

Comprehensive Analysis of Malaria: Causes, Incubation Period, Transmission Methods, Prevention, Control, and Treatment

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ABSTRACT

Malaria is causing a significant burden of diseases and deaths worldwide. It has become a global issue, and its long history can be traced back to the most ancient writings. Malaria is historically known as an infectious disease. Malaria cases are classified by several causes, and the pathogen of malaria can be found in various species of the mosquito vector and four different types of the parasite in the blood. The onset of the disease occurs just 48 hours after the bite of the female Anopheles mosquito, which is very characteristic and called cold in agues. This is also a typical symptom of a fever and one of the earliest biomolecular disease philosophies based on the life cycle model. Moreover, it is also a recurrent infection that is more severe than the previous infection and is not transmitted directly between humans. It is important to detect the pathogen immediately for the pre- and post-infection processing of infection and onset, as there is only a short time to perform medical treatment. However, the microscopic identification of the pathogen in the blood is very difficult, especially when the concentration of the pathogen in the blood is low. Most studies have developed diagnostic devices to detect the concentration of the pathogen. Methods: The key electronic sources of information included databases and websites. Information from earlier reviews was used, along with relevant articles within the chosen time frame and languages (Spanish, English, Portuguese, and French). Key search terms included epidemiology, malaria, plasmodium, prophylaxis, antimalarial, parasites, insecticides, vectors, and vector control. After deduplication, 1,001 titles were scanned, resulting in 231 abstracts. After assessing the abstracts, 82 articles were obtained, with 1 article unable to be obtained. Nine articles were excluded, leaving 72 articles for this review. Two articles published in Spanish were found. Historical sources were used to provide context. Conclusion: In recent years, malaria cases have been reduced significantly with the use of insecticide-treated nets, indoor residual spraying, and anti-mosquito inoculation on a large scale in most

countries throughout the world. However, many individuals in all 56 endemic countries have not benefited from these effective methods yet, particularly those individuals in impoverished and marginalized populations. We focus on the causes of malaria, incubation period and symptoms, transmission methods, prevention (personal, community-based, and protection from mosquito bites), control approaches, and treatment options. The purpose is to raise awareness in both societies and governments in which malaria is still a severe public health problem, to promote the importance of disease prevention and control activities, and to discuss them in a single paper together, which has not been studied comprehensively yet. We expect that about 100,000 malaria cases in at least 44 endemic countries can be prevented due to increased fundraising and elimination plans accordingly. In conclusion, the real reasons for the high-risk people who cannot access malaria-reduction methods should be investigated thoroughly, and hidden subpopulations should be revealed using further studies. Useful advice and measures about protection and control must be given to all the people who visit malaria-endemic countries. Not just the individuals in malaria-endemic zones, there is a duty to everyone to prevent invitations to misery. We all must accept social responsibilities for avoiding the continuity of health disasters because the health of everyone affects the health of everyone else. Increased fundraising in health research and malaria programs conducted by global organizations, foundation support, and extra funding from philanthropists is expected, and government support for affected countries is needed.

KEYWORDS: malaria, transmission, incubation period.

1. Introduction

Malaria is a significant health problem in many parts of the world, transmitted by female *Anopheles* mosquitoes. Malarial infection happens through the bite of female *Anopheles* mosquitoes. The parasites form together within red blood cells and rely on the host for nutrients to produce energy. If a person is not able to receive prompt and appropriate treatment, there is a risk of cases where the disease can be severe, life-threatening, and relapsing. When malaria is present in the blood, the human host can serve as a source for the further spread of the disease. Hence, controlling the spread of malaria pathogens in human hosts is very essential. To reduce malaria infections and to support anemia control, antimalarials are frequently distributed to protect pregnant women and infants from severe illness. Additionally, antimalarials are given to groups at high risk, such as seasonal workers. Malaria infection cannot spread from one human to another through casual contact. The main prevention measures comprise personal protection using bed nets and repellents, stripping and drainage of mosquito breeding grounds, and spray programs. While malaria infections are frequently acquired by traveling to foreign countries, infections can still occur in the United States through blood transfusions, organ donations, and the sharing of needles among drug users. Malaria can be transmitted from a pregnant mother to her unborn baby. Settings for the transmission of malaria are extensive and continue to put significant pressures on global health systems. Development of malaria tightly depends on environmental conditions, and as a result of accelerated

urbanization, early detection of malaria is absolutely mandatory. Early detection of the disease could then lead to prevention and removal of the risk, and eradication measures can be taken to protect the people to some extent. Treatment for malaria has been available for a long time. To evoke early prevention measures, even though the technology to produce the vaccines is already there, this type of vaccine has not been developed.

1.1. Definition and Background

Malaria, a serious disease caused by a parasite, is transmitted by female *Anopheles* mosquitoes. Among all the diseases carried by arthropod vectors that cause human illness, death, and economic hardship, malaria is the world's most serious. Malaria is a life-threatening disease. It is typically spread by the bite of the female *Anopheles* mosquito. It can cause death, especially in children. Malaria contributes to poverty and can limit economic growth. Pregnant women are at high risk. The malaria infection is caused by a parasite and is transmitted through the bite of infected mosquitoes. The parasite can lead to the rupture of red blood cells within 48 to 72 hours—that's when people experience fever and other symptoms. After a period of initiation of the blood cells, the parasites pass into the blood circulation, eventually invading the erythrocyte. The sexual aspect of the life cycle of malaria, associated with the spread of parasites to other individuals living in an area, contributes to maternal anemia and results in significant adverse effects. Death and disease are represented by malaria, and this is one of the most serious health problems worldwide today. Malaria has been present since the Tertiary Era, at least. However, only recently, has the detailed system of the parasite become fully recognized, and physicians have been trying to reduce the occurrence rate of malaria using experimental methods.

1.2. Global Impact

Malaria is a disease of low socio-economic and public health significance at the global level. It has a negative impact on the gross domestic product of the affected countries. Poverty is the cause of malaria, and malaria is the cause of poverty in many countries with a high disease burden. Malaria causes a burden on the economy of a country and specifically on its poorest members in terms of the direct cost of medical care and the indirect costs of lost wages because of working days lost in seeking medical treatment at health facilities or homes, loss of productivity while sick at home, and the high price of labor for agricultural activities due to the shortage of labor. The most important consequence of malaria is that it increases poverty, particularly in the population of poor households because they have limited options for seeking optimal health care services. Malaria impedes the development of the affected countries due to the direct and indirect costs of illness and premature death. Malaria continues to be a leading cause of disease and death in developing countries, particularly Africa. The disease affects the entire population, but its impact falls most heavily on poor areas, poor households within those areas, and poor individuals. The economic impacts of malaria, often described as the consequences of a broad range of individual, household, community, and national levels, impose considerable burdens on households and limit the overall economic development of affected countries. Malaria constitutes an obstacle to improving the development index of the

affected countries. Malaria continues to have a major impact on the economy across Africa. It is one of the leading factors inhibiting economic growth and development. This underperforms the opportunities for economic development. Malaria is a serious national problem that substantially inhibits economic development, worldwide production, and social welfare in the most populous countries in the world. The net impact of these economic costs on society is astronomical, significantly dwarfing the direct benefits of treating the disease.

2. Causes of Malaria

Malaria is caused by Plasmodium. Plasmodium is a type of organism. The parasite is known as Plasmodium. Humans are the intermediate host of Plasmodium, and female Anopheles mosquitoes are the definitive hosts. Four species of Plasmodium cause malaria cases: *P. falciparum*, *P. vivax*, *P. ovale*, and *P. malariae*. Human infections are caused by *P. knowlesi*. The most dangerous form of Plasmodium is *P. falciparum*. This form can be diagnosed with a differential white cell count. The disease spreads rapidly and can be fatal as well. The incidence of this disease is malaria. The chronic, relapsing nature of *P. vivax* and *P. ovale* infections is caused by the presence of dormant parasites in the liver. This is known as hypnozoites. To cure these hypnozoite infections, *P. vivax* or *P. ovale* requires a short course of primaquine. Malaria is an infectious disease caused by the parasite Plasmodium. The disease affects humans, birds, other mammals, reptiles, amphibians, and arthropods, but it is generally caused in humans and other mammals by a few species. It is suspected that human malaria originated during early human history when it came into contact with the disease, more likely after the near end of the last ice age and the invention of agriculture.

2.1. Parasitic Agents

Malaria, an infectious disease caused by parasitic agents of the genus Plasmodium, is transmitted by particular species of Anopheles mosquito. It is the chief cause of mortality and morbidity in the world and is common in tropical areas. Every year, it causes the death of approximately 2–3 million people worldwide. It maims millions of other humans and presents a great threat not only to human health but also to the social economy in tropical areas. Although the process of malaria has been extensively studied, the credit for it being associated with malaria was given to a famous Roman physician. In 1717, he wrote that fetid air from marshes was toxic and caused a syndrome named malaria (mal-aria in Italian). The word itself means bad air. From the beginning, people recognized a link between the condition of malaria, its diagnosis, and certain geoclimatic and topographic conditions. At the end of the 19th century, an Italian doctor distinguished between a series of types of fever in the Roman Campagna, moving away from the miasmatic theory and associating malaria with a microscopic, parasitical being identified in bird blood and, failing that, in human circulation. (Adugna et al., 2021)

At the end of the century, a doctor, in the Roman Campagna, and another in India, demonstrated the complete life cycle: from the mosquito to man and vice versa. The recognition of the unique causal agent of malaria was given in 1880 when a

researcher was studying the blood of patients experiencing phenomena of malaria in military hospitals. The coining of the binomial for the specific causal agent of malaria was derived from a visionary interpretation of the enthusiasm shown by doctors. The genus name originated from the long-known cyclical nature of the disease and the Latin root *malariis* (malaria). At present, four species of the genus *Plasmodium* are credited with infecting humans. These are *P. vivax*, *P. falciparum*, *P. ovale*, and *P. malariae*. A plasmodium-like but non-zoonotic species, namely *P. knowlesi*, is also infectious. It is a macaque parasite transmittable to humans. The complete genome of the target, the potential seed, and the vector are available for all species. Another highly significant aspect of malaria is related to the easy detection of the infection. Virtually, all inhabitants of malaria-endemic areas already have an immune system; hence, asymptomatic malaria carriers can be discovered in the community by qualitative and quantitative examination. Quantitative examination indicates the density of the plasmodia, while qualitative examination determines the presence of mature and fertile gametocytes.

Consequently, the treatment inadequately lacks correlation with seasonality, the incubation time, and the seasonal effect of plasmodia on pregnant women or the parasite's resistance to drugs. Recent analysis should be integrated not only with actual climate and geographic data but also with information on demographics, behaviors, and migration or immunity of species, as well as economic, financial, and other studies to establish different routes of transmission and distribution in many regions of the world.

2.2. Vector Transmission

Malaria is transmitted solely by the infective bites of *Anopheles* mosquitoes. Female mosquitoes of the genus *Anopheles* are solely responsible for the transmission of malaria to humans. About 70 different *Anopheles* mosquitoes are known to carry malaria in nature. Since their bites predominantly occur in the evening, at night, and in the early morning, most transmission takes place from evening to early morning. *Anopheles* mosquitoes carrying female malaria parasites find their way into human blood to satisfy their nutritional needs. The presence of female parasites in the blood causes fever in malaria patients, which further enhances the transmission of malaria to other persons by biting the same mosquitoes. Malaria is not transmitted by person-to-person contact involving substrates, contaminated water, or air. Human feces and sewage do not play a role in malaria transmission. In the absence of mosquitoes, direct man-mosquito contact occurs at night. In countries where transmission is less common and people are not immune to malaria parasites, the illness may also be transmitted from one person to another through blood transfusions and organ transplants. Even though malaria is a disease of poverty, it is often found among travelers returning from vacations, businessmen, students, and members of the diaspora who go on vacation in their own country. (Boussougou-Sambe et al.2022)(Mponzi et al.2022)(Degefa et al., 2021)(Markwalter et al.2024)

3. Incubation Period of Malaria

The incubation period of malaria generally ranges from 7 to 30 days. This period is

affected by a variety of factors, such as the infective stage of *Plasmodium* and the kind. After an infective *Anopheles* insect bites a human, some non-infective strains of *Plasmodium* propagate and transform into an infective strain *in vivo*. When the infective metabolic products overflow *in vivo*, the *Plasmodium* is sucked into the abdomen of the *Anopheles* insect by the blood-sucking of *Anopheles*. In the female mosquito midgut, the *Plasmodium* further develops into oocysts with the aid of the chitinolytic enzyme.

The oocysts act as asexual sporogenesis and multiply the asexual sporulation stages, so a large number of sporozoites are bisected in multiples. After the oocyst matures, the sporulated sporozoites burst out, fall into the body cavity, and then run to the brain along with the nutrient granule of the mosquito anus of *Anopheles* which has bitten the body, thus causing disease occurrence. Upon the first infection of the mosquito by the sporozoites, it needs about 10 to 12 days for the *Plasmodium* to develop from sporozoites to the sporogenesis of the oocyst, and then 3 to 5 days to develop into mature oocysts to burst, *in vivo* a large number of infective sporozoites and to bite again.

3.1. Definition and Duration

Malaria is a life-threatening disease that is caused by parasites that are transmitted to people through the bites of infected female *Anopheles* mosquitos. Malaria is caused by *Plasmodium* parasites. In humans, malaria is caused by *P. falciparum*, *P. malariae*, *P. ovale*, *P. vivax*, and *P. knowlesi*. *P. falciparum* and *P. vivax* are the most dangerous. Approximately 40% of the world's population lives in malaria-endemic areas and is at risk of *Plasmodium* infection. After being bitten by an anopheline mosquito infected with parasitic protozoans, the cycle of infection begins. The incubation period is about 7-30 days and depends on the *Plasmodium* species found in the bite.

There is yet no commercial vaccine against the disease; however, drugs can treat the disease and prevent the symptomatic progression of infection. Once people get ill, effective support treatment can help prevent death. In order to prevent the spread of malaria from one person to another and to protect against the disease, public health officials urge people to avoid mosquito bites and travel infested areas. There is a lot of drug resistance to medicines used to protect and cure malaria. Rapid diagnostic tools for identifying *Plasmodium* species that cause disease and their sensitivity to medicines will be tested. The number of sick hospitalized is high despite all these advances, 229 million people are at risk on the continent. Measures must be taken in Africa to bring this number to zero. Individual and community actions must be taken to improve health together.

4. Methods of Transmission

Malaria parasites can be transmitted from one person to another by a mosquito. The process of malaria being transmitted from one person to another involves the following: Female mosquitoes that bite a person release parasites from their salivary glands as they take a blood meal. These parasites travel to the liver, where they grow

and multiply into unknown numbers. When mature, they leave the liver and reenter the bloodstream, where they infect red blood cells and form the characteristic stages that are known to cause all the symptoms and common laboratory findings of malaria. When the infected mosquito bites another person, blood containing these stages is taken in during feeding, and the parasites begin the next cycle in the mosquito. Mosquitoes become so filled with parasites during the course of some infections that they can transmit malaria after biting only once. In fact, the insect needs to feed on three or more people to have enough parasites in the salivary glands to transmit malaria.

It is impossible for malaria to be transmitted directly from one person to another. Thus, no longer is it necessary to fear that casual contact with a person who has or is recovering from malaria will lead to malaria. For example, having sexual contact, cursory conversation, or sitting casually close to a person who has or is recovering from malaria does not lead to the transmission of malaria. Only if the indications listed earlier exist should the possibility of malaria from a blood transfusion or organ transplant be considered; a doctor can help decide whether malaria testing is necessary based on how likely it is that these circumstances exist.

4.1. Mosquito Vectors

A detailed understanding of disease transmission may help improve human awareness, leading to the application of methods for the prevention and control of malaria efficiently. One of the main responsibilities when studying potential vector groups is the examination of sylvatic reservoirs to find out if there are silent foci. The mosquito fauna of the sylvatic environment is rich and serves as an important link in the spread of the disease, as wild mosquitoes can enter homes in endemic forest areas. Mosquitoes from areas near homes in rural overlapping regions are of great importance due to their significant epidemiological relevance. Their presence may help assess the potential risk of new outbreaks occurring due to changes in open or marginal environments, which always result in more or less intense contact between wild vector populations in the disease transmission cycle in areas populated by humans. (Rakotonirina et al.2023)(Guégan et al.2020)(Loiseau and Sehgal2022)

Currently, seven species of the subgenus *Kerteszia* are recognized in Brazil. But what does this taxonomy and phytogeography have to do with the epidemiology of malaria in Brazil? Much is already known, but many unanswered questions encourage research on the subject. There is no doubt about the high epidemiological importance from the point of view of possible actions to prevent or control the disease. Despite their low density or abundance inside or near homes, knowledge of the ecological and biological aspects of *Kerteszia* species in the wild is important for planning, implementing, and showing the results of constant prevention and control of malaria in these transmission areas. Confirmed for social primates or humans, sylvatic transmission with involvement and development in the vectors of hepatic phases of *Plasmodium* in the subgenus *Kerteszia* anomalously extends the criteria usually adopted for understanding the transmission dynamics of the disease and even for post-discharge control of transmission.

5. Prevention Strategies

Malaria's risk can be reduced by limiting exposure to mosquitoes and using anti-mosquito procedures and medications. The latitude-adjusted breakdown of activities suggested for this purpose is straightforward. Among other measures, the broad use of swift testing should be mentioned. It makes it easier to follow up with confirmed cases and helps to avoid unnecessary treatments and negative tests while avoiding delayed onset of the disease caused by neglected patients. In the absence of rapid testing, medicines should be administered within 24 hours of the onset of symptoms. It is vital to educate patients about the symptoms of malaria and the possible risks associated with infection so that those living in endemic areas can recognize the early signs and act quickly. In areas of malaria resistance, physicians and microbiological laboratories should know that they could be invaded. In general, the attention paid to any feverish state of individuals should be increased to prevent emergence in many places. Resorting to homemade or professional mosquito safety devices can help in ensuring the boundary of the use of restricted prescription drugs to ensure practice, particularly in endemic areas.

The most effective tools for controlling malaria are vector control. The use of personal protection measures protects the vector from both day- and night-biting. For example, at gymnasiums, long-sleeved shirts and slacks should be worn, repellents should be used, and windows and doors should be secured. In indoor settings, pyrethroid neurotoxin-treated bed nets offer practically full protection. Towards evening, most of the female *Anopheles* species are blood-loving, and to mitigate the incidence of nocturnal bites and therefore to avoid becoming larvae, humans should prepare their bed nets every evening. House spraying should be supported consistently by indoor residual spraying. More durable techniques include eliminating mosquitoes through ecological or chemical means, such as stacking insect traps and rearing mosquitoes along with genetically modified mosquitoes. Scientists are looking for more contemporary methods to manage malaria cases that could operate against all types of mosquitoes and at various stages. Judicial governance also plays a vital part in the fight against insect attacks. Each nation is vaccinating against treatments, and national and international networks are being depleted by the use of human biology to monitor malaria, best recognize countries, and understand their epidemiological and malaria control requirements to provide today's different malaria risk measures.

5.1. Vector Control

Malaria, a vector-borne disease, is caused by parasites called *Plasmodium* spp. It is transmitted to humans by mosquitoes. The disease results in high fevers and chills, headaches, severe flu-like symptoms, and occasionally causes death. Control of the vector, such as through indoor residual spraying or the use of mosquito bed nets, is the most effective prevention and control tool. No vaccines have yet been approved for human use, but studies show promise for developing an effective human vaccine. Vector control is considered the component of the public health malaria program that works properly and is used extensively. It is widely used because it successfully reduces the densities of malaria vectors and minimizes malaria transmission and associated diseases and deaths. The fundamental basis of vector control depends on

mosquito behavior. It is also based on the varying feeding and resting behaviors of different species of malaria vectors. Data on human-biting trends, as well as zones where mosquitoes enter and leave, also play major roles in providing useful information about the behavior of malaria vectors. Proper knowledge of mosquito vectors and their behaviors should be known to any control tool that one chooses to execute. Personal protection and community protection are the two tools that keep malaria from receiving a sufficient number of blood meals. Vector control is used to kill a large number of mosquitoes, and this reduces their population. The availability of multiple tools to control mosquitoes has many advantages, besides serving as insurance when the tools are applied to resistant mosquitoes, diseases, and separate biological or spatial settings.

5.2. Chemoprophylaxis

Chemoprophylaxis - preventive drug treatment - can provide either full or partial protection against *P. falciparum* malaria in those at highest risk for severe outcomes (including death), such as young children, pregnant women, or those with no previous exposure to malaria: 1) Full protection. Sensitive malaria strains respond quickly and efficiently to treatment with chloroquine or another drug. With no remaining parasites in the body, the person is protected from illness for the life of the current malaria season. Malaria may recur during following malaria seasons if reinfection occurs. 2) Partial protection. Primaquine or another drug has been unable to kill all sensitive malaria strains within the body. The person may still experience all the ill effects of blood-stage malaria (fever, etc.) for an initial infection. However, they should be protected for the remaining duration of the malaria season and should not have any of the severe effects possible with their first malaria attack. Full and partial protection from malaria are temporary conditions, existing only for the length of the malaria season. Relapses may occur with people infected with any malaria type, but relapses should have no severe effects. No lifelong protection is provided by chemoprophylaxis.

6. Control Measures

Malaria is a life-threatening disease caused by *Plasmodium* parasites and is transmitted through *Anopheles* mosquitoes. Humans with *P. falciparum* internally produce the human infectious type. Also, in the human form of *P. vivax* and *P. ovale*, along with the development of the liver schizonts, some form a hypnozoite which can infect mosquitoes for a long time during periodic relapses. Severe or moderate malaria occurs when an individual does not use or build resistance against the different stages, such as the sporozoite attacking liver hepatocytes, the merozoite breaking the RBCs, and the gametocyte that lives in the RBCs. Ever since medieval times, in addition to traditional treatments such as quinine and artemisinin, many drug treatments have also been used. In some areas, the prevalence of malaria has decreased drastically by manual killing of mosquitoes, while in others, mortality has reached 90% due to uncontrolled margins. There are other control measures such as infecting mosquitoes with *Wolbachia* that hinder *Plasmodium*, injecting mosquitoes with a weakened *Plasmodium* strain, and eliminating only males. However, the different modalities of *Plasmodium* interaction with the mammalian and vector hosts

should be considered to inform vaccine development. *Anopheles* mosquitoes are responsible for the transmission of malaria, and this depends on their ability to feed on human or animal hosts. However, the preference of some species for different types of blood can make the vectors less competent. The implementation of agro-ecological techniques such as the use of mosquito-repellent herbal plant extracts at relatively low prices could also control the vectors. Preventive interventions to reduce mosquito population growth also include the application of combined nets and insecticides inside and outside houses, the use of traps and screens in windows, the careful selection of the breeding sites of mosquitoes and the placement of salients, and finally, the elimination of the aquatic stage mosquitoes from bromeliads. Accordingly, the introduction of strategies involving residents of endemic areas and schools is more frequently traded for less expensive integrated interventions. Finally, mosquitoes can also be genetically manipulated, as in replacing dengue vectors. However, the engagement of governments and other stakeholders is crucial to ensure the sustainability of these interventions.

6.1. Vector Control Programs

Malaria control programs in many countries rely less on any one method to be successful than on a carefully selected mix of methods under adequate management that are well coordinated to fit the integrated disease control concept. Nonetheless, vector control offers some attractive features for both short- and long-term gains. Vector eradication is not feasible anywhere but in a locally restricted and bug-infested area. Malaria eradication, although at the time mainly an ideal, cannot be achieved without attacking the personal or household environment of all potential vectors in every major endemic area at the same time. In the urban tropical environment, where malaria now causes some major health problems, uncontrolled aggressive vectors have been developing a resistance to certain insecticides. It becomes extremely difficult to mount a successful chemical vector eradication program. Much uncertainty still clouds both the causes and the cures of the urban 'explosion,' and future activities will have to be preceded by epidemiological investigations. Such well-executed localized investigations will also provide the basis for a selective choice of the acceptable vector control tactics. Even when vector control lodges in the background only, in view of future needs, urgent research is required for developing new and safe chemicals and increasing the success of applications.

6.2. Environmental Management

Mosquitoes breed in water. This knowledge has led various public health interventions to focus on environmental management. Land is reclaimed to reduce the area of water and vegetation, and other materials are placed in water to remove the oxygen by flooding. In contrast, larvae eat organic materials and are removed from the areas that people go to wetter places. This creates the cycle. Fish are also placed in water so that they can eat mosquito larvae. In particular, the introduction of certain fish, which has higher stability than other fish and of which offspring belong, is often used.

The flooding method is an effective way to create flight density equal to a lowland heavy place during the transmission period in an upland area, which has various non-

native trees and air with little water vapor pressure. Moreover, because mosquitoes fly around water at the time of preparation for childbirth in an upland area, flooding can be used to create conditions in other harmful environments. Unlike the larvicide method, the flooding method has the possibility of contamination of the beneficial mite habitat for the biological control of citrus aphids.

7. Treatment Approaches

Since the evolution of malaria, the disease has been proved to be a challenge to curtailment. Many factors are involved in this challenging circumstance, which range from insufficient infrastructure to manage its prevention to unfit medication to manage the disease. This difficulty has influenced the sustainable control and national task planning of the governments in malaria-ridden zones. It is worth mentioning that the refusal to take the medications as correctly directed usually leads to the build-up of malaria parasites' immunity in the body, and, in turn, the patients become resistant to any antimalarial drug.

There are numerous antimalarial drugs in the market today, but regrettably, there are only a few types because the malaria parasite has built up endurance to the medications. Despite the use of homemade or fake antimalarial drugs in treating the disease, it has been attested that to eliminate the malaria pathogen from the body and its symptoms, drugs known as Artemisinin Combination Therapy are the most effective drugs recommended and used by most authentic health providers recently. Artemether/lumefantrine has been attested to be successful in demolishing the parasite and has a speedy action against *Plasmodium falciparum*. It is taken orally after meals and should be kept out of sunlight and moisture. Amalar treatment is used for dense, mild to severe malaria attacks with the only stubbornness of *falciparum*. This drug is only taken orally and should never be used for self-medication. Another highly acceptable drug is Atovaquone/Proguanil. It is good for treatment. It should not be used during pregnancy. Atovaquone inhibits mitochondrial form 2 type of cytochrome B, which is used as an electron transfer to produce pyrimidine triphosphate in the case of *P. falciparum*. If used with other drugs, the antimalarial effect can be boosted, i.e., when it is used together with the Malarone packet for kids. The medications have side effects like a sore throat, kidney or liver problems, issues with vision, or hallucinations. In case of all these conditions, stop taking the medication forthwith and seek immediate medical attention. Pyrimethamine, Loperamide, Clarithromycin, Chloroquine, Cladarine, and Artecs are some good examples of suitable antimalarial drugs. The amendments proposed in the treatment of malaria would include discontinuance of the prevalence to make the parasite in humans and mosquitoes go extinct, but this would be economically challenging unless an effective vaccine or medication can be discovered. Antirelization - reversal of a country from malaria and stable disease progress.

7.1. Antimalarial Drugs

Antimalarial drugs are designed to manage the disease and its symptoms by interfering with the life cycle of malaria parasites in the blood and other tissues of

the human body. To be effective, these drugs must adequately raise the concentration of their active ingredient in the blood to levels that can eliminate the malaria parasites. The serum concentration of an antimalarial drug is achieved by either single-dose therapy or by ensuring that the patient takes the complete doses of the drug within the duration of the treatment regimen. The complete doses of the antimalarial drug must be maintained for as many days as are thought to be required to completely eliminate all the parasites present in the treated individual's body. Unfortunately, malaria parasites possess the ability to fend off the effects of the body's immune system as well as the effects of antimalarial drugs. This ability to resist or to become immune to the effects of antimalarial medicines is called antimalarial drug resistance.

There are three main categories of antimalarial medicine: - Drugs used to treat people who have malaria. - Medicines to prevent malaria. - Household and community-based products not registered as medicines but helpful for malaria protection, treatment, and prevention.

Most, if not all, antimalarial drugs are either plant or synthetic in origin, and the malaria parasites can become less sensitive or resistant to the drugs' effects. The four main mechanisms of resistance are: - Where the drug is pumped out of the parasite cell or its serum concentration is reduced. This type of resistance mechanism is characterized by reduced erythrocyte/plasmodium levels of the drug. - Where the drug breaks up before it can kill the parasite. The malaria parasites metabolize and deactivate some of these drugs in the gastrointestinal tract or in tissues in the human body. - Changes in the target proteins that the drug is designed to interfere with, so that the signal molecule which the drug normally binds to can no longer do so. Once the signal molecule is unable to bind with the target protein, the drug's inhibitory action is eliminated, so it is of no effect. This form of resistance is mainly due to protein mutations or structural changes in the parasites' target proteins. - Enhanced repair of damage caused by the drug.

7.2. Drug Resistance

The emergence and spread of resistance to chloroquine, and to a lesser extent to sulphadoxine-pyrimethamine, by *P. falciparum* has led to increased use of antimalarial drugs under the umbrella term combination therapy. Initially, drugs of different classes were used, but fear of interactions between the two drugs led to the development of fixed dose combinations. The principal disadvantages for the pharmaceutical companies developing these combinations have led to increased use of sulphadoxine-pyrimethamine, in which resistance was reported relatively recently, and to artesunate/mefloquine, despite reports of resistance to mefloquine. Many compounds undergo development including combinations delivered in a single pill. Some are in the developmental pipeline, but are so far only available as separate tablets.

Problems with the deployment of artemisinin and ACT are partly a result of the time available for resistance to develop, suggesting that this class may also fail. Consequently, it has been proposed that ACT should be combined with a long-acting drug to provide post-antimicrobial host-protective immunity. Trials with passive and active immunization might also be worthwhile, as might a greater understanding of

immune memory and its elicitation by defined subunit antigens.

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