#### REVIEW ARTICLE

#### FOSSIL-FUEL POLLUTION AND CLIMATE CHANGE

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# Climate Change and Vectorborne Diseases

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HE EFFECTS OF CLIMATE CHANGE ARE WIDESPREAD AND RAPIDLY INtensifying and are largely driven by greenhouse-gas emissions from burning fossil fuels. Global mean temperatures have already increased by 1.1°C since 1900, with most of the change having occurred in the past 50 years. The extent of change is most extreme in highland and polar regions (Fig. 1), and temperatures in tropical regions are creeping closer to the thermal limits of many organisms. Given the current policies and actions, a warming of 2.5°C to 2.9°C or more by the end of this century is expected.

Warming and other manifestations of climate change — including changes in precipitation, with increased flooding in some areas and drought in others — have important implications for vectorborne diseases through their effects on pathogens, vectors, and hosts, as well as on our ability to prevent and treat these diseases (Fig. 2). Yet attributing changes in the distribution and frequency of vectors and diseases to climate change is challenging because other factors, including land-use changes,<sup>3</sup> the abundance of reservoir hosts,<sup>4</sup> and control measures,<sup>5</sup> also contribute to these changes. Furthermore, it may be difficult to distinguish between natural climate variability and human-influenced change,<sup>6</sup> although scientific techniques to do so are emerging. Despite these complexities, it is clear that the components of vectorborne disease systems, including pathogens, vectors, and reservoir hosts, are highly responsive to the varied environments they inhabit and that observed changes in the rates of vectorborne diseases at given locations are often associated with concomitant changes in the local climate.

For example, warming temperatures affect the behavior, physiologic characteristics, and life history of both vectors and pathogens as well as the abundance and behavior of reservoir hosts and definitive hosts. The interactions among temperature, vector, and pathogen can change the risk of human-to-human disease spread and of spillover to humans from reservoir hosts. Thermal performance curves illustrate the ways in which temperature affects the physiological traits of pathogens, vectors, and reservoir hosts, which determine the rate of disease spread in a susceptible population. These curves are commonly used to predict the potential effects of rising temperatures resulting from climate change on vectorborne disease systems.<sup>7</sup> Curves for individual components of a disease system must overlap in order for transmission to occur. Thermal adaptation, acclimation to a warming climate, or both can potentially shift thermal performance curves and thermal tolerance limits, with important implications for expansion of the geographic range of certain diseases. Depending on their ability to adapt, vectors may no longer carry certain pathogens or may carry new ones as climate-mediated ecosystem changes bring different pathogens, vectors, and reservoir and human hosts together.8

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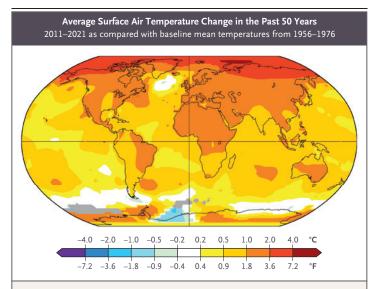


Figure 1. Mean Surface Air Temperatures.

Shown are mean surface air temperatures from 2011 to 2021 as compared with baseline mean temperatures from 1956 to 1976. Adapted from the National Aeronautics and Space Administration Goddard Institute for Space Studies (https://data.giss.nasa.gov/gistemp/maps/index\_v4.html).

# CLIMATE-SENSITIVE VECTORBORNE DISEASES

The Intergovernmental Panel on Climate Change reported with high confidence that the prevalence of vectorborne diseases has increased in recent decades and that the prevalences of malaria, dengue, Lyme disease, and West Nile virus infection in particular are expected to further increase during the next 80 years if measures are not taken to adapt and strengthen control strategies.¹ Table 1 describes these and additional examples of vectorborne diseases that are responding to a changing climate. Additional details are provided in Figure S1 in the Supplementary Appendix, available with the full text of this article at NEJM.org.

### MALARIA

Malaria, which is caused by plasmodium species and is transmitted between humans by infected female anopheles mosquitoes, is the most deadly and most studied climate-sensitive vector-borne disease. Despite control efforts, more than 600,000 deaths were attributed to malaria in 2020, predominantly among pregnant women and young children in Africa.<sup>26</sup> In many regions, malaria is a seasonal or epidemic disease that

responds to short-term changes in rainfall, humidity, and temperature. Temperature increases of 0.2°C per decade in the highlands of Colombia and Ethiopia have been associated with the spread of malaria to higher elevations in these countries. The frequency of droughts is also increasing as a result of climate change and may reduce the prevalence of malaria in certain regions. However, the broader effects of climate change on local livelihoods, food security, and migration may increase population vulnerability to the disease and undermine the effectiveness of control strategies, irrespective of the direct effects of climate change on transmission. Es

#### DENGUE

In recent decades, the geographic range of dengue, the most common mosquito-borne viral disease worldwide, has expanded substantially in response to declining vector-control programs and increasing global trade and travel.29 An estimated 390 million cases occur each year in more than 100 countries.30 The four serotypes of dengue virus are transmitted between humans the primary reservoir host — by infected female mosquitoes, most commonly Aedes aegypti and A. albopictus. Water-storage containers, which are commonly used in regions where a piped water supply is inadequate, or rainwater-filled containers (e.g., tires, pots, and tree holes) can become mosquito breeding sites and can thus drive epidemics.31 Transovarial transmission of dengue virus (from female mosquitoes to their offspring) and the long-distance dispersal of drought-resistant aedes eggs in suitable containers facilitate efficient expansion of the virus worldwide.32 The northward expansion of A. aegypti and A. albopictus thus far is best explained by human movement patterns within regions in which the climatic conditions are suitable for geographic expansion; however, by 2030, the dominant cause of expansion of these vectors is predicted to be climate change.33 The differential ability of A. aegypti and A. albopictus to survive normally lethal temperatures may influence their roles in future outbreaks.

## LYME DISEASE

Lyme disease (which is caused by the *Borrelia burgdorferi* sensu lato complex) is the most common tickborne illness worldwide, with an estimated seroprevalence of 14.5%; the reported

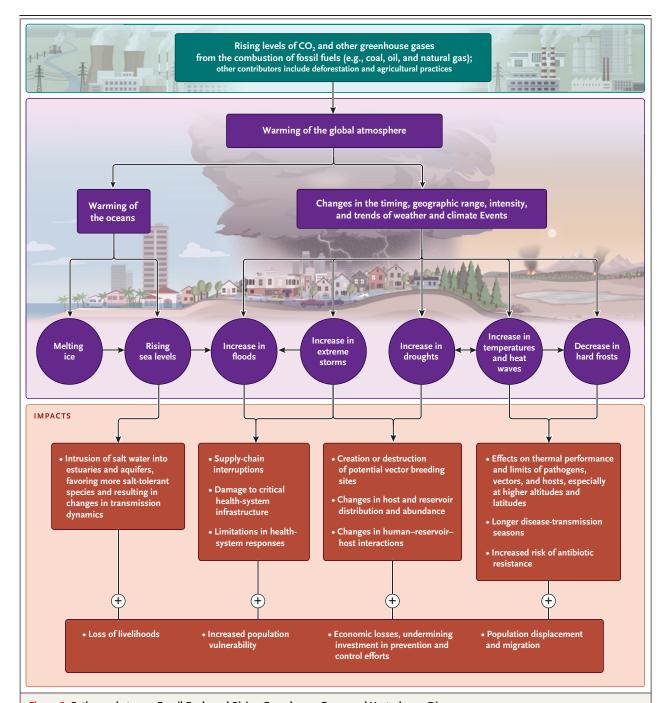


Figure 2. Pathways between Fossil Fuels and Rising Greenhouse Gases and Vectorborne Diseases.

The climate experienced at any location and time represents a combination of natural climate variability and, increasingly, climate change. As greenhouse gases accumulate and lead to increased global temperatures, extreme weather events are becoming more frequent, more severe, or both.

prevalence is highest in the temperate regions of Worldwide, Lyme disease involves four dominant central and western Europe and East Asia.34 tick species, although generally only one tick Without early treatment, infection can cause species is important in any given region.<sup>35</sup> Widedebilitating multisystemic chronic disease.34 ranging reservoir hosts — including mammals

	Examples of Observed or Predicted Effects	In the Ethiopian highlands, a temperature increase of 0.2°C per decade has exposed a growing population of nonimmune persons to the risk of malaria during the past three decades. <sup>9</sup> Malaria is reemerging in temperate regions in response to public health infrastructure decline, migration, and higher temperatures. <sup>10</sup>	Aedes vectors are increasingly emerging in temperate regions such as North America and Europe, <sup>12,13</sup> as evidenced by the occurrence of autochthonous dengue infections in Croatia and France in 2010. <sup>14</sup> The extreme flooding in Pakistan in 2022, which was attributed in part to climate change, <sup>15</sup> has resulted in a surge of dengue cases. <sup>16</sup> The emergence of Zika virus infection in Brazil in 2015 occurred during a period of severe drought and unusually high temperatures caused by El Niño, as well as short- and long-term warming trends. <sup>17</sup>	Lyme disease and other tickborne diseases have started to emerge in Canada <sup>19</sup> and more recently in the Arctic. <sup>20</sup> In 2015, sea birds in the arctic region of Norway were discovered to be carrying <i>Ixodes uriae</i> ticks infected with Lyme disease <i>Borrelia garinii</i> spirochetes.	Heat waves are associated with the emergence or reemergence of West Nile virus infection. The unprecedented upsurge in the number of human cases in Europe and Eurasia in 2010 was associated with an extreme summer heat wave. More recently, West Nile virus has been found to be overwintering in mosquitoes in Germany. <sup>22</sup> As a result of climate change, heat waves in Europe and Eurasia are expected to be more frequent and more intense, with milder winters in these regions.	Climate change is expected to weaken the powerful annual cycle of the intertropical convergence zone and move it southward. This may cause savannah black flies, which carry the most dangerous form of blinding onchocerciasis, to move from the savannah region to the forested areas of West Africa. <sup>22</sup>	Warming in the African highlands may reduce, rather than enhance, the transmission of bubonic plague because transmission is prevented when blood meals consumed by the flea coagulate and block pathogen transmission.	Climate change may lead to extinction of the vector (and therefore the human- and animal-associated diseases) in regions where temperatures are already close to the upper thermal limit of juvenile tsetse fly survival. <sup>25</sup>
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Change on Representative Arthropod-Borne Diseases.	Climate Drivers of Disease	In the absence of disease control and socioeconomic development, the spatial and temporal risk of malaria is largely governed by rainfall (temporary water bodies), temperature, and humidity. Although malaria is widely considered to be a tropical disease, it should be noted that it was previously common in temperate regions (e.g., Europe and North America). It has been eliminated in temperate regions mainly because of socioeconomic development and land-use changes.	Arboviral diseases are common in tropical and subtropical regions. <sup>11</sup> In drought conditions, households without access to secure piped water may store water in open containers in and around the home, which creates ideal domestic breeding sites for <i>A. aegypti</i> . After extreme rainfall events, outdoor natural and artificial containers provide ideal sites for <i>A. albopictus</i> mosquito egg and larvae development in urban and periurban areas.	The range expansion of Lyme disease is associated with a warming atmosphere and the effect of such warming on deer, mice, and tick populations. In the Sudano–Sahelian region of West Africa, tickborne disease may be associated with drought rather than with higher temperatures. <sup>18</sup>	The range expansion of West Nile virus infection is associated with a warming atmosphere, changing weather conditions, land-use changes (and their independent effect on bird migration), mosquito population dynamics, and flavivirus survival, replication, and virulence.	Simulium damnosum breeding sites are common in the whitewater river systems of West Africa, where the long-distance movement of adult black flies is governed by the intertropical convergence zone — a band of thunderstorms that moves across West Africa, bringing monsoon rains.	Climate-related factors influence localized outbreaks and worldwide pandemics. <sup>23</sup> Certain rainfall patterns favor large increases in rodent populations that support the flea population. Worldwide, 90% of cases of plague occur in Africa <sup>24</sup> in cooler highland environments, where temperatures below 27°C favor the transmission of Y. <i>pestis</i> from the most common vector, the <i>Xenopsylla cheopis</i> flea.	Tsetse flies are pervasive in the Zambezi valley of southern Africa. Populations of tsetse flies plummet during the hot, dry season; this decline has been exacerbated in recent decades by the observed warming in the Zambezi valley.
licted Effects of Climate Cha	Vectors and Transmission Pathway	Anopheles mosquito Direct transmission	Aedes aegypti and A. al- bopictus mosquitoes dominate Mostly direct transmission but zoonotic compo- nent in some contexts	Ixodes tick Zoonosis involving mice, small mammals, birds, and deer	Culex mosquito Zoonosis involving birds, horses, and other mammals	Simulium black fly Direct transmission	Flea Zoonosis involving small mammals (including rats)	Tsetse fly Direct transmission
Table 1. Observed and Predicted Effects of Climate	Disease and Pathogen	Malaria Plasmodium protozoan parasite	Dengue and Zika virus infection Flavivirus	Lyme disease Borrelia spirochete bac- terium	West Nile virus infection Flavivirus	River blindness Onchocerca volvulus nema- tode (helminth)	Plague Yersinia pestis bacterium	Human sleeping sickness Trypanosoma brucei proto- zoan parasite

(e.g., mice and squirrels), lizards, and birds—are part of the ecologic complexities of this disease; however, humans play no role in ongoing transmission.<sup>35</sup> The life cycle and prevalence of tick vectors, primarily *Ixodes scapularis* and *I. pacificus* ticks in North America and *I. ricinus* and *I. persulcatus* ticks in Europe, are strongly influenced by the abundance of reservoir hosts and by the ambient air temperature.<sup>36</sup>

Insurance records indicate that 470,000 cases of Lyme disease were diagnosed and treated in the United States during the period from 2010 to 2018, as compared with 329,000 cases during the period from 2005 to 2010.37. Lyme disease is most common in the Northeast and rare in the Southeast; although tick vectors are found in both regions, variations in the host preferences of the ticks (e.g., lizards or mice), in the hostseeking behavior of the ticks, and in the tick density help to explain this geographic pattern.<sup>36</sup> The increases in Lyme disease cases in the Northeast are largely attributed to the recovery of white-tailed deer populations,36 which are critical hosts for adult stages of the tick vector; however, increased human-tick interaction owing to the extended summer season resulting from climate change also contributes to the increases in cases. Warming temperatures have been associated with the expansion of ixodes ticks into Canada and Norway, with a corresponding increase in cases of Lyme disease. 19,38

### WEST NILE VIRUS INFECTION

West Nile virus causes potentially fatal neuroinvasive disease in humans and animals worldwide.<sup>39</sup> The virus is part of a complex ecosystem that is centered around a bird-mosquito transmission cycle involving more than 300 bird species and at least 65 mosquito vectors. Mammals, including humans and horses, can be incidentally infected. Human infections are mostly asymptomatic but can cause life-threatening illness in rare cases, predominantly in older adults and in immunocompromised persons.<sup>40</sup>

West Nile virus, which was first identified in the United States (in New York City) in 1999, is the leading cause of mosquito-borne disease in the continental United States. During the period from 1999 to 2016, nearly 7 million persons were infected.<sup>41</sup> The observed air temperature that results in a peak incidence of the virus among humans across the country was found to be

24°C, which closely matches the temperatures (which ranged from 24°C to 25°C) that were predicted by mechanistic models that were based on vector and pathogen thermal performance curves.<sup>42</sup> Warming temperatures are expected to shift transmission of this disease northward, as is already occurring in Europe; local transmission was recently discovered in Germany after unusually warm weather.<sup>40</sup>

## INEQUALITY AND VULNERABILITY

Climate change exacerbates inequalities, such as those driven by systemic economic injustice.<sup>43</sup> Persons living in less developed countries bear the greatest burden of most vectorborne diseases, a circumstance that reinforces health inequities and hinders socioeconomic development. Poverty, inadequate housing, poor environmental conditions, and limited access to quality health services exacerbate the effect. Children are particularly susceptible,44 owing in part to the effects of malnutrition<sup>45</sup>; women and older adults are also at increased risk. Vectorborne diseases during pregnancy are associated with particularly poor health outcomes among mothers and newborns from low-income or otherwise disadvantaged groups,46,47 as evidenced by the devastating effects of congenital infection with Zika virus during the explosive epidemic of Zika virus infection (which was spread by aedes mosquitoes) in Brazil in 2015.48

#### PUBLIC HEALTH INTERVENTIONS

Investments in surveillance and control have led to improvements in the public awareness, detection, prevention, and treatment of vectorborne diseases<sup>49</sup> and form the basis of adaptation strategies for a changing climate (Fig. 3).1 Specific measures to be taken vary according to disease, pathogen life cycle, and the level of risk and may include a combination of enhanced and new land-use management strategies, climate-informed early-warning systems, improved access to prevention measures (e.g., biologic mosquito control, personal protective measures, insecticides, and vaccines), and new and improved therapies<sup>50</sup> (Table 2). Figure S1 shows the projected benefits of adaptation strategies with respect to vectorborne disease rates. To be successful, interventions must include sustainable funding, as well

as community and household acceptance and uptake. A 2017 survey of 1083 U.S. vector-control programs showed that 84% of the programs were rated as "needs improvement" in one or more core competencies (e.g., insecticide-resistance testing).<sup>59</sup> The same year, the Centers for Disease Control and Prevention established five regional centers of excellence to help respond to emerging vectorborne diseases and to help create a new generation of vector experts.<sup>60</sup>

Malaria highlights several challenges that can occur in the implementation of adaptation strategies. After two decades of concerted international and national investment and consistent declines in malaria cases and malaria-related deaths, worldwide funding has stagnated; malaria is now resurgent in several countries, owing in part to increasing drug and insecticide resistance and, to a lesser extent, to service disruptions resulting from the coronavirus disease 2019 (Covid-19) pandemic.<sup>26</sup> Innovations are needed to keep up with biologic and socioeconomic challenges and to ensure equitable access to high-quality treatment in low- and middle-income countries.

The prevention of dengue and West Nile virus infection relies mainly on community-level mosquito-control programs; the implementation of such programs varies according to several factors, including funding and management. <sup>61</sup> Avoidance of the vector habitat during the transmission season as a result of public communication has long been an important prevention strategy for Lyme disease. <sup>62</sup> Various personal protective measures (e.g., insect repellent and protective clothing) and tick-control strategies (e.g., the culling of deer) have been proposed as approaches to reduce the risk of Lyme disease, but evidence of effectiveness is generally lacking. <sup>60</sup>

Vaccines have been successful in the prevention of three vectorborne diseases: yellow fever, Japanese encephalitis, and tickborne encephalitis. Despite the fact that vaccines approved for malaria and dengue in the past several years have had only limited success, efforts are under way to develop new and more effective vaccines that target vectorborne diseases. A recent phase 2 trial showed the effectiveness of a single infusion of a monoclonal antibody against *Plasmodium falciparum* infection over a 6-month follow-up period in Mali during malaria season. A for-

merly approved, effective vaccine for Lyme disease was withdrawn from the market, <sup>67</sup> but a new Lyme disease vaccine is currently being evaluated in a phase 3 trial (ClinicalTrials.gov number, NCT05477524). Similarly, a new dengue vaccine has shown promise in a phase 3 trial, and regulatory approval by European authorities is being sought (NCT02747927). According to the Intergovernmental Panel on Climate Change, successful vaccine development and uptake — although made more difficult by the growing worldwide challenge of vaccine hesitancy — have the potential to substantially offset the effect of climate change on vectorborne diseases.<sup>1</sup>

Better surveillance data and climate-informed early-warning systems are needed to enhance public awareness, facilitate the targeting of resources (human and financial) for improved responses,<sup>5</sup> and identify knowledge gaps and research needs. Adaptation plans must be time-sensitive and context-specific while also taking into account factors such as shifting disease patterns, extreme weather events, and current and future climate variations and trends.<sup>28</sup> This approach will require collaboration among various sectors, such as local health communities, urban planners, and climate experts.<sup>68</sup>

# IMPLICATIONS FOR CLINICAL PRACTICE

Education of health professionals is needed with respect to specific vectorborne diseases, particularly in regions in which diseases are newly emerging or anticipated to emerge. In many locations, clinicians are likely to see more cases of vectorborne diseases during longer transmission seasons, especially in regions with historically low levels of transmission. Awareness of local changes in disease rates and travel histories on the part of clinicians is important.<sup>69</sup> The nonspecific clinical manifestations of many vectorborne diseases often make diagnosis difficult.<sup>70</sup> Strategies for the prevention and treatment of vectorborne diseases are reviewed in Table 2. To help address the additional burden of health care delivery created by a changing climate, health professionals can advocate for more climate-resilient health systems<sup>71</sup> and for programs that focus on the current worldwide shortages of health professionals, including infectious-disease experts.

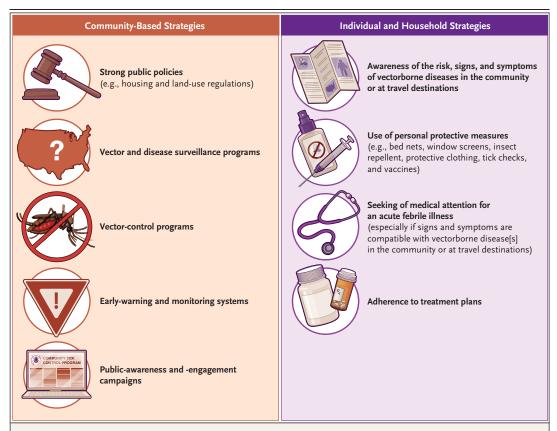


Figure 3. Key Adaptation Strategies for Responding to Vectorborne Diseases.

The panel on the left shows key community-based strategies for responding to the threat of vectorborne diseases. These include public policies that regulate land use and home construction, both of which can affect vector breeding sites and reduce the risk of indoor mosquito bites; surveillance for vector abundance and the incidence and prevalence of disease; and syndromic surveillance. Surveillance may be enhanced through multisectoral collaborations (e.g., meteorologic services that might predict conditions that are suitable for enhanced mosquito survival and parasite development). If early-warning systems can be developed, they could be used to enhance vector and disease surveillance in targeted areas. Then, if a surveillance threshold is exceeded, certain measures could be taken or reinforced (e.g., initiation of public-awareness campaigns that educate and provide guidance). The panel on the right shows the strategies that individual persons and households can use to prevent or respond to a vectorborne disease. For adaptation measures to be successful, community and household acceptance and uptake are needed.

#### MITIGATION OF CLIMATE CHANGE

Reducing the risks of vectorborne diseases and other health consequences of climate change requires not only adaptation but also a rapid and equitable transition away from fossil fuels. The signing of the Inflation Reduction Act of 2022 represents a necessary (although insufficient) move toward decarbonizing the U.S. economy in line with the goals set under the Paris Agreement in 2015. The health care sector, which contributes an estimated 4.9% of the total carbon footprint worldwide, must be part of the

process.<sup>72</sup> As trusted voices,<sup>73</sup> health professionals can weigh in regarding the importance and urgency of mitigation.<sup>74</sup>

## CONCLUSIONS

Climate change has substantial effects on pathogens, vectors, and reservoir hosts, with implications for the health sector worldwide. Many vectors are already expanding their latitude and altitude ranges, and the length of season during which they are active is increasing; these trends are expected to continue as the climate continues

Category	Adaptation Measures
Vector and disease surveillance programs <sup>51</sup>	Newer surveillance systems that incorporate satellite data, smartphone-based disease reporting, and integration of artificial intelligence algorithms
Early-warning systems <sup>52</sup>	Accurate and timely climate-informed early-warning systems that monitor climatic, environmental, and socioeconomic drivers of vectorborne diseases and thus ca be used to predict outbreaks
Vector-control programs <sup>53</sup>	Development of effective insecticides that are safe for humans and the environment†  Noninsecticidal strategies including microbial (e.g., the introduction of wolbachia-infected mosquitoes into field populations) and genetic (e.g., the introduction of genetically modified mosquitoes that transmit a deleterious mutation)
Public awareness and engage- ment <sup>54</sup>	Ongoing education of policymakers, public health officials, health professionals, and the public, particularly in regions where vectorborne diseases are emerging or anticipated to emerge
Health care systems <sup>1</sup>	Training of climate-educated health professionals to address the domestic and worldwide shortage of health professionals Climate-resilient health care systems Universal health care
Public policies <sup>55</sup>	Improved housing regulations (e.g., requiring household designs that restrict mosquito entry) Improved land-use management strategies (e.g., limiting deforestation in tropical areas to prevent increases in mosquito breeding sites)
Personal protective measures <sup>56</sup>	Use of safe and effective insect repellents Use of insecticide-treated bed nets Avoidance of tick habitats (e.g., moist, shady environments), especially in spring and summer
Household interventions <sup>55</sup>	Indoor application of safe insecticides in homes with cracks, crevices, or ill-fitting doors or windows and in homes without door or window screens Installation of window screens Improved housing design to ensure well-fitting doors, windows, and roofs
Vaccines <sup>1,57</sup>	Safe, effective, and affordable vaccines for vectorborne diseases that confer durable immunity  National programs to address vaccine hesitancy
Drugs <sup>57</sup>	Development and broad distribution of drugs for the treatment of vectorborne diseases, especially those predicted to have growing burdens of disease Surveillance for drug resistance and understanding of potential effects of climate change on antimicrobial resistance <sup>58</sup> Enhanced monitoring for counterfeit drugs
Diagnostics <sup>1</sup>	Inexpensive, accurate, and widely available diagnostics for climate-sensitive vector-borne diseases

<sup>\*</sup> The development and implementation of adaptation measures are associated with considerable challenges, including the need to ensure adequate funding for research and development and for the establishment and sustaining of programs and to ensure equitable access to adaptation measures.

to warm. Changes at the local level will be context- and disease-specific. Clinicians should be alert to changes in risk for the population they serve. To protect health and equity in a warmer world, investments are needed in vector control with respect to tailoring measures to rapidly emerging situations and in new forms of technology and approaches, including vaccines. Unfortunately, adaptation strategies will not be viable

as a long-term solution without the implementation of sufficient, urgent mitigation efforts to maintain global temperatures below critical thresholds.<sup>1</sup>

Disclosure forms provided by the authors are available with the full text of this article at NEJM.org.

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<sup>†</sup> Însecticides such as dichlorodiphenyltrichloroethane (DDT) have historically been the cornerstone of vector-control programs but have become less effective and have unacceptable environmental and toxicologic effects.

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