## THOMAS R. ODHIAMBO (\*)

# Malaria research in Africa: new approaches to vector control (\*\*)

The satistics regarding malaria throughout the tropical and sub-regorded regions of the world are horserodous, persistent, and certainly demonstrates that is the single most widespread vector-borne disease among the people of the developing tropical regions (Table 1). About 100 million cross occur annually, it is estimated conservatively that if minimal protection is to be provided for 60% of the peopletion as triak in 1985, which is estimated to be about 1.5 billion people, such protection would cost U.S. \$2.9 billion (McEivers, 1976) a year. Such an expeculture is a major benefit not provided developing countries, on the contraction of the Sadan, which produces—through a major irrigation between 50% and for the produces and the such as the contraction of U.S. 4 for million was carmacked for malaria control over the eight-year period, 1976-1984. This budgetary allecation a presented a per capital inventment of U.S.3 for its malaria control alone from a population whose overall per capita income is only shout \$11.2.

Such an investment in malaria control is undoubtedly heavy; it is, however, commensurate with the high profile this tropical disease occupies in the tropical developing regions of Africa and the world in general. Malaria, and other vector-borne tropical diseases, have serious social consequences:

They cause high mortalities, especially among the young;
 They cause long-term debility: development of anaemia due to malaria.

infection, impairing the productivity of labour, lowering the qualitative output

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Table 1 - Malaria situation in all endemic areas world-wide, 1965-1985 (in millions).

Number of people			Estimates	
1965	1970	1975	1980	1985
1,041	1,088	1,191	1,293	1,459
635	650	840	900	1,021
406	438	351	393	438
	1,041 635	1965 1970 1,041 1,688 635 690	1965 1970 1975 1,041 1,088 1,191 635 650 840	1965 1970 1975 1980 1,641 1,088 1,191 1,293 635 650 840 900

Source: McEvers (1976).

of workers due to their impact on intellectual fulfilment; and leading to losses due to absenteeism;

 They cause waste of current resources invested in health care services (septially among the young), education, and agricultural production in the rural areas.

Thus, the control of malaria is seen very much in the context of national development objectives in Africa.

The cost of mularia treatment and control is colousal and persistents. Considerable rates of submercious among malarious school children in Africa, scholastic retraductions, stanting of growth, and impairment of normal growth and development that the control in high exclusive areas made the desicnoisal system in a control of the contr

Yet, the conventional methods for malaria control are so inadequate that they cannot either reduce transmission or prevent repeated infection on a continuing basis.

### Current malaria control strategies

The centrel of malaria in Africa has reached crisis proportions due too development of resistance to conventional symbietic insacricides, unavailability of replacement inserticides without resistance problems, and the development by the main malaria agent in Africa (Paramation Astroparo) of resistance to deboe-quiete, until recently the main chemotherapeatic anti-malarial drug used throughout Africa.

Resistance of the main mosquito vector of maleris in sub-Salatara Africa, Anophelie genelite complex (now known to have 6 sibling species), was first recorded in Africa Western Solono province of northern Nigeria in November 1955, where delethin had been used to speay the size 3 times over the previous 18 months. Includes of resistance to delichin were then from sirphily in many parts of West and East Africa (Rown and Pa), 1971). Since then, one can save that delethin resistance is asserted throughout sub-Salatana Africa.

Resistance to DDT note much longer to become evident. Indeed, it is howen that inside at amongitor tentimes to DDT workfuled is concerned, DDT evidentance develops more slowly (Bown and Pal, 1971). The first DDT-resistance calculation of the property of

Marks of insecticial resistance reported in Africa (and elsewhere in the tropics) as a result, not of direct malants coursely insecticidal treatment, to relique in a result of the control of Control of Control of the Control of Control of the Control of the Control of the Control of Control of the Control

Such uncoordinated action, hetween plant protection gractices and malaria courted campaigns, have led to well documented near catarophes, spart from insecticide-resistance arising in hitherto unexpensed fashion, as recorded above. Most of these are related to the development of legar fringings schemes in order to increase agricultural production in Africa (and other tropical countries). In Kenpy's Above Dito Tirgition's Scheme, for instance, the application of dimercina (and application) of almost control and application of almost control and application of almost control and application of an extension researance of the first necessary and the state of the above and control and application of an extension researance of the first necessaries of the publy fields because of the disease of its inernal productors (Service, 1984). Thus, the integration of such agricultural schemes, involving plant protection scriptings, each to be integrated with short involving malarias country, and its overage, this is an important part of the strategy to control vector monopalous in such a numer as to interrupt malaria transmission on a continuing bear.

We have presently few insecticidal agents which are effective against malatia vectors in Africa, except perhaps temephos (a larvicide) and fenotrothion (an adulticide). This major setback has been made more acute by the emergence of chlorosyniae-enismat. P. Infeprene in Mrica Erin coxed some 10 years ago have more than \$1984. WHO's Malaria Action Programme was able to show on a map the buses to which chlorosyniae-enismate had gened in Africa overrige Arapia. For the Maria Action Programme Arapia (Albon, Service, Mangasea, Malaria and Monaniae, Calonia, Service, Mangasea, Malaria and Monaniae and amboquistics, Genous, Madagasea, Malaria and Monaniae and amboquistics of the Malariae and Malaria

The possibility of developing as nationalasis vectice is being held at the present time as the ultimate long-term solution of the mulaite problem. Certainly, the World Bank/UNDP/WHO Special Pragramme for Research and Training, in Tropical dissues is concentrating in research and development effort in immunological control of mulairs. But this approach is still a long way from being a particule possibility, as a evidenced by the wide-enging NSSMI/USAD/WPMI workshop on the "Immunology of Malieris" held in October 1978 in Bethenda, Marriad, USAS (WHO, 1979).

We are back to making do with vector control as still the most feasible method for large-scale malaria control for the forescendic future. What are the characteristics of vector mosquirous in sub-Subarra Africa which have made malaria control so difficult in the para? The about naver is to be found in the epidemiological characteristics of the Aar, gambiae complex, the major transmistation of the property of the work, should, Anopole for justice is another although mixer or the work, should, Anopole for justice is another although mixer or property of the work.

## Epidemiological perspective

An genthur plays the major role in the transmission of malaris, hasterdalian filtratinal (Wardersite Saversity) and opengrapous free or in tropical Africa; and there have therefore been considerable conlegical and behavioural studies understand in respect of those elements without would give relevant information occurs to the contract of the contra

Barber and Olinger (1931) first noticed slight variations in appearance and breeding habits of a melanic coastal variety of an An. gambiae mosquito form in

West Africa, known to be associated with salt-water breeding sites. These distinctions were later utilized to separate, as distinct species, An. gambiae sensu stricto and An. melas: that is, physiological differences in salt-water breeding characteristics, and morphological differentiation provided by an extra dark band on the palps of adult females of An. melas and differences in larval pecten teeth (Ribbands, 1944a, b). These species distinctions were carried further by Moinhead-Thomson (1948), who demonstrated the fact that An. melas eggs are significantly larger than those of the freshwater-breeding An. gambiae s.str., and that on cross-fertilizing the two species they showed the presence of hybrid sterility. This resolution of the salt-water breeding species in West Africa raised the question whether the salt-water forms previously recorded in Mauritius and the coastal areas of East Africa also belonged to An. melas (Muirhead-Thomson, 1951). Crossing experiments made by Paterson (1962) confirmed that the East African salt-water form was indeed different from the freshwater-breeding species; crosses made between West African An. melas and the East African saltwater form showed that, indeed, the two salt-water forms were separate species (Davidson, 1962), later known as An. merus.

The freshwater As, gambiae mongatines themselves proved to be heterogeneous. Cooling experiments carried out in the early 1960's clearly domonstrated the existence of two other shifting species, spart from As, gambiae start. Cenning experiments reported by Duridoun (1962) and Duridous and Jackson (1962) recognized the separate existence of As, arabitents. Similar counsing experiments made by Patrono et al. (1963) differentially recognized An quantizementary, perciously described from the high welds of Tennoval, South Africa. A deced later, a footh freshwater species, so-coiled species. Do was found from Semilici Forces along the Uganda-Zaire border (White, 1973), and seems not to have been recognized from elsewhere in Africa.

This factinating denetive story was throught to a close by the matterly and comprehenive taxonomic revision by Mattingly (1977), which reduce the the 6 shiling species of the As gambiae complex (Table 2), and provided the 6 shiling species of the As gambiae complex (Table 2), and provided their essential diagnostic characteristation. Since then, those dislutricious when the second the contraction of the contr

This paintarking transmonic study, spanning over 30 years, opened up a new reassessment of the ecological and behavioural information critical to the epidemiological differential importance of the various components of the Aze grawhise complex. For instance, a study of the distribution of the different athling species was instructive: while Aze garden satt, and Aze audientia see very widely distributed throughout undo-Saharan Africa, localiding the Indian Ocean islands, Aze, garden's star, prodominates in hundred and the Aze audientia in takinety and the Aze audientia in the Saharan Africa, localiding the Indian Ocean islands, Aze, garden's at statistically protein any even become yenipatrie, for example, in the likeside area of Kimmu, in Kenya, where Aze garden's sax; concumber Aze audientia in the Daniel Saharan Africa, and concurrent the Aze audientia indian bouses towards the end of the cool ofly season, but the

Table 2 - Sibling species of the Anopheles gambiae complex in Africa (after Mattingly, 1977).

Old name	New name, and where found originally
Species A	Anopholes gambiae sensu stricto Giles, 1902. Material originally collected in Gambia.
Species B	An. arabicusis Patton, 1905. Found in semi-stid and stid sones.
Species C	An quadrizenulerus Theobold, 1911. Original material collected from Onderstepoort, South Africa
An. melas	An. moles Theobold, 1903. Name unchanged. Species associated with West African brackish-water breeding sites
An mens	An. meruz Donitz, 1902. Name unchanged. Species associated with salt water beceding sites in East African coastal zones
Species D	Species D: still unnamed. Found in mineral water in Bwamba swamps

reverse situation obtains in the hot dry weather following the short rains (Service, 1970; White, 1972). In contrast to the Africa-wide distribution of these two sibling species. An. melas is restricted to the West African coast from Angola to Senegal; and An. merus is confined to the East African coast from northern Natal (in South Africa) to lower Juba-land (in southern Somalia), including the Indian Ocean islands (Pemba, Mauritius and the Comoros), although this latter species has been recorded as breeding 50-120 km inland in Swaziland (Paterson et al., 1964) and coastal Kenya hinterland (Mosha and Subra, 1982). In this latter respect, Mosha and Mutero (1982) found that Au. merus can occasionally breed in outdoor sain-water, and can even experimentally breed successfully in freshwater in the laboratory. It is therefore of interest to find that An morus and two freshwater sibline species (An. gambiae s.str. and An. arabiensis) are found in sympatric association in certain villages along the Kenya coast of differing proportions. For example, in one village (Jimbo), 96% of the An. gambiae complex mosquitoes consist of An. merus; while in another village (Jego), only 17.4% comprise An. merus, while An. gambiae s.str. constitutes 71.7%, and the rest An. arabiensis (Mosha and Petrarca, 1983). The remaining two sibling species of the complex, namely An. auadriannulatus and species D, show a patchy and relict distribution: the first species seems to be confined to Zanzibar, and the highlands of Ethiopia and southern Africa: while the second has so far only been collected from Semliki Forest in Bwamba (White, 1973).

The strong differences in distribution among the sibling species of the An gambiae complex, considered together with other differences subsequently found in the differing vectorial capacities of these selfsame species, have strengthemed the conviction that these characteristics are basic to the epidemiological understanding of these malaria vectors in Africa. Multibasi/Thomson (1931) first showed that Xan, nearst exhibits a lower vectorial perceptive for malaria than An, genidate satz. White (1974) followed up these pioners studies by demonstrating that the major vectors of malaria (as well as basecrotian fills ritabi) in Africa were in fact An, genidate xatz., An endownist and An melas, room chough the latter was confined no countal areas of West. Africa. It is unlikely that, because of its very restricted dustribution, species D is epidemiologically with the parameter of the product of th

The feeding behaviour of the different tilling species somewhat follows the pattern of findings from vectorial capacity studies. As a gambia state perfects human blood, and is normally enable feeding inside bosser, As media does has strong perfectence for human blood, but blets both indoors and outdoors. It is a set of the pattern of the perfect of the pattern of the perfect of the pattern of

Information on resting behaviour of the different sibling species shows that the overall epidemiological picture is likely to be highly complex, while demonstrating clearly why An. gambiae s.str. is preemmently the leading malaria vector in sub-Saharan Africa. An. gambiae s. str. is the most endophilic of the sibling species, predominantly preferring to rest in houses or man-made shelters. Three other species (An. arabiensis, An. quadriannulatus and species D) rest either indoors or outdoors, depending on the availability of resting sites next to where feeding takes place. The two salt-water breeding species (An. melas and An. merus) are definitely exophilic. Thus, of those An melas which may feed indoors during the night, 90% leave for outdoor resting sites by dawn (Gelfand, 1955), where they rest in rodent burrows, shaded earth banks, etc. Similarly, while only 2% of An. gambiae s.str. left houses after feeding, 35-40% of An. seerus did so (Muirhead-Thomson, 1951); and the latter then go on to rest in crab holes, under fallen leaves, in shaded termite mounds, heavily shaded mangrove and mangrove trees, etc. (Iyengar, 1962), or coconut husks (Mutero, Mosha and Subra, 1984). Indeed, An. merus feeds mostly outdoors (Mutero, 1980; Mutero, Mosha and Subra, 1984), with peak biting activity between 2400 and 0100 hours, whether indoors or outdoors. If indoors, most biting occurs while the people are asleep, and then the An. merus mosquitoes move out shortly afterwards (Mutero, Mosha and Subra, 1984). This work shows clearly that residual insecticidal spraying of houses at the Kenya coast may kill off An. gawbiae s.str., but the largest proportion of An. merus population will escape the treatment.

The other major factor of epidemiological importance is the estimation of narrival trace of field populations of the individual species or flantist vectors. The most thorough study so far done in Africa is that of Matters (1985) understand as at the ICIPE on the three siding periods symptotically associated at the Kenya coasts. In brief, his findings show that: (a) the day season population of the gardine start, restational a longer life than the verseason population of the gardine start, restational a longer life than the verseason population of the gardine start, inc. the because of the relatively restricted eco-groupshical distributions ange of the mera, both because of the relatively restricted eco-groupshical distributions of this species, the much more subsequent day, in other beds of the restriction of the species start, in, on the whole, the more important vector of malatis, in spite of its vectorial capacity being 4 times less than that of the mora factorial capacity being 4 times less than that of the mora factorial capacity should be substantially the vectorial capacity, becomes calculated and the substantial as a factorial vector takes much longer.

The epidemiological perspective being adduced now since the choledsion of the taneounic compositive of the malaria veners in sub-Saharan Africa is spidly giving us a better understanding of the vector-parasite-bost transmission factors that night eventually lead to more rational integrated control of malaria. The biological control factor of this problem has not yet received much amendion, and substantially administration of the factor of integrated veneric control strategies in Africania; no be decorated to this factor of integrated veneric control strategies in Africania; no be decorated to this factor of integrated veneric control

## Potential biological control of mosquito sectors of malaria

More than 1,000 natural enemies of mosquitoes have been recorded worldwide. These include: larvivocous fish (such as the minnow, Gambasia effinis), which has been utilized for mosquito control since the early 1900's, but has lately declined in its effectiveness in some regions due to the advent of DDT and similar synthetic insecticides, predation by other fish, and decimation by fungal diseases; larvicidal plants, such as rooted algae of the family Characeae (especially Chara and Nitella) which exude toxin into the water, and other plants (such as Phytolacca and Melia) whose fruit contains anti-larval compounds; parasitic nematodes (especially mermithids) in larvae, for example Reesimermis nielseni; parasitic protozoans, especially spore-forming species, such as Nozema stegomysae, that kill larvae, pupae and adults, or reduce adult longevity; pathogenic bacteria, especially the spore-forming Bacillus thuriengiensis; invertebrate predators, especially the predator larval mosquitoes of the genus Toxorbynchites living in tree-holes or other small containers, and feeding on mosquito eggs, larvae and pupae; and pathogenic fungi, particularly those belonging to the genus Coelomonyces (National Academy of Sciences, 1973). There is recent resurgence of interest in the use of Toxorbynchites and Coelomontyces in Africa for malaria control.

Tourselymether is primarily a tropical genus, which takes no blood in the ability and teals requised interpolar a prey. The most adults and teals requised interpolar as prey. The most studied species so far, T. Investigably (Theodolds), breeds in tree-boles and feels statis, and has wide distribution, throughout sub-Subaran Africa, where it is most abundant in forest vegatation, and not in peridomentic situations or cubi-vated ind (Lomolius, 1979). The large author observes that this predature classity is in contrast to the pery density (unish) consisting of Aceler mosquitonity, which is highest in the cubinvast across. Even though T. Verrapicali lower colonisated in the laboratory, and mass. Even though T. Verrapicali lower colonisated in the laboratory, and mass. 1973), and prey population reduction by Terrapical lower in the colonisated and prey population reduction by the colonisated of the colonisated colonisated and the colonisated colonisated and the colonisated colonisated to the displacement between its population and that of its prey, and the fast that its moughind due to the displacement to its usual more arrived only included to its uniform the carbolic, with Are gambiar complex being appearantly only includent to its sund more and the superior and the colonisated colonisation.

The potential for the use of Coelomomyces in malaria vector control is somewhat higher. The genus, of which 40 species are known, is widespread in all the continents, except the Antarctica, but is virtually restricted to mosquito hosts, causing high mortality in infected larvae (Laird, 1967). Among the mosquito species it attacks, are An. gambiae s.lato. All Coelomomyces species require an intermediate crustacean host, usually a species-specific copepod or ostracod, and a definitive host, a mosquito larva. Because of this circumstance, it has been possible to have in vivo mass cultures of this fungus, since mid-1970's, my maintaining separate cultures of the infected copepeds and mosquito larvae (Federici and Chapman, 1977). So far, no successful attempt has been made for in vitro culture. If this were accomplished, and because of the obligate parasitic nature and restricted host range of Coelomomyces, the latter would have a high potential for the control of malaria vectors in Africa in at least two possible ways: as a microbial insecticide (the way B. thuringiensis toxin-bearing spores are presently being used); and as a suppressant of mosquito populations on a long-term basis when the fungus is introduced into areas in which it was previously absent.

That possibility has been raised recently by the discovery of epitocetics of  $Aar_{\rm gardots}$  complex berow being caused by a high infection by  $\ell$ . raiset in Kongseves and Rabai, at the Kerspa coast (Odineo  $\sigma at$  al, 1980). Sampling theoughout the pares aboved that infection varied between 33 and 100%. This finance and its copped intermediate bosts, have been rareed in the laboratory regularly, and its copped intermediate bosts, have been rareed in the laboratory regularly and it has been formed the the expression value in the laboratory regularly and it has been formed the three copped overhild in 8 bosts or in flooding 65 lower at at, 1990). There is still a long way to bring the Costomoroper potential to the text, but a good start has been made.

#### Conclusion

It is imperative that the direction the current research in Africa on malaria control has taken should be pushed to the limit. It is much more sensitive to coolgical and environmental issues, the coordination between plant protection and vector control, and the long-range goal of integrated vector and malaria control than during the sen of exclusively inserticials control regimens.

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