Mosquitoes of Etiological Concern in Kenya

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Kenya is among the most affected tropical countries with pathogen transmitting Culicidae vectors. For decades, insect vectors have contributed to the emergence and distribution of viral and parasitic pathogens. Outbreaks and diseases have a great impact on a country's economy, as resources that would otherwise be used for developmental projects are redirected to curb hospitalization cases and manage outbreaks. Infected invasive mosquito species have been shown to increasingly cross both local and global boarders due to the presence of increased environmental changes, trade, and tourism. In Kenya, there have been several mosquito-borne disease outbreaks such as the recent outbreaks along the coast of Kenya, involving chikungunya and dengue. This certainly calls for the implementation of strategies aimed at strengthening integrated vector management programs. Here we look at mosquitoes of public health concern in Kenya, while highlighting the pathogens they have been linked with over the years and across various regions.

Keywords: Aedes ; Anopheles ; Culex ; Mansonia ; pathogens

1. Introduction

The term "vector-borne" has become a commonly used term, especially in tropical and subtropical countries where emerging and re-emerging vector-related diseases frequently occur. One-sixth of human diseases is associated with vector-borne pathogens, with approximately more than half of the global population currently estimated to be in danger of contracting these diseases [1]. Hematophagous mosquitoes are the leading vectors among arthropods because of the significant role they play in disseminating microfilariae, arboviruses, and *Plasmodium* parasites that seem endemic to sub-Saharan Africa [2,3]. The majority of these pathogens are maintained in zoonotic cycles and humans are typically coincidental dead-end hosts with a none-to-minimal role in the cycle of the pathogen [4].

Mosquito vectoral ability is greatly influenced by the availability of conducive breeding grounds, which is in turn, influenced by the spatial heterogeneity as well as the temporal variability of the environment [5]. Mosquitoes, pathogens, and hosts each endure and reproduce within certain ideal climatic conditions and changes in these conditions can greatly alter these pathogen transmission/competences. In this scope, temperature and level of precipitation are the most influential climatic components, but other factors such as sunshine length, sea level elevation, and wind have been shown to have considerable effects [6,7]. These vectors often adjust to changes in temperature by changing topographical distribution. For instance, the advent of malaria cases in the cooler regions of East African highlands may be attributed to climate change, which has led to an increase in mosquitoes in the highlands as they warm up [8]. Variability in precipitation may also have a direct influence on distribution of mosquito-borne diseases. When precipitation increases, the presence of disease vectors is also expected to rise due to the expansion of the existent larval habitat and emergence of new breeding zones [6]. Each mosquito species has unique environmental resilience limits dependent upon the availability of favorable aquatic larval habitats and the closeness of vertebrate hosts that serve as their source of blood meals. This reliance of mosquito species on aquatic environments is a constant part of their lifecycle and the availability of a suitable aquatic domain, which is a requirement for the development of eggs, larvae, and pupae, and basically determines the abundance of mosquito species.

Kenya represents a topographically diverse tropical/subtropical country which harbors a large diversity of mosquito species of public health importance. Many factors contribute to the extensive proliferation of mosquitoes ranging from global warming, sporadic floods, improper waste disposal, irrigation canals, presence of several lakes/rivers, and low altitudes around coastal regions. Consequently, an upsurge in emerging and re-emerging mosquito-borne pathogens over the years has been recorded with increased research efforts geared towards mosquitoes and their pathogens [9]. Increased urbanization, tourism, and international trade have led some of these species and pathogens to cross local and international borders to new territories. This dispersal poses both local and global health threats if proper mitigation measures are not put in place. Nevertheless, not all mosquito species are associated with human diseases, thus, this review highlights species of the main mosquito genera to which pathogens have been associated/detected and their

countrywide distribution based on published data and reported cases. Additionally, this information provides a guide to proper mosquito control strategies by comparing the methods currently applied in the country and proposed alternative methods applied in other countries affected by mosquito disease burden.

2. Mosquito-Borne Disease Endemic Regions in Kenya

Kenya, as a tropical country, is among the most affected sub-Saharan regions with mosquito-related ailments. The countrywide distribution of mosquito species is, however, not well documented with most studies focusing on disease endemic regions. Some of these regions include Garissa, Mandera, and Turkana situated in the north and north-eastern parts of the country and characterized as arid and semi-arid regions (Figure 1A). During rainy seasons, flood water acts as breeding areas for mosquito species. Rift Valley fever outbreaks were recorded in this area in 1997/98 and 2006/07 [10,11]. Other viruses reported from the regions include West Nile virus (WNV), Ndumu virus (NDUV), Babanki virus (BBKV), and orthobunyaviruses isolated from the flood water *Aedes* and *Culex* species [4]. For decades, the coastal region of Kenya (Kwale, Kilifi, Mombasa, Lamu) has also reported multiple outbreaks resulting from mosquito-borne pathogens making it one of the most endemic regions. The contributing factors are majorly the low altitude which provides a conductive environment for mosquito breeding and high human population composed of both locals and global tourists. Mosquitoes of interest in this coastal region in terms of pathogen transmission include: *Aedes aegypti* that transmits dengue fever and chikungunya [12,13]; *Anopheles* species (*Anopheles gambiae, Anopheles arabiensis, Anopheles funestus, Anopheles merus*) which are associated with malaria and bancroftian filariasis sporozoites [14,15], and *Culex quinquefasciatus* [16] among others as demonstrated in Figure 1A–C.

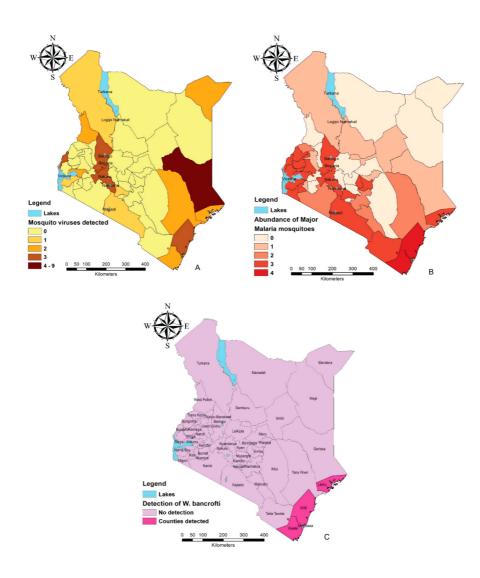


Figure 1. Mosquito-borne disease endemic regions based on distribution of pathogens and associated mosquito species (maps were constructed using the free and open-source Quantum GIS software (https://qgis.org/en/site/) using data compiled from <u>Table 1</u> and <u>Table 2</u>. (A) Abundance of mosquito-borne viruses detected/isolated in various counties. (B) Distribution of the major malaria vectors in numbers in different counties. (C) Counties in which *Wuchereria bancrofti* has been detected from mosquitoes.

Genera	Species	Virus Isolated/Detected ¹	County of Virus Detection	Reference
Aedes	A. aegypti	DENV, CHKV	Mombasa, Mandera, Kilifi, Lamu, Busia	[12,13,33,55,57,58]
	A. africanus	YFV	Baringo,	[<u>59]</u>
	A. albicosta	DENV, CHKV	Mombasa, Kilifi, Lamu, Kwale	[<u>33]</u>
	A. circumluteolus	RVFV, BBKV, NDUV, SMFV	Garissa	[<u>4,36]</u>
	A. fryeri	DENV	Mombasa, Kilifi, Lamu, Kwale	[<u>33]</u>
	A. fulgens	DENV, CHKV	Mombasa, Kilifi, Lamu, Kwale	[<u>33]</u>
	A. keniensis	YFV	Baringo	[<u>59]</u>
	A. Luridus	NDUV	Tana River	[<u>4]</u>
	A. mcintoshi	RVFV, NDUV, PGAV, BUNV, BBKV, PGAV, SMFV, NRIV	Garissa	[<u>4,11,36,40]</u>
		DENV, CHKV	Mombasa, Kilifi, Lamu, Kwale	[<u>33]</u>
	A. ochraceus	RVFV, NDUV, BUNV, BBKV, SNBV, SMFV	Garissa	[<u>4,11,36,37,38]</u>
		DENV, CHKV	Mombasa, Kilifi, Lamu, Kwale	[<u>33]</u>
	A. pembaensis	RVFV	Kilifi	[4]
		DENV, CHKV	Mombasa, Kilifi, Lamu, Kwale	[<u>33]</u>
	A. sudanensis	BBKV, SNBV, WNV	Garissa	[<u>36]</u>
		NDUV	Tana River	[<u>39]</u>
Anopheles	An. funestus	ONNV	Kisumu	[<u>59]</u>

Genera	Species	Virus Isolated/Detected ¹	County of Virus Detection	Reference
		BUNV	Kajiado	[<u>4]</u>
		NRIV	Tana River	[<u>4]</u>
	An. gambiae	BUNV	Homabay	[20]
	An. squamosus	RVFV	Garissa	[11]
Culex	Cx. bitaeniorhynchus	RVFV	Kilifi	[11]
		NDUV	Tana River	[<u>39]</u>
	Cx. cinereus	NDUV	Busia	<u>[4]</u>
	Cx. pipiens	USUV	Kisumu	[<u>4]</u>
		NDUV	Garissa, Tana River	<u>[38,39]</u>
	Cx. poicilipes	RVFV	Kilifi	[11]
	Cx. quinquefasciatus	RVFV	Baringo, Garissa	[<u>11,60]</u>
		WNV, SNBV	Garissa	[<u>36,60]</u>
	Cx. rubinotus	NDUV	Baringo	[<u>4</u>]
	Cx. univittatus	RVFV	Baringo	[11]
		BUNV	Homa Bay	[20]
		SNBV	West Pokot, Nakuru, Busia	<u>[4,37,61]</u>
		WNV	Garissa, Turkana, West Pokot	<u>[4,61]</u>
	Cx. vansomereni	NDUV	Tana River	[<u>39]</u>
		BBKV, SNBV	Nakuru	[<u>4,37]</u>
	Cx. zombaensis	RVFV	Nakuru	[62]
		BBKV	Kiambu	[4]

Genera	Species	Virus Isolated/Detected ¹	County of Virus Detection	Reference
Mansonia	Mn. africana	RVFV	Nakuru, Baringo, Garissa	[<u>11,60,62</u>]
		NDUV	Baringo	[<u>4</u>]
	Mn. uniformis	RVFV	Baringo, Garissa	[40,60]
		NDUV	Baringo	[<u>36]</u>

Genera: *A, Aedes; An, Anopheles; Cx, Culex; Mn, Mansonia.* ¹ Abbreviations: DENV—dengue virus, CHKV—chikungunya virus, RVFV—Rift Valley fever virus, NDUV—Ndumu virus, PGAV—Pongola virus, BUNV—Bunyamwera virus, BBKV— Babanki virus, SMFV—Semliki Forest virus, NRIV—Ngari virus, YFV—yellow fever virus, SNBV—Sindbis virus, WNV— West Nile virus, USUV—Usutu virus, ONNV—O'nyong'nyong virus.

Parasite	Associated Human Ailment	Dominant Mosquito Species	Counties of Vector Distribution	Reference
Plasmodium falciparum	Malaria	An. gambiae s.s., An. arabiensis, An. funestus, An. merus	Kwale, Kilifi	
		An. gambiae ss, An. arabiensis, An. funestus	Taita-Taveta, Lamu, Kajiado, Embu, Nakuru, Baringo, Bungoma, Kirinyaga, Kiambu, Busia, Siaya, Kakamega, Vihiga, Homabay, Migori, Kisii, Kisumu, Nandi	
		An. gambiae ss, An. arabiensis	Narok	[<u>71,75,81,93,94</u>]
		An. arabiensis, An. funestus	Tana-River, Makueni, Machakos, Trans-Nzoia	
		An. funestus	Samburu, Isiolo, Garissa, Mombasa, Uasin-Gishu, Nyamira	
		An. arabiensis	Turkana	
		An. gambiae ss	Tharaka-Nithi	
Wuchereria bancrofti	Bancroftian filariasis	An. gambiae sl, An. funestus, Cx. quinquefasciatus	Kwale, Kilifi, Lamu	[<u>50,91,95]</u>

Table 2. Summary of the dominant malaria and Bancroftian filariasis mosquito vectors distributed in Kenya.

Mosquito-related studies have been carried out in some of the mainlands surrounding lakes in Kenya, such as Lake Victoria and Lake Baringo due to the prevalence of malaria [<u>17</u>]. Other outbreaks such as Rift Valley fever near Lake Baringo also led to intensified surveillance on arbovirus mosquito vectors in the surrounding regions [<u>4,18,19,20</u>]. The favorable tropical climate and swampy areas around the lake regions throughout the year is among the factors favoring the high diversity of mosquito species [<u>18,21</u>]. A study by Lutomiah et al. [<u>22</u>] which recorded high numbers of *Culex pipiens* and *Culex univittatus* around the two lake regions additionally associated this abundance to highly humid conditions and the presence of migratory birds. Other lake regions in Kenya that are important for mosquito-related studies include areas around Lake Naivasha [<u>23,24</u>], Lake Bogoria, and Lake Nakuru [<u>20,24</u>] (<u>Figure 1</u>A,B).

Over the years, several malaria and arbovirus seropositive cases have been recorded in Kakamega and Busia located in the western part of Kenya [25,26,27]. In a study conducted in 2013, Kakamega and Busia showed high numbers of potential arbovirus vectors, with *A. aegypti* at 91.8% and 45.6%, respectively, of the total mosquitoes collected in the two regions [22]. Additionally, NDUV has been detected from *Culex* species collected from Busia [4]. Kakamega's climate is characterized by high precipitation occurring throughout the year, most likely an influence of the Kakamega tropical rainforest while Busia is known to experience warm temperatures (annual mean temperature of 25 °C). These factors, among others, contribute to the high mosquito densities and consequent disease occurrence in the regions.

3. Overview of the Main Mosquito Genera of Public Health Importance in Kenya and Their Geographical Distribution

The main mosquito species associated with human etiologies in the country belong to four genera (*Aedes, Anopheles, Culex, and Mansonia*) as discussed below. Their distribution in Kenya has mainly been studied in disease endemic regions and as a result of outbreaks. These data are summarized in <u>Table 1</u> and <u>Table 2</u>.

3.1. Aedes

Aedes represent a genus of mosquito that were originally known as tropical and subtropical mosquito species but have now been reported in almost all parts of the world apart from Antarctica. This genus comprises over 950 species normally identified by their black and white body coloration, preferential breeding in open water containers, and their daytime feeding, especially mornings and evenings [28]. Much interest in these mosquito species has been due to their role as major vectors of arboviruses across the globe. For instance, in various islands of the south pacific, *A. polynesiensis*—a primary vector of *Wuchereria bancrofti*—has also been identified as an endemic dengue vector [29]. *Aedes scutellaris*, a native species of Papua New Guinea, has also been linked to dengue outbreaks around the region and other pacific islands [30,31]. *Aedes aegypti* and *A. albopictus* of African and Asian origins, respectively, are globally known for their invasive nature and role in transmitting viruses that cause chikungunya, dengue fever, yellow fever, Zika, and encephalitis [32]. The success of these species in virus transmission has been attributed to three mechanisms: interaction of vector and the host, transovarial transmission, and sexual mating [28,33].

Kenya harbors more than 80 different *Aedes* species (http://www.wrbu.org) but not all have been associated with the spread of mosquito viruses. An important species is *A. ochraceus* which is highly distributed in the arid regions of north-eastern Kenya and linked to the spread of Rift Valley fever virus (RVFV), NDUV, Bunyamwera virus (BUNV), Babanki virus (BBKV), Sindbis virus (SNBV), and Semliki Forest virus (SMFV) in the region [4.11,34,35,36,37,38]. Additionally, this species has also been reported to be associated with RVFV, dengue virus (DENV), and chikungunya virus (CHKV) in coastal counties (Kwale, Kilifi, Mombasa, Lamu) [11,33,39], and some parts of in the western region [22]. *Aedes mcintoshi* and *A. sudanensis* are also important arbovirus vectors occasionally reported in coastal and north-eastern regions [4.11,14,22,33,36,40], while *A. circumluteolus* have been reported in Busia and Garissa counties [4.22,36]. Other *Aedes* secondary vectors of arboviruses such as: *A. tricholabis* [4,33], *A. Albicosta* [33], *A. fulgens* [33], *A. fryeri* [33], *A. pembaensis* [33], and *A. Luridus* [4] have been identified in various parts of the country and mostly along the Kenyan coastline. *Aedes africanus* and *A. keniensis* sampled from Baringo county represent the only field collected *Aedes* spp. in which successful yellow fever virus (YFV) has been detected in Kenya [41]. Moreover, other *Aedes* spp. such as *A. vittatus* and *A. bromeliae* have shown competence to DENV and YFV, even though no field detection has been recorded [42,43].

Aedes aegypti, the principal vector of dengue, chikungunya, and other emerging arboviruses is also widely distributed in Kenya [4,33,44] but not uniformly, with more occurrence being recorded in the lowlands [22]. It exists in two forms that

were found coexisting sympatrically in Rabai, along the coast of Kenya [45]: the domestic, light-colored *A. aegypti aegypti* and the sylvatic form, *A. aegypti formosus*, which is dark in color [45,46]. In addition to morphological differences, definite behavioral variabilities have also been observed between the two forms of *A. aegypti* [46]. They include a longer developmental time and indoor breeding tendency of the domestic form as opposed to the ancestral sylvatic form that usually breed in forests tree holes and develop within a shorter period [47,48]. This explains the occurrence of *A. aegypti formosus* in vegetated ecosystems and previous documentation in western Kenya near Kisumu city and in the Kakamega forest [46]. Additionally, the sylvatic type is predominantly zoophilic as compared to the domestic type, which prefers humans for their blood meals.

Rapid urbanization in Kenya, facilitated by the increase in both domestic and international trade has led to major ecological and social changes [14]. These transformations have attracted *A. aegypti* to urban centers with up to three times the population size compared to rural areas due to the accumulation of non-biodegradable human-made containers that serve as conducive breeding sites to these vectors [12,49,50]. It is, thus, evident that if effective vector control strategies are not enhanced, the rate of arbovirus disease transmission is bound to increase, especially in urban centers.

Sporadic arboviral disease outbreaks associated with *Aedes* spp. have occurred over several decades in Kenya of which some have shown a tendency to recur in the same or new populations. For example, dengue fever in 1982 (Malindi, coastal region) [51], 2011 (Mandera, north-eastern) [52], and 2013–2014, 2017 (coastal region counties) [12,53]; Rift Valley fever in 1997–1998 (Garissa, north-eastern) [10], 2006–2007 (Garissa and Baringo) [11], 2014–2015, 2018 (north-eastern) [54]; yellow fever in 1992–1993 (Kerio Valley, Rift Valley region) [41]; chikungunya in 2004 (Lamu, coastal region) [55], 2016 (Mandera, north-eastern) [56], and 2018 (Mombasa, coastal region) [13].

3.2. Anopheles

This genera comprises mosquito species that are commonly identified by long palps, dark-brown body coloration, discrete black and white scales located at the wings and a characteristic resting position (stomach area pointing upwards) [63]. Over 460 anopheline species have been recognized across the world, with approximately 7% known to transmit *Plasmodium* parasites that cause malaria in humans [64]. Malaria is endemic in 91 countries, with about 40% of the world's population at risk. The latest World Health Organization (WHO) data showed that in 2017, there were 219 million cases of malaria and estimated 435,000 malaria deaths across the world. Out of these, Africa accounted for the most cases and deaths (92% and 93%), respectively [65].

Africa is home to more than 140 anopheline species, of which six of them are primary vectors of malaria parasites (*P. falciparum, P. vivax, P. ovale,* and *P. malariae*) [66,67]. They include *Anopheles gambiae, Anopheles arabiensis, Anopheles funestus, Anopheles nili, Anopheles moucheti,* and *Anopheles coluzzii* [67,68]. However, other than the primary vectors that contribute to 95% of malaria transmission in Africa, secondary *Anopheles'* vectors have also been recorded and contribute 5% of the overall transmission. For instance, *An. pharoensis* and *An. ziemanni* were found to harbor the malaria parasite *P. vivax* in Ethiopia at irrigated rice plantations [69]. In Tanzania, *An. ziemanni, An. Coustani,* and *An. squamosus* have also been linked to malaria transmission as secondary vectors [70].

In Kenya, *Anopheles* spp. are a major concern in the public health sector due to their role as malaria transmission agents. In a country-wide study by Okiro et al. [71], out of 166,632 hospital admissions, western Kenya had the highest number of cases (70%) with the Rift Valley highlands (45%) and Kenyan coast (22%) coming in at 2nd and 3rd, respectively. Each year, approximately 3.5 million new malaria cases and 10,700 deaths are recorded, with those residing in the western part of Kenya being at high risk of contracting the disease [72].

The main vectors of malaria in Kenya include the *An. gambiae* complex (*An. gambiae* ss, *An. arabiensis, An. merus*) and *An. funestus*. These transmit all four *Plasmodium* parasites with *P. falciparum* being the most common, accounting for 99% of all malaria cases within the country [73]. *Anopheles arabiensis* and *An. gambiae* have been shown to coexist sympatrically, especially in the coastal counties Kilifi and Kwale, and Nyanza counties Kisumu and Siaya [74]. However, evidence has shown that over time *An. arabiensis* started becoming the more predominant vector where the two coincide [75,76,77]. *Anopheles merus* is largely distributed within a 25 km radius from the Kenyan coast due to its need to breed in saline water [71,78]. See Figure 1C and Table 2.

Secondary vectors of malaria are also wide spread within the country. For instance, *An. pharoensis* sampled from a rice irrigation scheme in Mwea, central Kenya, constituted 1.3% and 0.68% *P. falciparum* by ELISA (enzyme-linked immunosorbent assay) and dissection methods, respectively [79]. This species has also been reported in western region highlands [80]. *Anopheles nili* has been identified in 26 scattered sites along the coastal areas: Trans Nzoia, Taita and Taveta area, Kaimosi Forest in Vihiga county, Mwea tebere rice plantation, and Thika [81]. *Anopheles coustani* and *An.*

ziemanni sampled from Taveta and western regions, respectively, have also been implicated as potential secondary vectors [14,82]. Regardless of the country-wide distribution of these potential secondary vectors, their actual role in the transmission of malaria parasites in the country is not well known.

Kenya has been stratified into four malaria epidemiological zones with risk factors determined usually by temperature, altitude, rainfall patterns, and malaria prevalence [83]. These zones are: a) The endemic areas in coastal regions and Lake Victoria in the western region. These areas are characterized by altitudes ranging from 0–1300 m above sea level and stable high malaria prevalence throughout the year. b) Highland epidemic prone areas that experience seasonal malaria transmission with considerable year-to-year variations around the western highlands and between 5–20% prevalence. c) Seasonal malaria transmission areas which include arid and semi-arid areas in the northern and south-eastern parts of the country where they experience short episodes of intense malaria transmission during the rainy seasons. d) Low malaria risk areas in the central highlands of Kenya such as Nairobi, where temperatures are too low for the survival of malaria *Plasmodium* parasites [83]. However, with changes in annual climatic patterns and increase in global warming, previously malaria-free regions may soon become endemic areas. For instance, mosquito species such as *An. arabiensis* which were not known to inhabit central Kenya highlands are now being reported [66].

Although the government of Kenya has established various measures that have seen a reduction in malaria prevalence especially in major endemic regions [83], this has not been the case for some counties. In October 2017 and February 2018, malaria outbreaks hit the arid counties of Marsabit, Baringo, and West Pokot leading to numerous hospitalization cases and deaths [84]. This calls for more cohesive policies in the fight against malaria and their respective anopheline vectors.

Apart from the spread of malaria, *Anopheles* spp. are associated with spread of other vector-borne diseases such as: canine heartworm (*Dirofilaria immitis*) common in the Americas [85,86], lymphatic filariasis, and O'nyong'nyong fever. The first case of O'nyong'nyong virus (ONNV) was reported in Uganda in 1959 and spread through the Victoria basin to western Kenya by 1960 [87]. From this time onwards, this virus has repeatedly been isolated from *An. funestus* and *An. gambiae* in different localities within Kenya providing strong evidence that these two mosquito species are indeed the primary vectors [59,88]. Other mosquito-borne viruses detected from *Anopheles* spp. in the country include: Ngari virus (NRIV) in the Tana River (*An. funestus*) [4] and BUNV in Homabay (*An. gambiae*) [20], as summarized in <u>Table 1</u>. Additionally, *An. funestus* and *An. gambiae* are also associated with the spread of lymphatic filariasis in the Kenyan coast from Lamu to Kwale where ecological factors favor its transmission [89,90,91,92].

3.3. Culex

Drab in color, the *Culex* genera comprises up to 1000 species with a fairly weak flying tendency and generally found in all zoogeographical areas ranging from the tropics to cooler temperate regions [96]. Due to their proximity to human settlements, these species are often referred to as common house mosquitoes. They have also been shown to prefer their blood meals from birds that feed at dawn and dusk [97]. Among the multiple *Culex* spp., *Cx. pipiens* complex is the most widely distributed across the globe. It comprises *Cx. Pipiens* L, *Cx. quinquefasciatus* Say, *Cx. pipiens pallens* Coquillett, *Cx. australicus* Dobrotworsky and Drummond, and an autogenous *Cx. pipiens* subspecies *Cx. pipiens molestus* Forskal. Worth noting is that the complex species are not easily distinguished morphologically and can only be separated through a combination of molecular techniques, and behavioral and physiological traits [98,99].

Culex spp. are globally known as vectors of human, avian, and animal pathogens such as lymphatic filariasis primarily vectored by *Cx. pipiens* [100,101]. This disease is caused by a nematode called *W. bancrofti* which is currently in more than 52 countries where 856 million people live in disease endemic regions (South-East Asia (66%) and African regions (33%)) being at high risk [102]. *Culex* spp. are also associated with transmission of viruses, with WNV being the most common. The virus is often amplified by *Cx. pipiens quinquefasciatus* and maintained in a cycle involving humans and birds although other species of the *pipiens* complex have shown competence in spreading the virus [60,97,103,104]. Originally isolated in Uganda in 1937, this virus has spread globally and has been reported in almost all continents [105]. Other pathogens vectored by *Culex* spp. include the Saint Louis encephalitis virus (SLEV) [106], equine encephalitis virus (EEV) [107], dog heartworm (*Dirofilaria immitis*), SNBV, *Plasmodium relictum* (bird malarias) [103], and RVFV. Whether or not Zika virus is vectored by *Culex* spp. has been a topic of debate. Sammy et al. [108] noted the possibility of *Cx. quinquefasciatus* playing a vectorial role in Zika prone areas in Brazil due to their higher abundance compared to *A. aegypti*, which is a commonly known Zika transmission vector. Later on, Amraoui et al. [109] and Huang et al. [110] rejected this possibility. However, a recent study by Phumee at al. [111] evidenced that vertical transmission of Zika virus by *Cx. quinquefasciatus* was indeed possible.

According to the Walter–Reed Bioinformatics Unit (WRBU), Kenya harbors approximately 42 *Culex* spp. (<u>http://www.wrbu.org</u>). The ones with the most public health significance include members of the *Cx. pipiens* complex, *Cx. pipiens* L. and *Cx. quinquefasciatus Say,* whose success in colonizing both urban and rural environs have been attributed to their larval ability to develop in a variety of aquatic habitats [<u>14</u>]. These mosquitoes are highly distributed in Kenyan coast counties and associated with the spread of bancroftian filariasis [<u>90,91,95</u>] and cases of RVF [<u>11,112</u>]. A study by Atoni et al. [<u>17</u>] recorded high numbers of *Cx. quinquefasciatus* in Kwale county, which had abundant vertebrate viruses based on metagenomic analysis. These species have also been implicated in entomologic studies focusing on WNV, SNBV, and NDUV in Garissa county [<u>38,39,60</u>], RVF in Baringo, Garissa, and Nakuru counties [<u>11,19,60</u>], and Usutu virus in Kisumu [<u>4</u>].

Culex univittatus is another common *Culex* spp. distributed within the country. Its first vertical natural transmission of WNV was evidenced in the Rift Valley province in a study by Lanciotti et al. [61]. In Baringo county, it has been implicated as a vector of the RVFV [19] and BUNV in the Lake Victoria region [20]. In an entomologic survey by Lutomiah et al. [22], on the abundance of potential mosquito vectors, this species was found to be highly abundant in Kisumu (37%) and Naivasha (21.6%), even though no pathogen screening was conducted.

Other disease important *Culex* spp. in the country include *Cx. bitaeniorhynchus*, *Cx. cinereus*, *Cx. poicilipes*, *Cx. rubinotus*, *Cx. Vansomereni*, and *Cx. zombaensis* in which various etiological pathogens have been isolated as summarized in <u>Table 1</u>. Vector competence involving some of these species has also been witnessed. For instance, in a study by Turell et al. [23], *Cx. zombaensis* collected around the Lake Naivasha region showed high competence for RVFV with an 89% infection rate.

3.4. Mansonia

The *Mansonia* mosquito species belong to the sub-family Culicinae and are normally characterized by their large body size and asymmetrical wing scale structure with sparkling on their wing veins and legs. They show some resemblance with *Aedes, Culex,* and *Coquillettidia* mosquito but the simplicity of their tarsal claws in structure and a truncated abdomen in females sets it apart [113]. Furthermore, *Mansonia* preferably breed in ponds and permanent waters with aquatic plants where larvae can burrow into the dead plants at the bottom or cling to the roots of the live ones [114]. Its genera comprise 25 globally distributed species in two subgenera: *Mansonioides* (10 species) and *Mansonia* (15 species).

The distribution of these species comes with increased vectorial role of human etiologies. *Mansonia titillans*, a common species in the Americas, is a known vector of Venezuelan equine encephalitis [<u>115</u>]. Recently, Argentina detected SLEV and BUNV for the first time from this species, proving its competent nature to transmit the viruses [<u>116</u>]. In south-east Asia, *Mn. bonneae* and *Mn. dives* have been linked to various cases of Brugian and lymphatic filariasis [<u>117</u>]. Africa, as one of the most affected regions of vector-borne diseases, harbors two main and probably the most common *Mansonia* spp. (*Mn. africana* and *Mn. uniformis*). These species have been reported to vector *W. bancrofti*'s lymphatic filariasis [<u>118</u>,119], RVF [<u>34</u>], Zika [<u>120</u>], among other diseases. Kenya is not an exception, as studies have shown that the two *Mansonia* spp. are distributed within the country and harbor pathogens of human health importance [<u>4</u>,11,36,40,60].

LaBeaud et al. [60] showed that *Mansonia* mosquitoes could have significantly contributed to the vectorial transmission of RVF in the 2006–2007 outbreak in the north-eastern region. Out of the 12,080 mosquitos collected during the study, 682 were identified to be *Mansonia* spp. Further investigation based on molecular screening identified three out of eight pools of *Mansonia* spp. to be positive for RVFV. *Mansonia*, like many other female mosquitoes, feed on vertebrate blood for egg development. These vertebrates can lead to amplification of a disease in case of outbreaks. Virus screening from bloodfed *Mansonia* spp. by Lutomiah et al. [40] during this same RVF outbreak, showed that goats and sheep were the greatest amplifiers of the virus.

Owing to the fact that *Mn. uniformis* and *Mn. africana* preferably breed around permanent water bodies, flooded lagoons, and swamps, they have been shown to dominate lake regions in the country. A study by Lutomiah et al. [22] on the abundance of mosquito vectors in various ecological zones in the country revealed high abundance of *Mansonia* spp. in Baringo (71%) and Kisumu (23%), the home places of Lake Baringo and Lake Victoria. Interestingly, it is here (Baringo) that positive strains of RVFV and NDUV had earlier been detected in *Mn. uniformis* and *Mn. africana* [11,36]. This was further evidenced by Ajamma et al. [18], where these species dominated in abundance in the islands and mainlands of both Lake Victoria and Lake Baringo.

Even though Zika virus and *W. bancrofti* have been detected from *Mansonia* spp. in some African states [<u>118</u>,<u>120</u>], this has not been the case in Kenya. In a study at the Kenyan Tana delta on vectorial potential of *Mansonia* spp. to transmit *W. bancrofti*; all 236 species collected tested negative on PCR (polymerase chain reaction) analysis [<u>121</u>]. However, the study recommended further assessment to be done by infecting *Mansonia* spp. experimentally with *W. bancrofti* microfilaria to identify whether or not they can support development of microfilaria up to the infective stages.

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