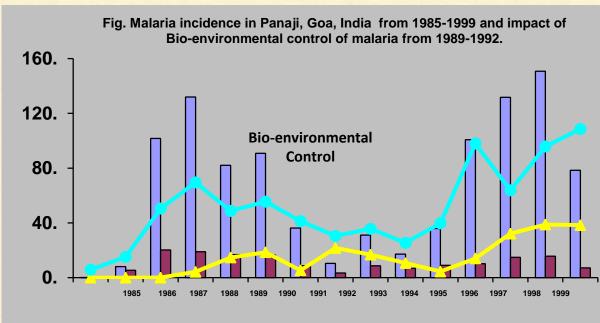


Goa Port: Anopheles stephensi Breeding Habitats and Interventions









Challenge with Adult *Anopheles stephensi* Resting Collection by Hand Catch: Sumodan et al, 2004 JAMCA

Collected 38 mosquitoes resting on 15 different surfaces

#### CDC Light Trap Deployment, Mosquito collections and Identification: More Effective for Collections/Vector Surveillance











## Anopheles stephensi: a catalyst for multi-sectoral action

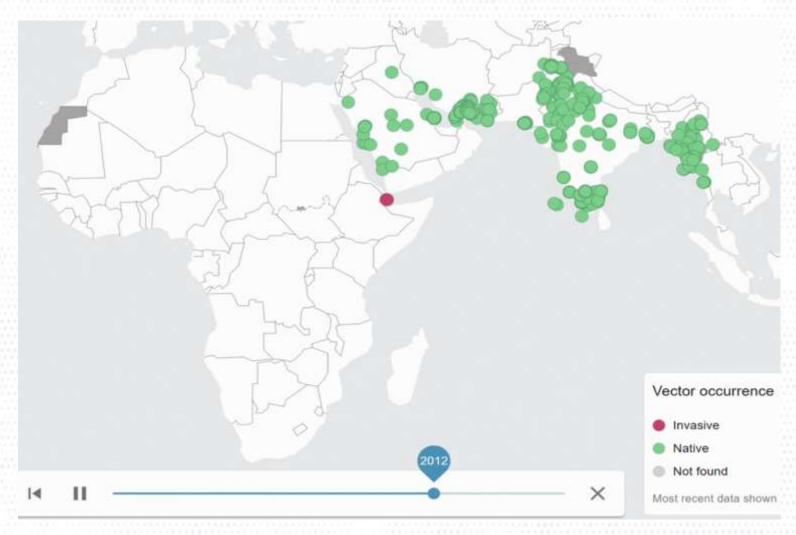
#### OUTLINE

- Unique vector
- What is PMI doing?
- Multi-sectoral opportunities
- Indian Ocean example
- Global engagement and advances in public health entomology



#### An. stephensi IS STILL SPREADING IN AFRICA

- Djibouti (2012)
- Ethiopia (2016)
- Sudan (2016)
- Somalia (2019)
- Nigeria (2020)
- Kenya (2022)



#### A UNIQUE VECTOR CREATES CHALLENGES OPPORTUNITIES

- Urban and rural adapted
- Thrives in artificial habitats
  - Shared with dengue mosquito, Aedes aegypti
  - Water storage for personal use, agriculture, construction, etc.
- Persists through dry periods
- Often collected in proximity to livestock
- Transmits-Plasmodium falciparum and P.

  □ ফাফের্মেণ্ট্রাইউভিউউভি শ্রাইউভিউউভি শ্রাইউভিউভিউভি শ্রাইউভিউভিউভি শ্রাইউভিউভিউভি শ্রাইউভিউভি শ্রাইউভিউভিউভি শ্রাইউভিউভিউভি শ্রাইউভিউভি শ্রাইউভিউভিউভি শ্রাইউভিউভিউভি শ্রাইউভিউভি শ্রাইউভিউভিউভি শ্রাইউভিউভি শ্রাইউভি শ





#### WHAT IS PMI DOING?

Alignment with WHO initiative











- PMI An. stephensi interagency Task Force formed
- USG coordination across 7 agencies
- PMI LSM policy revised for rapid response
- Action plan developed, shared w/ 27 countries, will be online
- Leverage PMI partnerships-12 PMI countries began activities
- Supported modeling study on potential impact in Ethiopia
- U. Surveillance Agantral needs have identified new partners to engage with surveillance and response capacity



#### PMI & CDC GLOBAL COORDINATION

- WHO urban malaria technical advisory group
- WHO An. stephensi initiative
- WHO EMRO/AFRO
- Roll Back Malaria-Vector Control Working Group
- Global Fund
- Indian Ocean Commission
- Academic partnerships
- West African Aedes Surveillance Network
- Pan African Mosquito Control Association



#### WHAT IS CDC DOING?

- WAASuN trained Aedes programs in 18 countries
- PAMCA trainings and response in Djibouti
- Provide support to non-PMI partner countries
- Genomics-identified source populations and reintroductions (Carter et al. 2021)
- Collaboration within CDC- One Health, border health, FETP, geospatial training support opportunities
- CDC BAA call for An. stephensi population genomics and control innovation



# PMI Action Plan 2.0

#### **Approach**

 Mitigation of impacts of An. stephensi utilizing enhanced vector and disease surveillance, coordinated intervention implementation, and improved monitoring

#### Geography

- Scenario 1 An. stephensi is established
- Scenario 2 At risk of invasion

#### Activities that can be conducted immediately

- Investigate anomalous Anopheles spp. from routine collections and IRM
- Align with Aedes surveillance
- Strengthen urban and port vector surveillance activities
- Habitat suitability maps provided to guide efforts

#### **Guidance and activities proposed**

- Surveillance, monitoring and evaluation
- Vector control
- Social and behavior change (SBC) guidance developed
- Multisectoral coordination-One Health, WASH, trade/commerce, <u>population mobility</u>
- Regulatory needs
- Community engagement

PMI Action Plan to respond to the threat of Anopheles stephensi in Africa



Last more lawying (A. 2023)



### PMI ETHIOPIA

- Invasive An. stephensi entomological surveillance and monitoring
   Insecticide resistance trends

  - 48 detection sites
  - Sporozoite and bloodmeal analysis
- Training support for: Kenya, Benin, Nigeria, Yemen, Sudan, Somalia, Djibouti
- Control launched in eight towns beginning Aug 2022
- Support to investigation of 2022 malaria Dire Dawa outbreak led by AHRI, NMEP
- Support to NMEP and national action plan for An. stephensi elimination





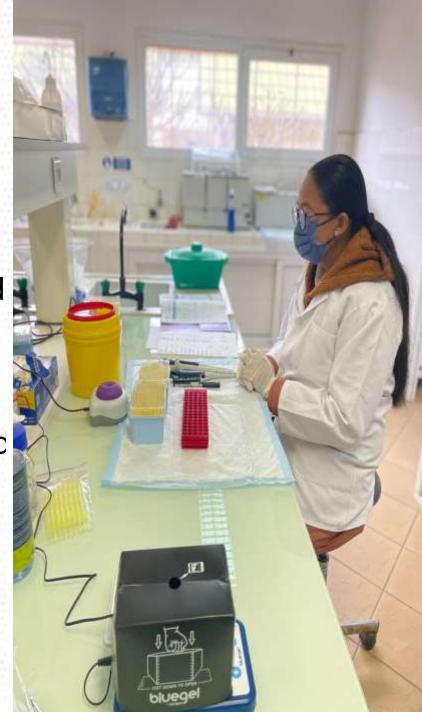
### **OPPORTUNITIES**

- Leveraging enhanced surveillance and control monitoring
- Cross-border coordination and collaboration to accelerate response and learning opportunities
- Coordination with non-malaria programs to fill gaps
- Accelerate GVCR and preparedness and response for mosquito borne diseases
- Urban mosquito borne disease surveillance and control
- Enhance public health entomology workforce



#### SURVEILLANCE

- Improved scientific quality of routine entomological data by reporting and investigating anomalies
- Leverage sequencing capacity
- Negative detections are beneficial for tracking spread
- Understanding urban vector dynamics when not An. stephensi
- GVCR accelerated by leveraging Aedes/arboviral and Anopheles/malaria programs
  - Aedes data in África enhanced
- WHO urban malaria framework alignment



# **INVASIVE SPECIES**

- Opportunities to leverage invasive species efforts and consortia
- US National Invasive Species Council (NISC)
- Biosurveillance, trade, economics, regulations, containment, elimination thresholds
- Invasive mosquitoes often thrive in containers and approaches can be integrated
- Cross-border training to identify species outside of typical keys
- Species distribution models to predict likelihood of introduction/establishment



# TRADE/COMMERCE

 2011 maritime trade data identified **Djibouti** and **Sudan** as countries at greatest risk of introduction (Ahn et al. 2023)

• 2020 marine trade (introduction) data combined



## POPULATION MOBILITY

- Identify points of entry/points for control
- Leverage existing data on pinch points from communicable disease (Medley e al. 2021)
- PopCAB Toolkit in PMI Action Plan



#### Legend



# INDIAN OCEAN EXAMPLE: Anopheles and Aedes

Mauritius connects Asia and Africa through maritime trade

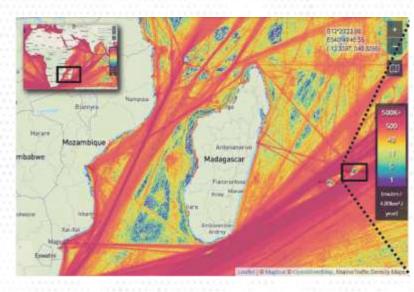
- Strong Aedes surveillance program leveraged (Diana Iyaloo)
  - An. stephensi surveillance reveals Anopheles diversity
- Indian Ocean Commission
- In 1 yr five island nations are coordinating vector programs
- Port surveillance in coordination with Port Authorities

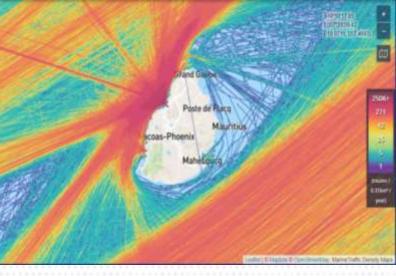












### WHAT HAVE WE LEARNED?

- Strengths in global partnerships, evidence-informed decision making, and localization initiatives when combined with efforts outside of malaria can accelerate the rate of action
- Cross-cutting engagement can allow for progress in parallel
- Multisectoral engagement can advance initiatives and activities outside of organizational scope
- Gaps and competing resources limit existing surveillance, research, and response capacity
- It is critical to drive more action and address questions around a future with *An. stephensi*



# **THANK YOU!**

#### MESA FORUM | 21 FEBRUARY 2023, 2 PM CET

# RESPONDING TO THE THREAT OF ANOPHELES STEPHENSI INVASION IN AFRICA



#### **Pre- Forum Questions**

- 1. What is the threat of *An. stephensi* in other regions of Africa, what is the basic step for proximal countries to take and why are most countries more interested with detecting it rather than starting to put strategies to prevent its introduction?
- 2. Is there any finding that indicates the current malaria transmission by *An. stephensi* and what molecular tools are being used for detection in developed urban areas?
- 3. Is there any change in the vector control interventions since the detection of An. stephensi?
- 4. There are parallels with *Aedes aegypt*i control and behaviour. What have we learnt from that species?
- 5. What options are there for action by individuals and organisations in response to the *An. stephensi* rising danger to global eradication?