



Environmental changes and emerging vector-borne diseases: A review

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ABSTRACT : Vector-borne infectious diseases are resurging as a result of changes in public health policy, demographic and societal changes, insecticide and drug resistance, shift in emphasis from prevention to emergency response, genetic changes in pathogens in the last two decades of the twentieth century. Each environmental change, whether occurring as a natural phenomenon or through human intervention, changes the ecological balance and context within which disease hosts or vectors and parasites breed, develop, and transmit disease. Climate changes also can influence the emergence and reemergence of these diseases which are malaria, dengue, yellow fever, plague, filariasis, louse-borne typhus, Lyme disease, trypanosomiasis, leishmaniasis and viral diseases. West Nile virus is just the latest example of this type of invasion by exotic virus. This paper provides an overview of the distribution, reasons of emergence or reemergence of the important vector-borne diseases throughout the world.

Keywords : Malaria, Yellow fever, Dengue, Plague, Typhus, Arbovirus

INTRODUCTION

During the history of human cultural evolution, population dispersal around the world and subsequent inter-population contact and conflict there have been several distinct transitions in the relationships of *Homo sapiens* with the natural world. Each of these transitions in human ecology and inter-population interaction has profoundly changed the patterns of infectious disease. In the first transition, human settlements allowed countless novel strains of bacteria and viruses to make the jump from domesticated herds of naturally gregarious animals to humans. Anthropological research in Africa has shown that malaria-protective sickle-cell trait arose when iron tools and slash-and-burn agriculture were introduced, presumably by creating new breeding sites for *Anopheles gambiae*, the major mosquito vector for falcifarum malaria (Michael, 2001).

The second great transition occurred from around 2500 to 1000 years BC with travel and military movements between the contiguous land masses of Europe, Northern Africa and Asia. And the third great transition began around five centuries ago and entailed the trans-oceanic spread of disease.

These three great historical transitions were processes of equilibration between first humans and animal species and later, among regional human populations. As new ecological niches were created by changes in human cultural practices, microbes exploited those niches. As new contacts were made between previously isolated civilizations, infectious diseases have been shared. In today's globalizing world, populations everywhere are becoming interconnected economically, culturally and physically, enhancing the mixing of people, animals, and microbes from all geographical areas.

In today's world, no country is safe from infectious diseases. Human mobility has escalated dramatically in volume and speed between and within countries. Long distance trade facilitates the geographical redistribution of pests and pathogens. Changing social, cultural and environmental conditions have been related to the new distribution pattern of the diseases, emphasizing vector-borne diseases, to clarify the above question.

Historically, vector-borne diseases such as malaria, dengue, yellow fever, plague, filariasis, louse-borne typhus, *trypanosomiasis*, and *leishmaniasis* were responsible for more human disease and death in the 17th and the early 20th centuries. (Gubler, 1998). After successful control programs by the 1960s, vector-borne diseases were no longer considered major public health problems outside Africa. However, the benefits of vector-borne disease control programs were short-lived, and in the last thirty years, there has been a dramatic global resurgence of infectious diseases. Some important diseases were recognized for the first time, including HIV/AIDS, Hantavirus pulmonary syndrome, Ebola, Lyme disease and *ehrlichiosis*. Equally important was the rising incidence of many recognized diseases that had been effectively controlled since 1960s. New combinations of diseases have also been noted, such as the appearance and spread of co-infections of HIV virus and *leishmaniasis*. By 2004, vector-borne diseases like malaria and dengue hemorrhagic fever are among the most serious public health problems in countries where there are no properly-trained vector biologists in the Ministries of Health (WHO, 2004).

Although the reasons behind this dramatic resurgence of arboviral diseases are complex, Gubler (2002) listed the following factors :

- (i) Demographic changes including global population growth, population movements, and unplanned and uncontrolled urbanization;
- (ii) Societal changes such as human encroachment on natural disease foci, modern transportation, containerized shipping;
- (iii) Agricultural changes including changes in land use, irrigation systems, deforestation;
- (iv) Changes in pathogens due to increased movement in humans and animals and genetic changes leading to increased epidemic potential;
- (v) Change in public health, including lack of effective vector control, deterioration of public health infrastructure to combat vector-borne diseases, surveillance of disease, prevention programs and possible climate change (Gubler, 2002).

It is clear that certain demographic and societal changes in the past 30 years had a major impact on the ecology of vector-borne diseases. Modern transportation ensures faster and increased movement of humans, animals, and commodities and their pathogens between regions and population centers of the world. For example, in the past twenty years, four exotic mosquito species have been introduced and established in the United States; three of these are potential vectors of local diseases (Russel, 2001). *Aedes albopictus* (Asian tiger mosquito) of Asian origin, first spread to the America and African continents, and is now established in North and South America, Africa, Oceania and Europe. It was first detected in Europe in 1979 in Albania, after which it has been reported from many different countries (WHO, 2004). By late 2001, *Aedes albopictus* had rapidly become the most important pest species in northern Italy. In 1996, another introduced mosquito species *Aedes atropalpus* was discovered in the Ventao region of northern Italy. More recently, *Ochlerotatus japonicus*, an Asian mosquito, has been reported from Normandy, France where it was found breeding in tires, like the above *Aedes* species (Schaffner, *et al.*, 2001).

There is an increasing trend of Tick-borne *encephalitis* cases in Europe, partly due to changes in human behavior that bring more people into contact with infected ticks. Fluctuations in mean temperature values also effect the distribution of many tick species in Northern Europe. Dipteran larvae, a cause of myiasis, are often found on travelers returning from tropical countries. Cockroaches are ubiquitous pest aboard ships and aircraft, and exotic species are quite often transferred from one country to another (Gubler, 2002). The high rates of transmission by different vectors to different vertebrate hosts in new geographic areas can result in selective pressures that lead to genetic changes in the pathogen. These new strains of virus may have greater epidemic potential and virulence. For example, the unique susceptibility of young domestic geese in Israel during 1997-2000 to West Nile virus and the isolation of similar strains from migrating White storks in Israel and Egypt suggest that the recent isolates are more pathogenic for certain avian species (Malkinson and Banet, 2000).

Deforestation provides new ecological niches and conditions for proliferation of newly arriving and/or adaptive vectors and their parasites. The replacement of forests with crop farming, ranching and raising small animals can create a supportive habitat for parasites and their host vectors. New settlers to a deforested area are particularly vulnerable as they lack immunity to the zoonotic parasites endemic to the area. In adapting to changed environmental conditions, including reduction of non-human population and increased human population, some vectors display a conversion from a primarily zoophyllic (bites to animal) to primarily anthropophilic (bites to human) orientation. Water control projects can create new breeding habitats for mosquitoes, snails, their larvae and their parasites. The construction of new roads provides access for new human, livestock, vector and parasite populations (Patz *et al.*, 2002).

The International Council of Scientific Unions Intergovernmental Panel of Climate change, established by the World Meteorological Organization and the United Nations Environmental Program, have estimated that by the year 2100, average global temperatures will have risen by between 1.0°C to 3.5°C. Important ecological changes may come in future due to the global warming. The distribution and seasonality of diseases that are transmitted by coldblooded insects or ticks are likely to be affected by climate change (Gubler, 2002; Patz *et al.*, 2002). Changes in climate is responsible for the recent resurgence of vector-borne diseases

The incidence of mosquito-borne diseases, including malaria, dengue, and viral encephalitis are among those diseases most sensitive to climate (Epstein, 2000). Climate change would directly affect the transmission of the diseases by shifting the vectors geographic range and increasing reproductive and biting rates and by shortening the pathogen incubation period. Human susceptibility to infections might be further compounded by malnutrition due to crop failure caused by climate stress (Patz *et al.*, 2002). Temperature can affect both the distribution of the vector and the effectiveness of pathogen transmission through the vector. Temperature can affect survival of the vector, the rate of vector population growth, feeding behavior, susceptibility of the vector to pathogens, incubation period of the pathogen, seasonality of vector activity, and the seasonality of pathogen transmission. For example, the relation between El-Nino events and increased malaria risk is partly due to increased temperature and partly due to increased rainfall leading to increased mosquito breeding sites because of surface water collections (Hunter, 2003). Decreased rainfall has been shown to be associated with epidemics of SLE (St Louis encephalitis) when the vector *Culex pipiens* breeds effectively in urban drainage systems. Unusual abundance of some vectors, such as ticks or changed behavior of other pests, such as higher anthropophily of *C. pipiens* may in some areas is attributed to climate changes in Europe (Rettich, 2002).

The northern range of the occurrences of endemic malaria has been estimated to coincide with a summer isotherm of 16°C. Climatic factors that increase the inoculation rate of Plasmodium parasites, as well as the breeding activity of *Anopheles* mosquitoes, are considered the most important cause of epidemic outbreaks of malaria in no endemic areas. According to malaria models under climate change scenarios, the risk of malaria epidemics would rise substantially in tropical and temperate regions (Hunter, 2003).

MALARIA

Malaria is the most important tropical disease with more than half of the world's population living in areas of risk and with an estimated 300-500 million cases and 1.5-2.7 million deaths each year. It has been on the rise in many parts of Africa, with the mortality in young children almost doubling from the 1980s to the 1990s (Gubler, 2002).

Widespread drug resistance in the parasites and insecticide resistance among *Anopheles* mosquito vectors have complicated malaria control. Malaria is endemic in all parts of India except at elevations above 1800 m and in some coastal areas (Sharma, 1996). The principal vectors, which cause malaria in most parts of India, are the *An. culicifacies* – a rural vector, *An. stephensi* – an urban vector and *An. fluviatilis* a resident of hilly-forested areas. Periodic epidemics of malaria occur every five to seven years (Sharma, 1996). As recent as in the year 1998, about 20,000 people and an estimated 577,000 DALYs (disability-adjusted life years) were lost due to malaria in India (WHO, 1999).

Malaria is also the most common imported disease in the United States, where *Anopheles* mosquito vectors still exist (Gubler, 1998). Until after the end of World War II, malaria was endemic throughout much of southern Europe. The Balkans, Italy, Greece and Portugal were particularly affected, although seasonal epidemics or outbreaks occurred as far north as Scandinavia, Finland, Norway and southern Sweden. The area of malaria distribution in Europe peaked at the beginning of the twentieth century. Soon after the war, intensive control measures were initiated and by 1970 the WHO declared malaria eradicated from Europe. However, populations of potential *Anopheles* vectors of malaria remain high in many countries of the continent and their presence poses the risk of renewed transmission. The WHO Regional Office for Europe reported a total of 15,528 cases of imported malaria in Europe in the year 2000 (Gubler, 2002). Local transmission has also frequently occurred in Europe in the form of airport malaria. This refers to the transmission of malaria as a result of the inadvertent transport of live, malaria-infected mosquitoes aboard aircraft arriving from tropical malaria-endemic countries (Gratz, 2000). The most serious problem of resurgent malaria is in the newly independent states (Azerbaijan, Tajikistan) and in Turkey; due to an influx of refugees from malaria endemic areas, the breakdown in health services, and the lack of vector control measures in most of these states, as well as in the failure to carry out

adequate malaria surveillance and control measures (Alten, 2001).

LEISHMANIASIS

Leishmaniasis, which comes next after malaria is a parasitic infection transmitted by the bite of an infected female sand fly whose hosts are animals, such as dogs or rodents, or human beings. Leishmaniasis is a highly focal disease with widely scattered foci. The parasite may survive for decades in asymptomatic infected people, who are of great importance for the transmission since they can spread visceral leishmaniasis indirectly through the sand flies. The parasites can also be transmitted directly from person to person through the sharing of infected needles which is often the case with the *Leishmania*/HIV co-infection (Desjeux and UNAIDS, 1998). The disease has four main forms, depending on the parasite species and the cellular immune system of the patient:

Cutaneous leishmaniasis produces skin lesions mainly on the face, arms and legs. Although this form is often self-healing, it can create serious disability and permanent scars. After recovery or successful treatment, cutaneous leishmaniasis induces immunity to re-infection by the species of *Leishmania* that caused the disease.

Diffuse cutaneous leishmaniasis is difficult to treat due to disseminated lesions that resemble leprosy and do not heal spontaneously. This form especially is related to a defective immune system and it is often characterized by relapses after treatment.

Mucocutaneous leishmaniasis, also called 'espundia' in South America, causes disfiguring lesions to the face; it destroys the mucous membranes of the nose, mouth and throat. Reconstructive surgery of deformities is an important part of therapy.

Visceral leishmaniasis, also known as 'Kala azar', is characterized by irregular fever, weight loss, swelling of the liver and spleen and anaemia. It is the most severe form of Leishmaniasis, and is usually fatal if left untreated. The incubation period can be months or years and, unlike the cutaneous forms of leishmaniasis, it involves the internal organs. After treatment and recovery, patients may develop chronic cutaneous leishmaniasis that requires long and expensive treatment.

Leishmaniasis has a long history. Designs on pre-Colombian pottery and the existence of thousand-year old skulls with evidence of leishmaniasis prove that the disease has been present in the Americas for a long time. Today, an estimated 12 million cases of leishmaniasis exist worldwide with an estimated number of 1.5-2 million new cases occurring annually; 1-1.5 million cases of cutaneous leishmaniasis and 500 000 cases of visceral Leishmaniasis (WHO, 1993).

The geographical distribution of leishmaniasis is restricted to tropical and temperate regions, the living area of the sand fly. The leishmaniasis is considered to be endemic in 88 countries (16 developed countries and 72 developing countries) on four continents. 90% of cases with

cutaneous forms of leishmaniasis occur in Afghanistan, Algeria, Brazil, Iran, Peru, Saudi Arabia and Syria, while 90% per cent of visceral leishmaniasis cases are found in Bangladesh, Brazil, India, Nepal and Sudan (WHO, 1996).

Historically, African sleeping sickness transmitted by the tsetse fly, has been a major impediment to the social and economic development of Central and East Africa. In the past 20 years, major epidemics have occurred in East and Central Africa, mainly because control programs were disrupted by war (Gubler, 1998).

Tick-borne diseases are also sensitive to climatic conditions but favor cooler temperatures (Petz *et al.*, 1996). Documented increases in the incidence of Tick-borne *encephalitides* over the last decades have been reported from many countries in Europe, as the result of increased densities of vector tick species (Gubler, 2002).

LYME DISEASE

Lyme disease, transmitted by *Ixodes scapularis* or *pacificus* ticks is caused by *Borrelia burgdorferi*. Discovered in the United States in 1975, the disease has continued to increase in incidence and geographic distribution since national surveillance was started in 1982. It is the most commonly reported vector-borne disease in Europe and North America; its incidence is clearly increasing, and new species of *Borrelia* are continuously being found. Among the reasons for this increase is an ecological change such as reforestation that favors greater densities of tick vector populations and increased exposure of persons to tick bites when visiting endemic areas (Nurdan, 2005).

PLAGUE

Plague, is the original emerging disease, having caused major pandemics. Like many other vector-borne diseases, plague was controlled with antibiotics, insecticides and rat control in the latter half of the 20th century. In recent years, however, epidemic plague has resurged, most notably in Africa, with an average of nearly 3,000 cases reported annually (Gubler, 1998).

ONCHOCERCIASIS

Onchocerciasis or river blindness vector requires fast-flowing water for successfully reproduction and the adult vector can be spread by wind. According to the increased temperature and precipitation levels, black fly populations may increase by as much as 25% at their current breeding sites (Patz *et al.*, 2000).

SCHISTOSOMIASIS

Among other vector-borne diseases affected by climate changes, *Schistosomiasis* has increased in prevalence in arid warm regions primarily from expansion of irrigation systems where snails serve as the intermediate host (Hunter, 2003).

ARBOVIRAL DISEASE

Viral diseases transmitted by blood-feeding arthropods (arboviral diseases) are among the most important of the emerging infectious diseases public health problems facing

the world at the beginning of the third millennium (Gratz, 2003). These include Dengue, Dengue hemorrhagic fever, Yellow Fever, Japanese Encephalitis, West Nile Fever, Kyasanur Forest disease, Venezuelan equine encephalitis, Epidemic poly arthritis, Barmah forest disease and Mayaro Rift Valley Fever, Oropouche, and Crimenean-Congo hemorrhagic fever. There are currently 534 viruses registered in the International Catalogue of Arboviruses, of which 40% are known or probable arboviruses; 25% have caused documented illness (Gubler, 1998). Although some arboviruses that cause human disease such as Barmah Forest disease in Australia have been newly recognized, the greatest problem is by far the viruses mentioned above. Arboviruses have a worldwide distribution. But, in the last two decades, the geographic distribution of both vectors and viruses has expanded globally. The geographic distribution of each Arbovirus is restricted by the ecological parameters governing its transmission cycle. These cycles are usually silent and undetected in nature until some environmental change allows the virus to escape the primary cycle via a secondary vector or vertebrate host, or when humans invade or encroach on the nidus of infection. Mosquitoes are by far the most important vectors of arboviruses, and birds and rodents are the most important vertebrate reservoir hosts. These zoonotic viruses circulate in nature either in an *Aedes-mammal*, *Anopheles-mammal* or *Culex-bird* transmission cycles (Lundstrom, 1999).

DENGUE

The Dengue viruses are the most widespread geographically of the arboviruses and are found in tropical and subtropical areas where 2.5-3 billion people are at risk of infection with 50-100 million cases of Dengue fever and 500,000 cases of DHF. It is an old disease that distributed worldwide in the tropics during the 18th and 19th centuries when the shipping industry and commerce were expanding. Both the principal mosquito vector, *Aedes aegypti* and the viruses responsible for Dengue fever was spread via sailing ships. In those times because of the slow mode of transportation, epidemics were infrequent with intervals of 10-40 years. The global epidemiology and transmission dynamics of dengue viruses were changed in Southeast Asia during World War II. Today, dengue fever causes more illness and death than any other Arbovirus disease of human (Solomon, 2001). New dengue virus strains and serotypes will likely continue to move between areas where *Aedes aegypti* occurs in infected air travelers, resulting in increased frequency of epidemic activity and increased incidence of Dengue hemorrhagic fever. Although funding for dengue vaccine development has been sparse, good progress has been made over the past ten years (Kinney and Huang, 2001).

Currently, Dengue viruses are being transmitted in the tropics between 30° north and 20° south latitude, since frosts or sustained cold weather kills adult mosquitoes and overwintering eggs and larvae. Warming trends therefore

can shift vector or disease distribution to higher latitudes or altitudes as was observed in Mexico when dengue reached an altitude of 1700m during the unseasonably warm summer in 1998. Temperature also affects the transmission dynamics of dengue, because warmer temperatures reduce the size of *Aedes aegypti* individuals, smaller adults must feed more frequently to develop egg batches, resulting in double feeding (Hunter, 2003).

YELLOW FEVER

Yellow fever is an old, well-known disease that caused major epidemics in the Americas and in Africa from 17th to 20th centuries. Like Dengue and DHF, these were primarily urban epidemics transmitted by *Aedes aegypti* (Gubler, 2002).

WEST NILE VIRUS (WN VIRUS)

West Nile virus is the latest example of invasion by exotic viruses. West Nile virus was first isolated in 1937 from the blood of a febrile patient in the West Nile district of Northern Uganda. During 1994-2000, epidemics of WNV meningoencephalitis occurred at a new alarming rate in North Africa, Europe, North America and the Middle East (Deubel *et. al.*, 2001; Campbell *et al* 2002).

West Nile virus was first detected in the United States in September 1999 during the investigation of an outbreak of encephalitis in humans in New York City (White *et al.*, 2001). The dramatic appearance of epidemic meningoencephalitis in New York City in 1999 is an unsettling reminder of the ability of viruses, including arboviruses, to jump continents and hemispheres. WNV rapidly spread across the United States and invaded California during 2003 (Reisen, *et. al.*, 2001). The continued movement west into large population centers in California increases the urgency of continued efforts to develop intervention methods (Campbell *et. al.*, 2002).

WNV is mainly transmitted by *Culex mosquitoes*. Compared with most arboviruses, however, WNV has been identified from an exceptionally broad range of species. *Aedes albopictus* and *Culex erraticus* were recorded as the vector species for the first time in New York State (Kulasekera, *et. al.*, 2001). After that, in North America only, the virus has been identified in 43 different species of mosquito. With such a broad vector range, expansion of the geographic distribution of WNV into new areas and its establishment seems inevitable (Granwehr, 2004).

It was recorded that Israeli researchers in the 1950s were the first to characterize the clinical presentation of WN fever, but by the end of the 20th century, WN virus infection was an almost forgotten disease in Israel. The vectors of WN virus in Israel were observed as *Culex* species, *Culex Pipiens* and *Culex Perexiguus* (Weinberger *et. al.*, 2001).

Birds are the natural reservoir hosts for WN virus, which has been shown to infect at least 111 bird species in North America only (Campbell *et. al.*, 2002). In North America, WN virus seems to be particularly virulent in species belong to

the family Corvidae (*e.g.*, crows and jays) (Komar 2000) and these have a central role in dead-bird based surveillance programmes for detecting and tracking the virus (Campbell *et al.*, 2002) Bird deaths due to WN virus are unusual outside North America, with the exception of deaths of geese in Israel and pigeons in Egypt (Bernard *et. al.*, 2000) Whether high avian death rates in the United States are due to higher virulence of the circulating strains or to higher susceptibility in North American birds require further evaluation (Petersen and Roehrig, 2001). Then it was recorded that the unique susceptibility of young domestic geese in Israel in 1997-2000 to WNV and the isolation of similar strains from migrating White storks in Israel and Egypt suggest that the recent isolates are more pathogenic for certain avian species. Infected birds have recently been found in the Caribbean and the infection will no doubt continue to spread through the Americas (Gubler, 2002).

Transportation of WN virus strains between different areas by migratory viraemic birds along established flyways is probably a common occurrence. In the eastern hemisphere, WN virus is thought to be regularly introduced in Mediterranean and European countries by birds (Rappole *et. al.*). Campbell *et. al.*, 2002 emphasized that WN virus will almost certainly continue to spread into the contiguous western parts of the USA over the next years, primarily via the movement of viraemic birds.

A broad range of mammalian species are also susceptible to natural or experimental infection with WN virus. In the USA during 1999-2001, nine mammalian species (human beings, horses, cats, rabbits, skunks, squirrels, chipmunks and two species of bats) were found to be naturally infected with WN virus. The role, if any, that mammals play in the WN virus transmission cycle is unknown (Komar, 2003).

No human vaccine for WN virus is currently available. So, effective prevention of the disease depends on integrated arboviral surveillance and vector mosquito control programmes to reduce the density of vector species, including those that might serve as a bridge from birds to human beings. Cases were also of WNV transmission transplacentally by transfusion, by lactation, by organ transplants as new routes of transmission (Granwehr *et al.*, 2004). Werner pointed that the scope of infectiology keeps widening, while the threat of bioterrorism cannot be neglected (Werner, 2001).

CONCLUSION

Without sustained vector control in urban areas, even the world's affluent cities are at risk for epidemic arboviral diseases. We do not know how many potential organisms will share our planet with us that are awaiting the right conditions for their chance at stardom by becoming emerging infectious diseases. Millions of deaths occur annually due to these vectors all over the world and reemergence of

certain vectors are posing a potential threat and challenge to the humanity. Control measures for vector-borne diseases are important, these include undertaking personal protective measures by establishing physical barriers such as house screens and bed nets; wearing appropriate clothing (boots, apparel that overlaps the upper garments, head nets, etc.); and using insect repellents. Environmental modification to eliminate specific breeding areas or chemical biological control measures to kill arthropod larvae or adults may also be undertaken. Areas such as ports and airports should be rigidly monitored, with control measures utilized to prevent important arthropod disease vectors from entering the country. The department of Health and Family Welfare should launch curative, preventive and educative role to prevent and control vector borne diseases. Emphasis should be laid on new strategies for prevention and control of vector-borne diseases. Integrated Vector Management approach will reinforces linkages between health and environment.

REFERENCES

- Alten B, Caglar S.S. (2001). Ozer N. Malaria and its vectors in Turkey. *The European Mosquito Bulletin.*, **7**: 27-33,
- Aranda C, Sanchez-Seco M.P, Escosa, *et. al.*, (2003). Arbovirus mosquito surveillance in Spanish Wetlands. 14th European Conference of the Society for Vector Ecology. Bellinzona, Switzerland., 3-6.
- Bernard K.A, Maffei J.G, Jones S.A, *et. al.*, (2001). West Nile Virus Infection in birds and mosquitoes, New York State, 2000. *Emerging Infectious Disease.*, **7**: 679-685.
- Campbell G.L, Marfin A.A, Lanciotti, R.S, *et. al.*, (2002). West Nile virus. *The Lancet.*, **2**: 519-529,
- Desjeux P. and UNAIDS. (1998). *Leishmania and HIV in gridlock.* World Health Organization.
- Deubel V, Fiette L., Malkinson, M. *et. al.*, (2001). West Nile virus: a deadly emerging virus. 3rd International Congress of Vector Ecology. Barcelona, Spain. 16-21.
- Epstein P.R. (2000). Is Global Warming Harmful to Health? *Scientific American.* August, 50-57,
- Granwehr B.P, Lillibridge K.M, Higgs S, *et. al.*, (2004). West Nile virus: where are we now? *The Lancet*, **4**: 547-556.
- Gratz N.G. (2000). Steffen R, Cocksedge, W. Why aircraft disinsection? *Science Direct. Bulletin of the World Health Organization.*, **78**(8): 995-1004.
- Gratz N.G. (2003). The Global burden of the arboviruses. 14th European Conference of the Society for Vector Ecology. Bellinzona, Switzerland., 3-6.
- Gubler D.J. (1998). Dengue and Dengue Hemorrhagic fever. *Clinical Microbiology Reviews.*, **11**: 480-496.
- Gubler D.J. (1998). Resurgent vector-borne diseases as a global health problem. *Emerging Infectious Diseases.*, **4**: 442-450.
- Gubler D.J. (2002). Epidemic dengue / dengue hemorrhagic fever as a public health, social and economic problem in the 21st century. *Trends in Microbiology*, **10**: 100-103.
- Hunter P.R. (2003). Climate change and water-borne and vector-borne diseases. *Journal of Applied Microbiology*, **94**: 35-465.
- Kinney R.M, Huang C.Y.H. (2001). Development of new vaccines against dengue fever and Japanese encephalitis. *Intervirolgy*, **44**: 176-197.
- Komar N. (2003). West Nile virus: epidemiology and ecology in North America. *Adv. Virus Res.*, **61**: 185-234.
- Kulasekera V.L, Kramer L, Nasci R.S, *et. al.*, (2001). West Nile Virus Infection in Mosquitoes, Birds, Horses and Humans, Staten Island, New York, 2000. *Emerging Infectious Diseases*, **7**: 722-725.
- Lanciotti R.S, Kerst A.J, Nasci R.S, *et. al.*, 2000. Rapid Detection of West Nile Virus from Human Clinical Specimens, Field-Collected Mosquitoes and Avian Samples by TaqMan Reverse Transcriptase-PCR Assay. *Jr. of Clinical Microbiology*, **38**: 4066-4071.
- Lundstrom J.O. (1999). Mosquito-borne viruses in Western Europe: A review. *J. of Vector Ecology*, **24**: 1-39.
- Malkinson M, Banet C. (2002). The role of birds in the ecology of West Nile virus in Europe and Africa. *Current topics Microbiology Immunology*, **267**: 222-309.
- McMichael A.J. (2001). Human Culture, Ecological Change and Infectious Disease: Are We Experiencing History's Fourth Great Transition? *Ecosystem Health*, **7**: 107-115.
- Nurdan, Z. (2005). Emerging Vector-borne Diseases in a Changing Environment. *Turk J Biol.*, **29**: 125-135.
- Patz J.A, Epstein P.R, Burke T.A, *et. al.*, (1996). Global Climate Change and Emerging Infectious Diseases. *The Journal of the American Medical Association.*, **275**: 217-223.
- Patz J.A, Graczyk T.K, Geller N, *et. al.*, 2000. Effects of environmental change on emerging parasitic diseases. *International Journal for Parasitology*, 1-11.
- Petersen L.R, Roehrig J.T. (2001). West Nile Virus: A Reemerging Global Pathogen. *Emerging infectious diseases*, **7**: 611-614.
- Rappole J.H, Derrickson S.R, Hubalek Z. (2000). Migratory birds and spread of West Nile virus in the Western Hemisphere. *Emerging Infectious Diseases.*, **6**: 319-28.
- Reisen W.K., Lothrop H.D., Chiles, R.E., *et. al.*, (2004). Invasion of California by West Nile Virus. *Emerging Infectious Diseases.*, **10**: 1369-1378.
- Rettich F. (2002). Vector Ecology and vector control-public health implications. *VIIth European Congress of Entomology*, 7-13 October, Thessaloniki, Greece.
- Russel R.C. (2001). Introduction and establishment of exotic mosquitoes in Australia and New Zealand. 3rd International Congress of Vector Ecology. Barcelona, Spain. 16-21 September.
- Schaffner F, Bouletreau B, Guillet B, *et. al.*, (2001). *Aedes albopictus* (Skuse, 1894) established in metropolitan France. *The European Mosquito Bulletin*, **9**: 1-3.
- Sharma, V.P. (1996). Re-emergence of malaria in India. *Indian J. Med. Res.*, **103**: 26-45.
- Solomon T, Mallewa M. (2001). Dengue and other emerging flaviviruses. *The J. of Infection*, **42**: 104-115,
- Weinberger M, Pitlik S.D, Gandacu D, *et. al.*, (2001). West Nile Fever Outbreak, Israel, 2000. *Emerging Infectious Diseases*, **7**: 686-691.
- Werner G.H. (2001). The worldwide challenges of ÒnewÓ or reemerging communicable diseases at the dawn of the 21st century. *Ann. Pharm. Fr.*, **59**: 246-277.
- White, D.J, Kramer L.D, Backenson P.B, *et. al.*, (2001). Mosquito surveillance and Polymerase Detection of West Nile Virus, New York State. *Emerging Infectious Diseases*, **7**: 643-649.
- World Health Organization (1993). The Leishmaniasis, CTD/MIP/WP.93.8.
- World Health Organization (1996). Fighting Diseases, Fostering Development. WHO, Geneva, Switzerland.
- World Health Organization (1999). Making a Difference. The World Health Report, Geneva.
- World Health Organization (2004). The Vector-Borne Human Infections of Europe. WHO, Geneva, Switzerland.