

3 February, 2020

Results from community level evaluation of Imergard wettable powder (WP) a new potential molecule for IRS.

Sarah J Moore

Ifakara Health Institute and Swiss Tropical and Public Health Institute

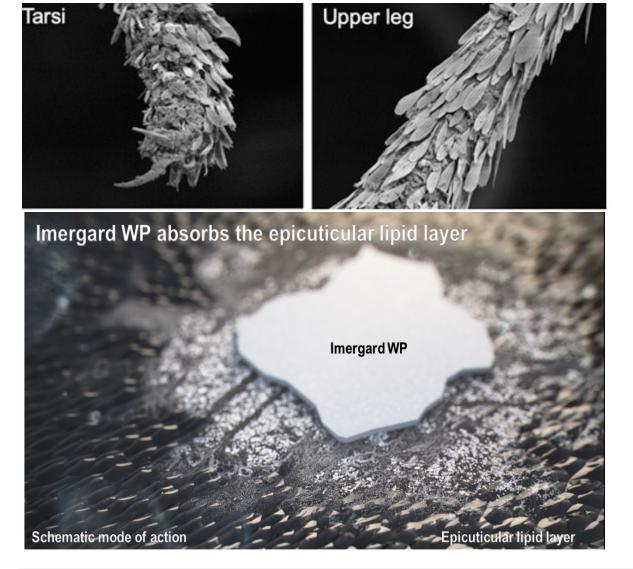
RBM Partnership To End Malaria

If IRS is to remain an important element in an integrated vector control approach for sub-Saharan Africa, alternative insecticides, particularly ones with novel modes of action (MOA) are urgently needed

RBM Partnership To End Malaria Imergard WP 3 February, 2020

What is a Mechanical Insecticide?

Industrial MINERAL particles that produce a mortal response on contact



Imergard WP is 100% perlite

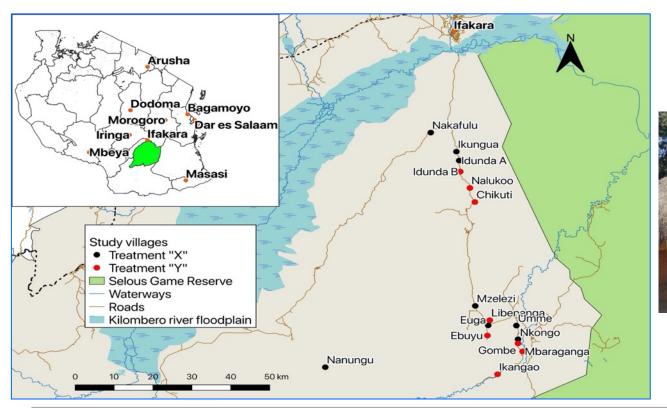
RBM Partnership To End Malaria Imergard WP 3 February, 2020

Trial design

Single-blinded 2 arm randomized community level evaluation

- 1) Imergard™ WP applied at 6 g ai/m2; and
- 2) Positive control: Actellic 300CS applied at 1 g ai/m2.

All households had Olyset LNs from universal coverage campaign.





RBM Partnership To End Malaria Imergard WP 3 February, 2020

Outcome measures

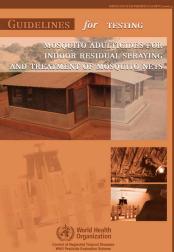
Sporozoite rate: the proportion of sporozoite positive mosquitoes (primary endpoint)

Mosquito density: nightly mosquitoes trapped by CDC light trap, human landing catch

Residual efficacy: the proportions of mosquitoes exposed in cone bioassay and held for up to 72 h with access to sugar

solution after 48 hours

Operational feasibility and safety











| Sampling method | Outcome | | Replicates per house | Frequency of sampling | Total sampling effort |
|------------------------------------|--|----|-------------------------|--|-----------------------------|
| Filter paper samples / gravimetric | Spray quality | 5 | 4 | 1 | 140 |
| WHO cone bioassays | Residual efficacy of IRS | 4 | 5 | 7 days post spray + monthly for 8 months | 2800 |
| WHO tube assays | Insecticide susceptibility of target species | 1 | 1 | baseline + 8 months post spray in 4 clusters per arm | 16 |
| Indoor Prokopack collections | Vector mortality | 2 | 1 | monthly for 8 months | 256 |
| CDC LT, HDT | Vector density Sporozoite rate (ELISA) | 80 | 1 | 4 houses each night, 20 nights a month for 8 months | 6,720 |
| HLC | Vector density / biting rate Sporozoite rate (ELISA) | 3 | 1 | monthly for 8 months | 236 |

| Factor | Imergard | Actellic 300CS | p value |
|---------------------------|------------------|------------------|---------|
| Number of participants | 6418 | 7122 | |
| Number of households | 1819 | 1991 | |
| Average Cluster Size | 259.86 | 284.43 | |
| Household Size | 3.53 (3.44-3.61) | 3.58 (3.49-3.66) | 0.4274 |
| | | | |
| <5 | 0.49 (0.46-0.52) | 0.52 (0.49-0.55) | 0.114 |
| 5 to 18 | 1.17 (1.11-1.23) | 1.19 (1.13-1.25) | 0.6314 |
| >18 | 1.87 (1.83-1.91) | 1.86 (1.83-1.90) | 0.821 |
| Pregnant Women | 0.04 (0.03-0.05) | 0.03 (0.02-0.04) | 0.1667 |
| | | | |
| Walls: Mud | 738 (40.57) | 842 (42.29) | 0.282 |
| Walls: Bricks | 1081 (59.43) | 1149 (57.71) | |
| Roof: Grass/Leaves/Palms | 906 (49.81) | 1025 (51.48) | 0.302 |
| Roof: Iron Sheets | 913 (50.19) | 966 (48.52) | |
| Floor: Mud | 1568 (86.20) | 1757 (88.25) | 0.058 |
| Floor: Cement | 251 (13.80) | 234 (11.75) | |
| Eaves: Open | 1072 (58.93) | 1191 (59.82) | 0.578 |
| Eaves: Closed | 747 (41.07) | 800 (40.18) | |
| | | | |
| Number of Sleeping spaces | 1.82 (1.78-1.86) | 1.79 (1.75-1.83) | 0.3057 |
| Household density | 2.04 (1.99-2.08) | 2.09 (2.04-2.14) | 0.0959 |
| Nets per household | 1.75 (1.70-1.80) | 1.76 (1.71-1.80) | 0.8101 |
| Net coverage | 0.55 (0.54-0.56) | 0.56 (0.54-0.57) | 0.5351 |



Results summary

Trial was conducted at acceptable quality

- No imbalance between treatment arms
- Spray quality within 50% of target dose measured gravimetrically and by filter paper
- The study was powered to detect a difference in sporozoite rate with 51,200 mosquitoes collected per arm. This was reached in the Imergard arm but not in the Actellic arm

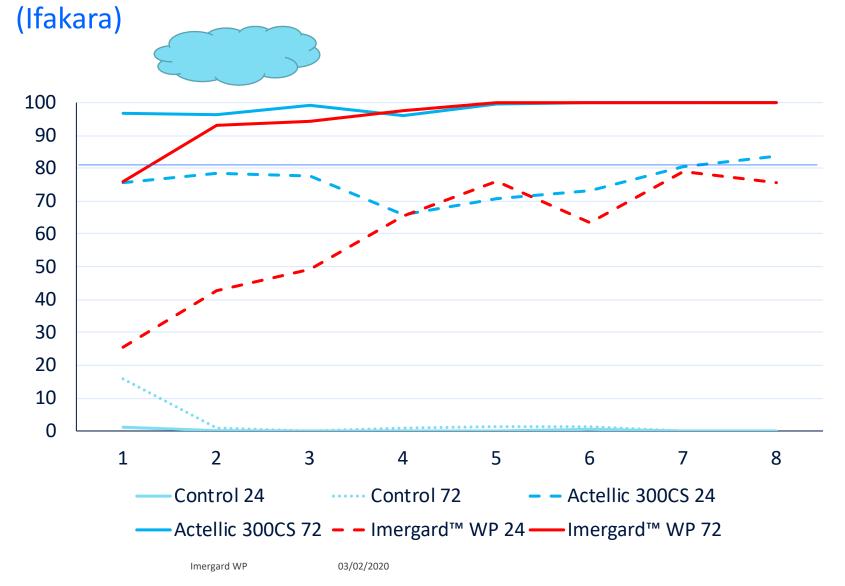
Resistance

An. funestus and An. arabiensis remained susceptible to pirimiphos methyl (100% and 98% mortality).

Safety and acceptability

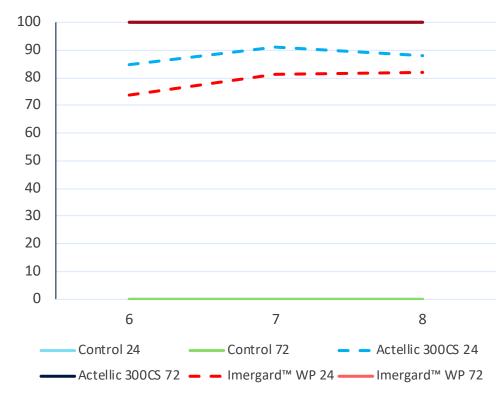
- There were no serious adverse events in either treatment arm among sprayers or residents.
- Both treatment arms were well received by local residents

Control corrected mortality for 8 months after spraying for Actellic and Imergaard with 24 hour and 72 hour holding times measured by cone bioassay with a susceptible *Anopheles gambiae* s.s.



Control corrected mortality for 6-8 months after spraying for Actellic and Imergaard with 24 hour and 72 hour holding times measured by cone bioassay with a resistant *Anopheles arabiensis* (Ifakara)





Residual efficacy

(≥80% mortality) of Imergard an Actellic with *An. gambiae* Ifakara was not met at 24 hours, however, with 72-hours holding time, residual efficacy was 8 months with susceptible and resistant strains.

Imergard WP 03/02/2020 12

SPOROZOITE RATE – not different

| | Actellic 300CS | | | Imergard | | | Odds Ratio with Actellic 300CS as the reference | | |
|--------------|----------------|---------|--------------------|----------|---------|--------------------|---|------------------------|---------|
| Species | Positive | Sampled | Sporozoite Rate | Positive | Sampled | Sporozoite Rate | % difference | Odds Ratio (95% CI) | p value |
| An. | | | | | | | | | |
| arabiensis | 5 | 4,258 | 0.00117 | 12 | 4,226 | 0.00284 | 58.64 | 4.14 (0.38-45.77) | 0.246 |
| An. funestus | 66 | 20,259 | 0.00326 | 274 | 56,556 | 0.00484 | 32.75 | 1.72 (0.70-4.26) | 0.233 |
| Overall | 71 | 24,517 | 0.00290 | 286 | 60,782 | 0.00471 | 38.45 | 2.04 (0.83-5.01) | 0.118 |

Primary endpoint:

proportion of malaria infected mosquitoes (sporozoite rate) measured in 85,299 samples was not statistically different between the treatment arms for both *An. funestus* and *An. arabiensis*

Overall sporozoite rate was not significantly different between Actellic and Imergard treatment arms (0.00290 vs 0.00471; 2.04 OR (95% CI 0.83-5.01); p=0.118).

ENTOMOLOGICAL INNOCULATION RATE

Mosquito densities different so EIR is higher in Imergard arm Mosquito densities not different between arms adjusted for cluster

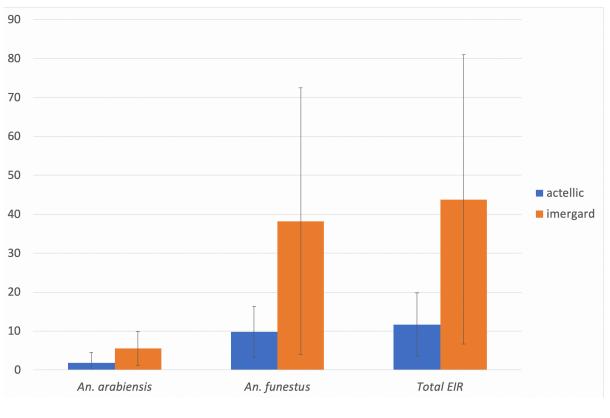
| | An. ard | abiensis | An. funestus | | |
|--|-------------------|----------|-------------------|----------|--|
| | Actellic 300CS | Imergard | Actellic 300CS | Imergard | |
| Total Number Mosquitoes collected by HLC | 846 | 1064 | 2205 | 6046 | |
| Total Number of HLC Nights | 236 | 240 | 236 | 240 | |
| Average Landing Rate per night by HLC | 4.3 | 4.1 | 6.7 | 14.2 | |
| Total number of mosquitoes analysed for Pf CSP* | 4,258 | 4,226 | 20,259 | 56,556 | |
| Total number of sporozoite positive mosquitoes | 5 | 12 | 66 | 274 | |
| Sporozoite Rate | 0.00117 | 0.00283 | 0.00325 | 0.00484 | |
| Proportion of all sporozoite positive mosquitoes | 1.40 | 3.36 | 18.49 | 76.75 | |
| Annual EIR | 1.84 | 4.25 | 7.93 | 25.11 | |
| % EIR Contribution | 4.71 | 10.86 | 20.27 | 64.17 | |

ENTOMOLOGICAL INNOCULATION RATE

Secondary endpoint

An. funestus densities were higher in the Imergard arm but not statistically different between the two arms (adjusted by cluster).

The mean EIR by arm for *An. arabiensis* was 1.8 and 5.5 and for *An. funestus* was 9.8 and 38.2 in the Actellic and Imergard treatment arms, respectively but not statistically different between the two arms (adjusted by cluster).



Imergard WP 3 February, 2020

15



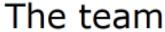
Amanda Ross

Swiss TPH

- Jason Moore
- Emanuel Mbuba
- Adam Saddler



- Carly Marshal
- Hassan Ngonyani
- Rose Philipo
- Olukayode Odufuwa
- David Kaftan
- Kyeba Swai

































Imergard WP 03/02/2020 16