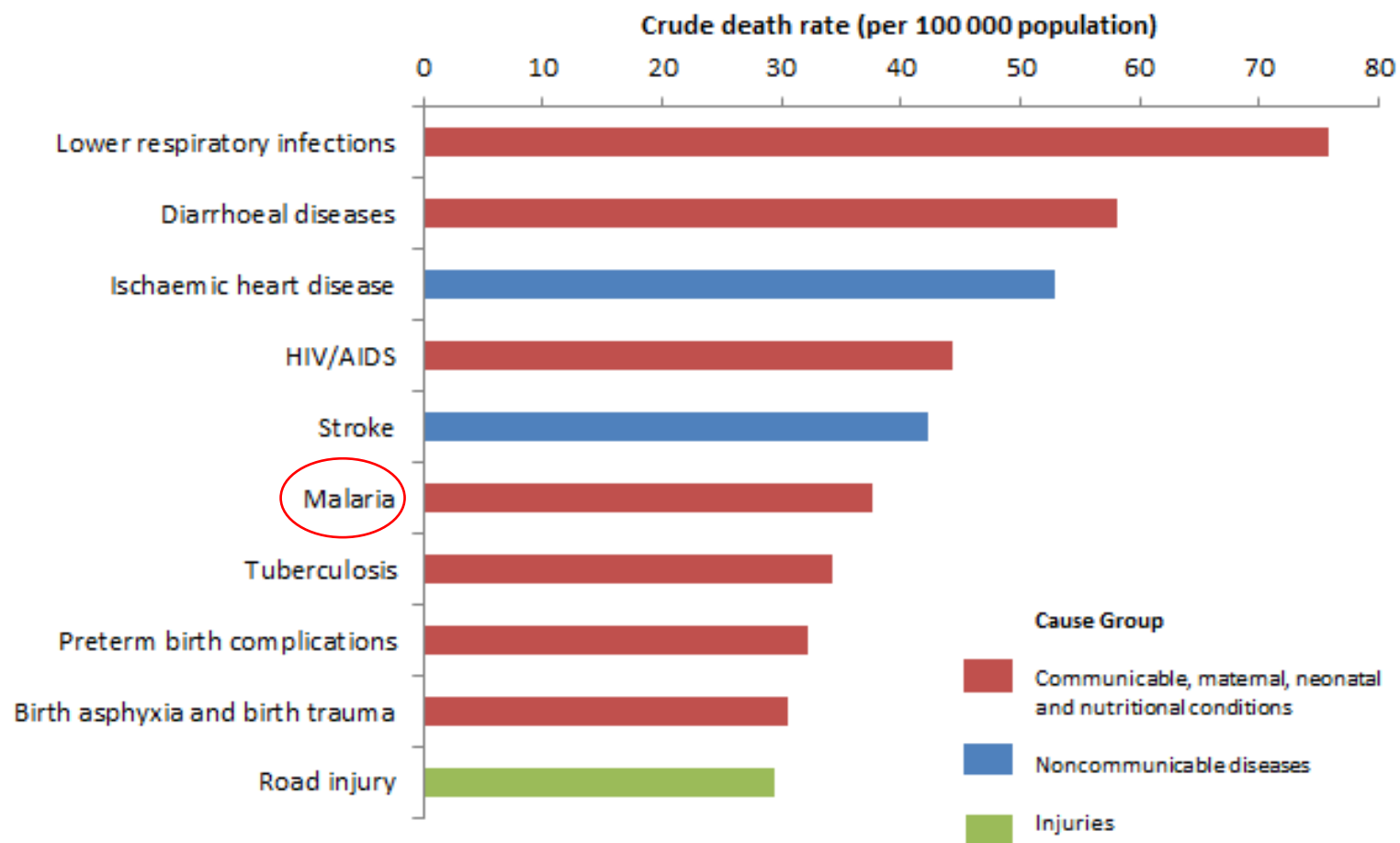


## Top 10 causes of deaths in low-income countries in 2016



Source: Global Health Estimates 2016: Deaths by Cause, Age, Sex, by Country and by Region, 2000-2016. Geneva, World Health Organization; 2018.  
World Bank list of economies (June 2017). Washington, DC: The World Bank Group; 2017 (<https://datahelpdesk.worldbank.org/knowledgebase/articles/906519-world-bank-country-and-lending-groups>).

# LA MALARIA IN NUMERI

**400 mila**

Le vittime ogni anno

**2/3**

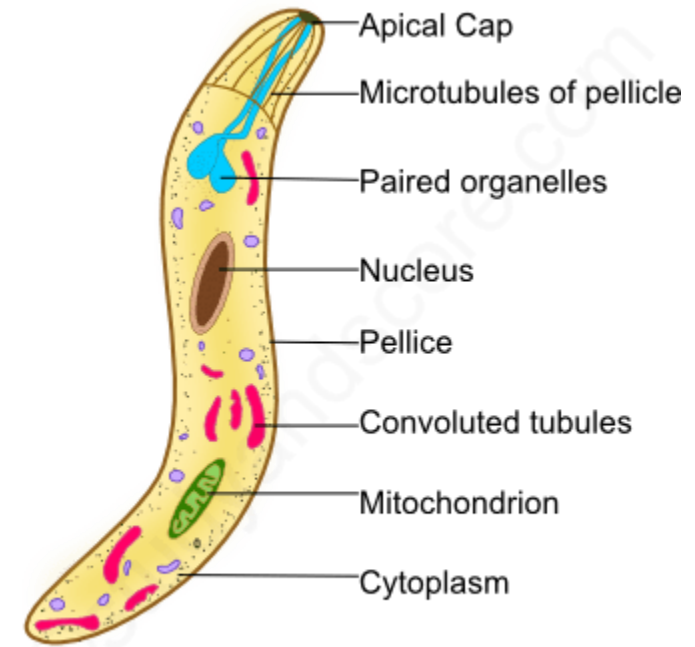
Le vittime di età  
inferiore a 5 anni

**94%**

Dei contagi e dei morti  
si registra in Africa

**87**

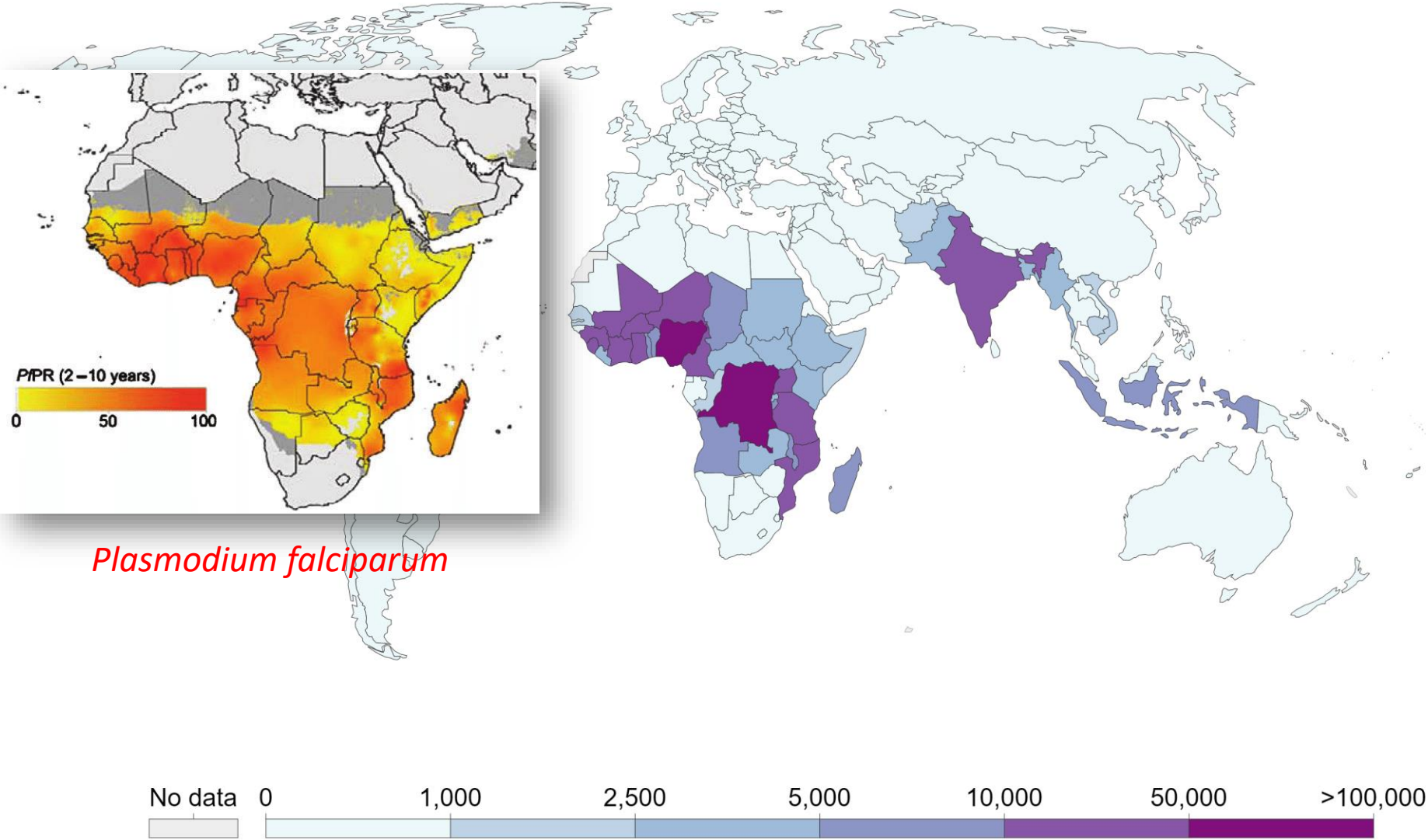
I paesi in cui è  
considerata endemica



FONTE:  
WHO Malaria Report 2020

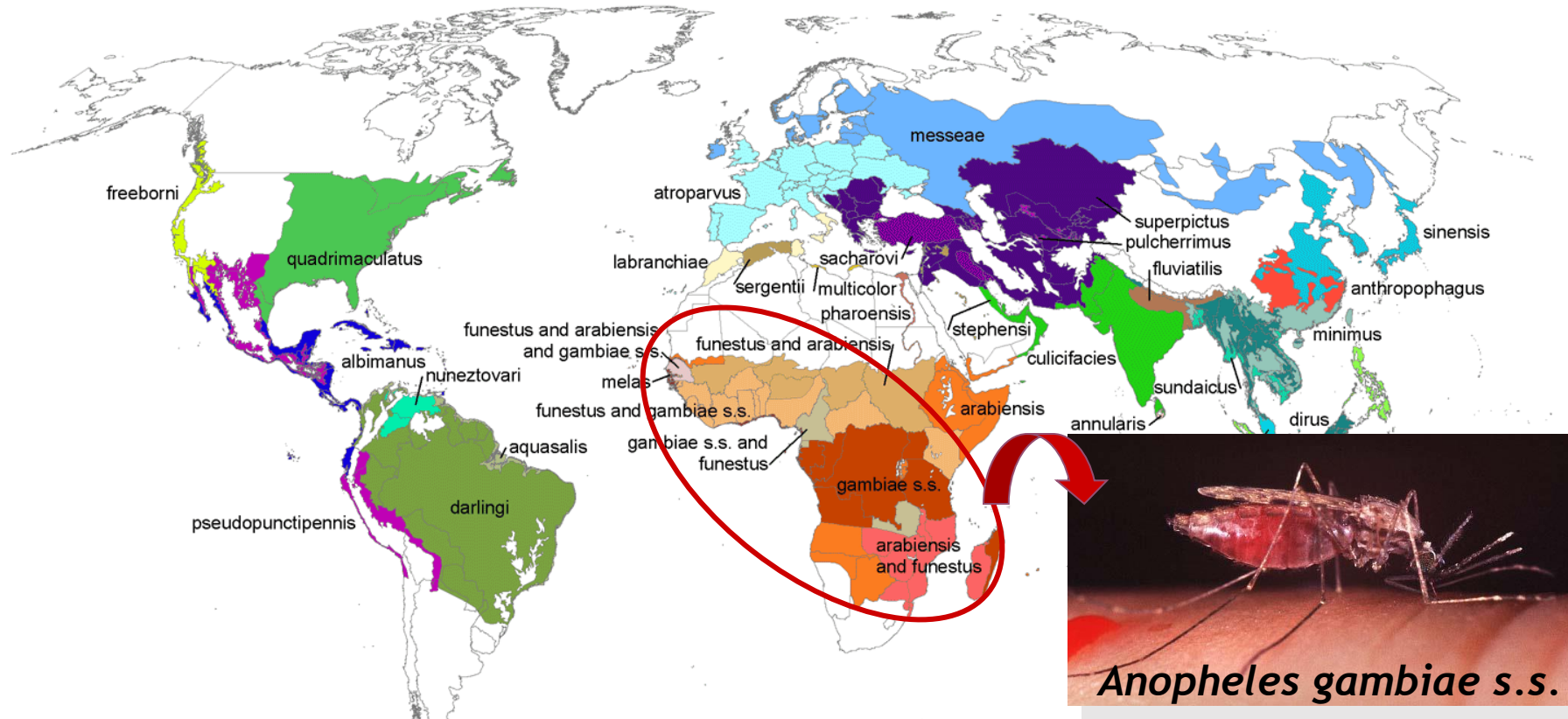
ISPI

# Number of deaths from malaria



Source: Institute for Health Metrics and Evaluation (IHME)

# Principali vettori della malaria (genere *Anopheles*)

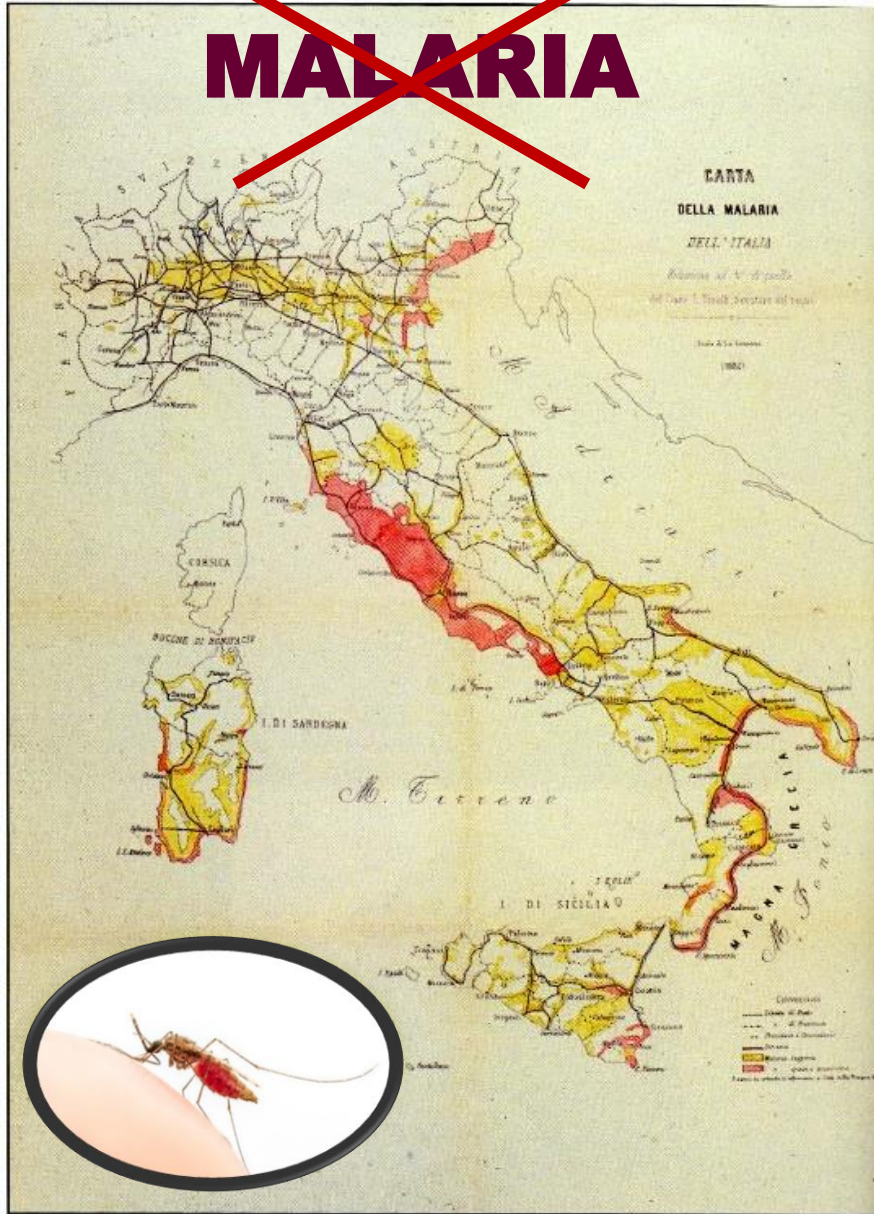


## Anopheles

- |                           |                |   |                      |                   |
|---------------------------|----------------|---|----------------------|-------------------|
| ○ No vector               | ● barbirostris | ● funestus and arabiensis               | ● melas              | ● pulcherrimus    |
| ● albimanus               | ● culicifacies | ● funestus, arabiensis and gambiae s.s. | ● messeae            | ● quadrimaculatus |
| ● annularis               | ● darlingi     | ● funestus and gambiae s.s.             | ● minimus            | ● sacharovi       |
| ● anthropophagus          | ● dirus        | ● gambiae s.s.                          | ● multicolor         | ● sergentii       |
| ● arabiensis              | ● farauti      | ● gambiae s.s. and funestus             | ● nunez-tovari       | ● sinensis        |
| ● arabiensis and funestus | ● flavirostris | ● labranchiae                           | ● punctulatus group  | ● stephensi       |
| ● aquasalis               | ● fluviatilis  | ● maculatus                             | ● pharoahensis       | ● sundaicus       |
| ● atroparvus              | ● freeborni    | ● marajoara                             | ● pseudopunctipennis | ● superpictus     |

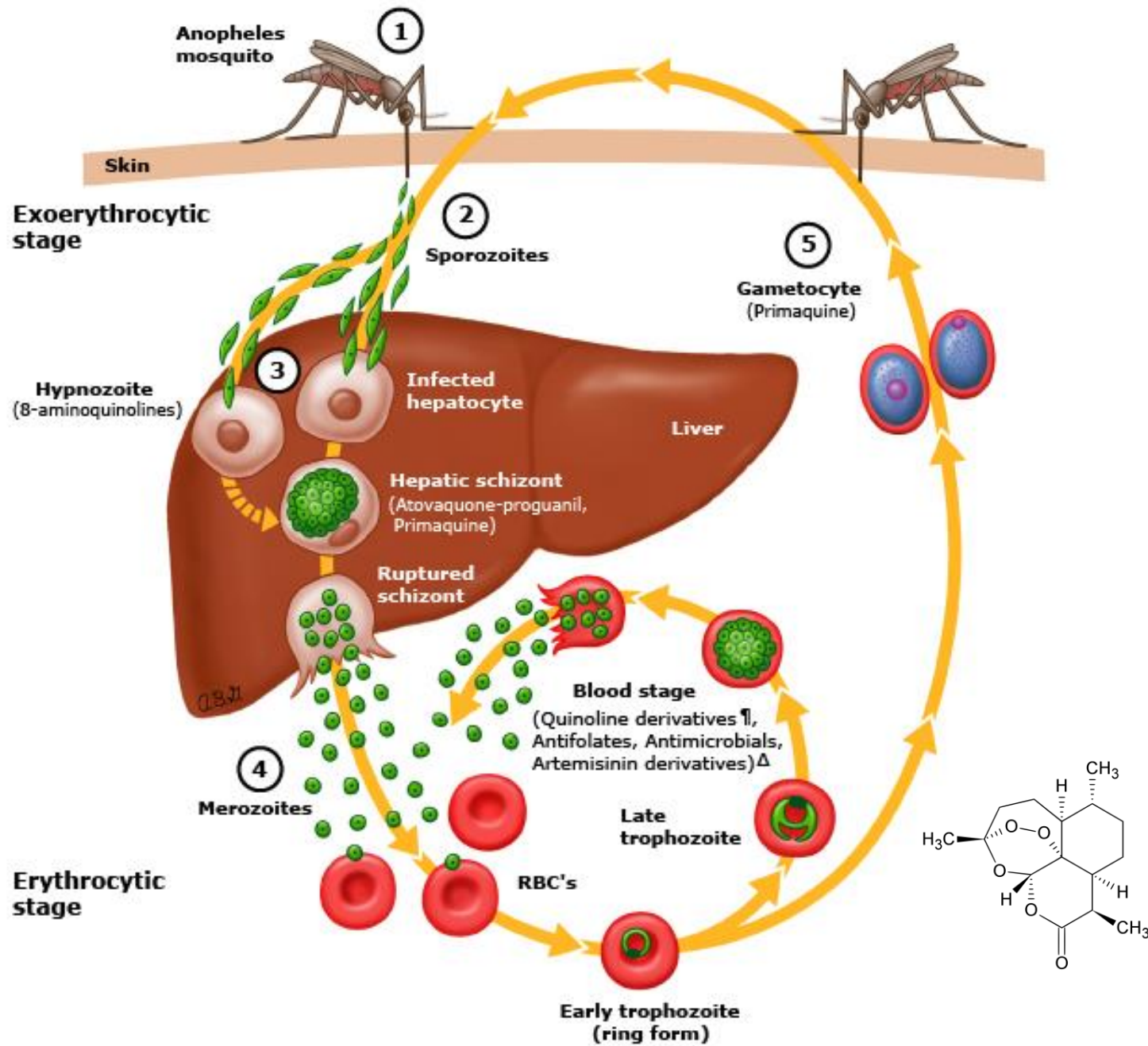
# E in ITALIA??

~~MALARIA~~

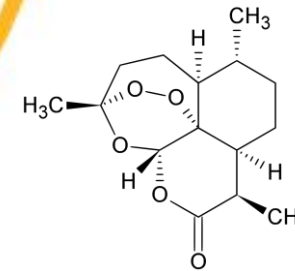


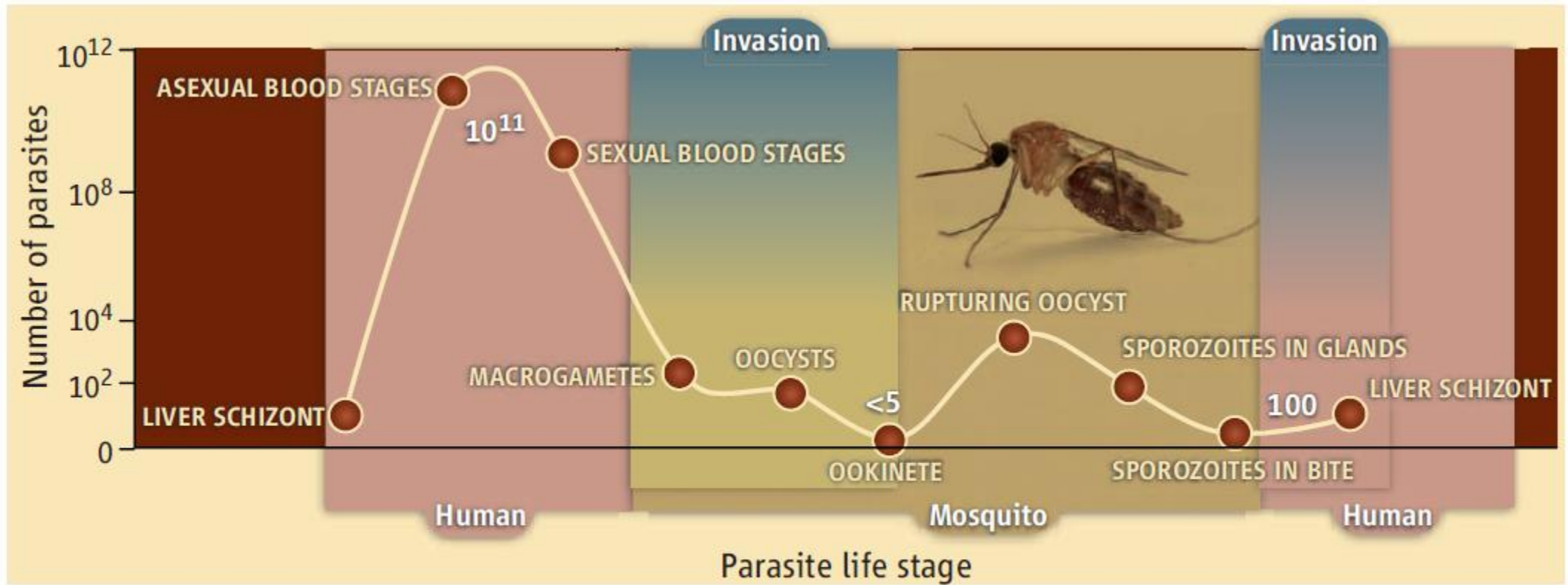
L'Italia, in data **21 settembre 1970**, fu iscritta dall'O.M.S. nei registri ufficiali dei Paesi liberi da malaria.



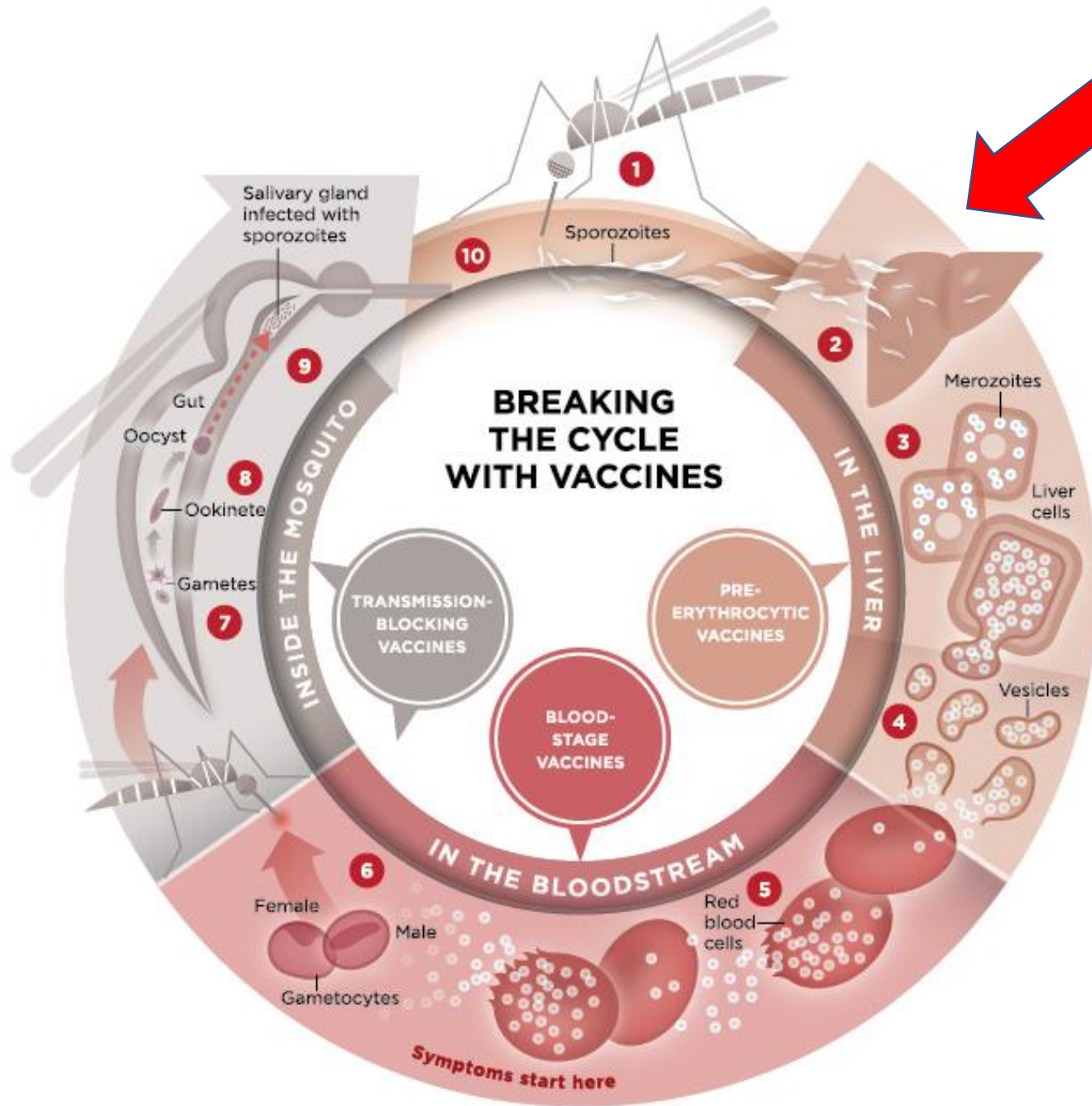


Ciclo della malaria  
(e farmaci  
antimalarici)





**Vulnerable target.** The malaria parasite's population ranges from about five inside the mosquito to trillions in the human bloodstream.

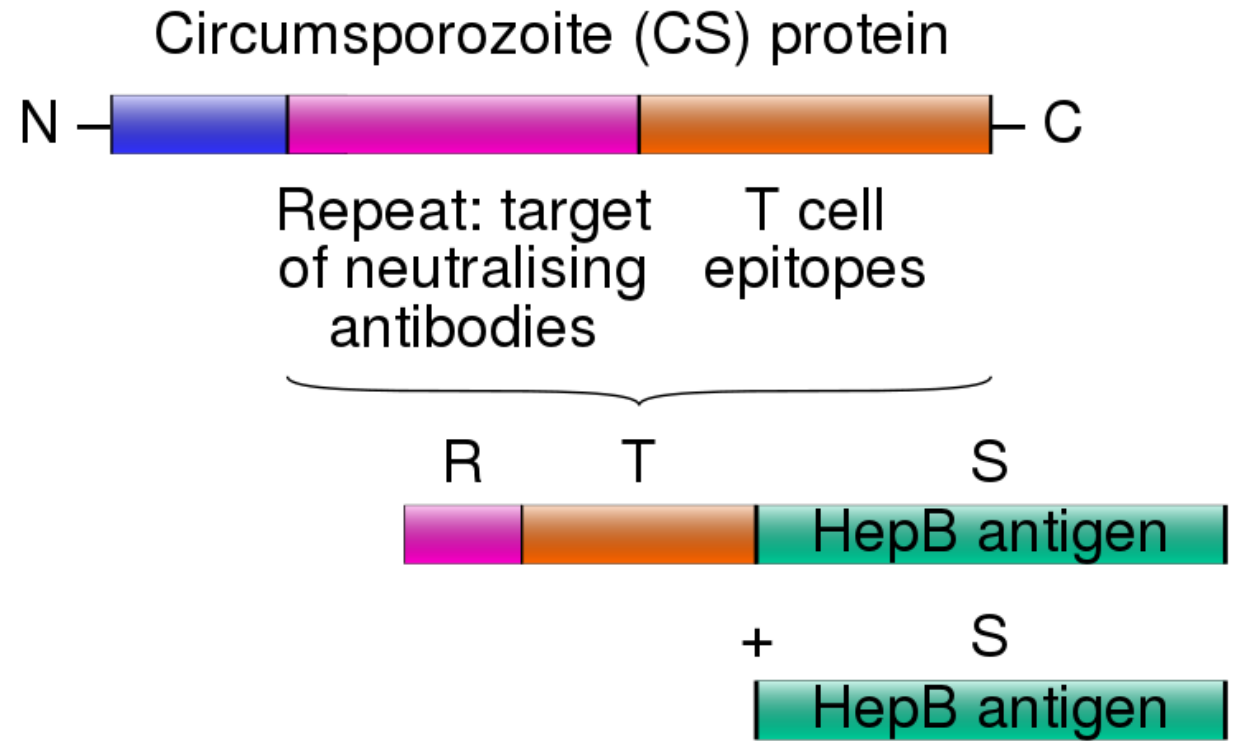


- Vaccini che portano alla produzione di anticorpi in una fase pre-eritrocitica
- Vaccini che agiscono a livello del flusso sanguigno
- Vaccini che bloccano la trasmissione

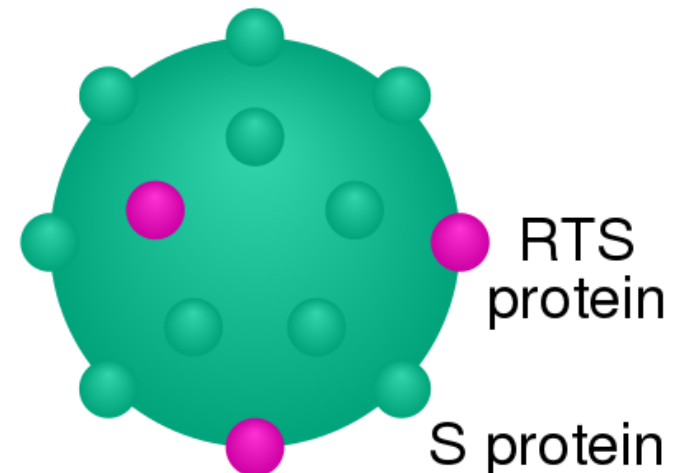


# Mosquirix (RTS, S/AS01) vaccine

- Proteina circumsporozoitica del *Plasmodium*  
+
- Antigene superficiale del virus dell'epatite B  
+
- Lisosomi contenenti un antagonista del Toll receptor 4 (per attivare la risposta immunitaria)



Co-expression of **RTS** (fusion protein) and **HBS** protein that assemble into mixed particles, in *S. cerevisiae*



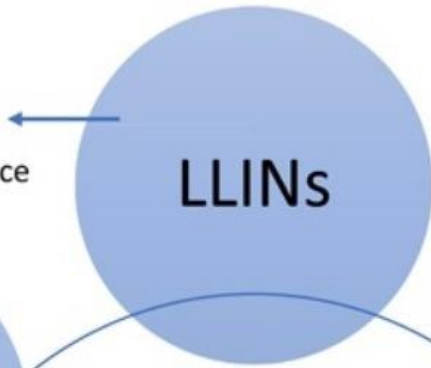
# Mosquirix (RTS, S/AS01) vaccino

- Primo studio nel **2015**, **120.000 bambini** di **7 paesi africani** (e prima approvazione dell'E.M.A.)
- Nell'ultima fase di una sperimentazione condotta in **3 Paesi** pilota (Ghana, Kenya e Malawi) su circa **800.000 bambini**, Mosquirix si è mostrato efficace nella riduzione dei contagi infantili soprattutto entro i 12 mesi successivi alla terza iniezione: prevenzione di malaria grave per bambini 5-17 mesi = 56%; per bambini 6-12 settimane = 31%  
**(approvazione W.H.O., 6 Ottobre 2021)**

**Margine di efficacia basso (circa 30% nel ridurre le forme di malaria grave nei neonati) MA grande risultato (ci vogliono 4 dosi...)!**

(potrebbe prevenire circa 25.000 morti l'anno nei bambini sotto i 5 anni!)

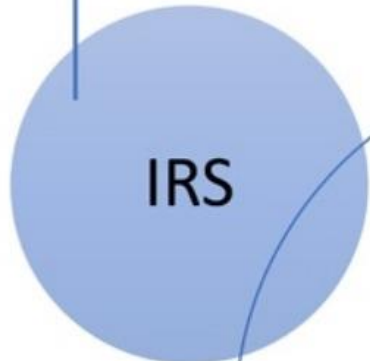
- Reduce malaria incidence
- Little impact on malaria prevalence
- Do not protect from outdoor transmission
- Growing levels of insecticide resistance



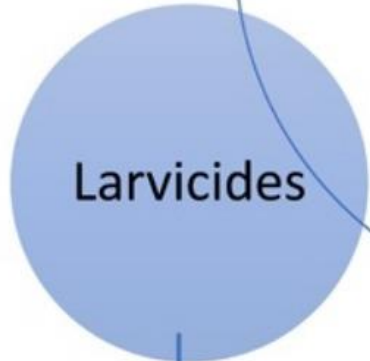
- Effectiveness, field suitability
- Must have epidemiological impact
- Non-target effects?



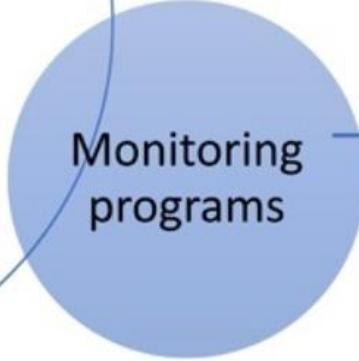
- e.g.
- Eave tubes
  - ATSB
  - Ivermectin



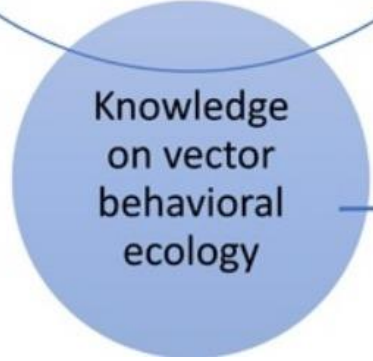
# Malaria vector control



Not suitable for rural areas



Environmental changes: impact on population dynamics



- Behavioral avoidance
- Vector biodiversity
- Competitive and food web interactions
- Dispersal
- Mating and sugar-feeding behavior



# The emergence of (various) resistance mechanisms

