



[Editing file](#)

# Malaria & travel medicine



In this Lecture the doctor only focused on Malaria disease and skipped the rest of the Lecture

## Objectives :

- ★ Learn the epidemiology and etiology of malaria
- ★ Know the clinical presentation
- ★ Know Risks to travelers
- ★ Know Malaria and pregnancy
- ★ Know Diagnostic work up
- ★ Learn the treatment and prophylaxis

### Color index

Original text

Females slides

Males slides

Doctor's notes <sup>438</sup>

Doctor's notes <sup>439</sup>

Text book

Important

Golden notes

Extra

# Malaria

## Definition

Malaria is a protozoal infection caused by a few plasmodium species that spread by mosquito, once it enters the bloodstream it infects and destroys mainly liver cells and RBCs causes a variety of symptoms.

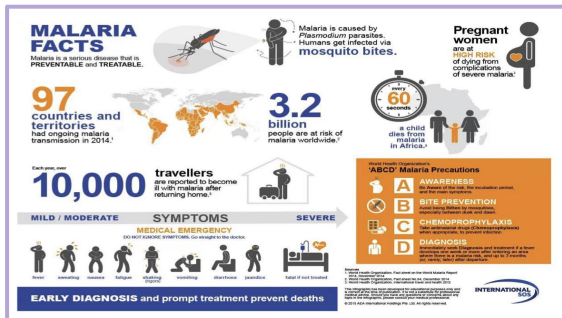
## World facts

97 countries have malaria.

216 million malaria cases worldwide in 2018.

445,000 malaria deaths worldwide in 2018.

10,000 world travelers become infected every year.



Goals, milestones and targets for the *Global technical strategy for malaria 2016–2030*

GOALS	MILESTONES			TARGETS
	2020	2025	2030	2030
1. Reduce malaria mortality rates globally compared with 2015	At least 40% 18% reduction achieved 22% off track	At least 75%	At least 90%	At least 90%
2. Reduce malaria case incidence globally compared with 2015	At least 40% 3% reduction achieved 37% off track	At least 75%	At least 90%	At least 90%
3. Eliminate malaria from countries in which malaria was transmitted in 2015	At least 10 countries On track	At least 20 countries	At least 35 countries	At least 35 countries
4. Prevent re-establishment of malaria in all countries that are malaria-free	Re-establishment prevented On track	Re-establishment prevented	Re-establishment prevented	Re-establishment prevented

Global technical strategy for malaria 2016-2030.

## Malaria in Travelers

Non-immune travelers exposed to mosquito bites between dusk and dawn.

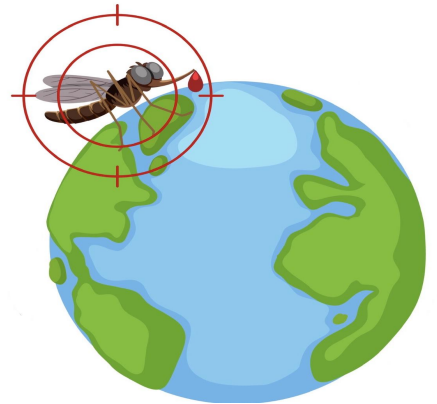
This includes previously **semi-immune** travelers who have lost or partially lost their immunity during **stays of 6 months** or more in countries or **areas of no risk**.

Travelers who have migrated to countries and areas of no risk are at particular risk when they travel to malarious areas to **visit friends and relatives**. Those travelers used to live in malaria endemic areas and got malaria during their childhood and think they have immunity against malaria and don't take proper precautions but there immunity is gone and they often get severe malaria when they get infected again.

Most cases of falciparum malaria in travelers occur because of poor adherence or inappropriate prophylactic malaria drug regimens and failure to take adequate precautions against mosquito bites.

Adherence to chemoprophylaxis can be improved if travelers are informed of the risk of infection and believe in the benefit of prevention strategies.

**Late-onset vivax and ovale** malaria may occur despite effective prophylaxis.



## Epidemiology

### Stable transmission:

including much of sub- **Saharan Africa**, transmission occurs consistently year round. The bulk of the mortality is seen in **children**, while those who survive to adulthood acquire significant immunity; **low- grade parasitaemia** is still present but causes few symptoms.

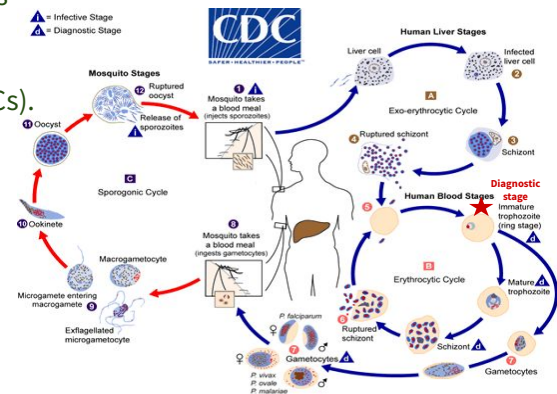
### Unstable transmission:

occurs when there is erratic, **seasonal or low- level transmission** (e.g. in the **Sahel belt**, where mosquitoes feed only in the rainy season). Little protective immunity develops and symptomatic malaria occurs at **all ages**. Changes in environmental or social conditions in such areas can lead to epidemics with substantial mortality in all age groups.

## Incubation period & Life cycle

- Sporozoites reach the liver within 1-2 hours following Female Anopheles mosquito bite.
- Patient is asymptomatic for 12-35 days until RBCs stage of parasite life cycle.

- 1) Malaria is mainly carried by **female anopheles mosquito**. the infected mosquito will bite to **take** a blood meal and **injects** the “Sporozoites” of the malaria parasite from its **salivary gland** into the human.
- 2) the sporozoites travel immediately to the liver where it starts the Human-Liver stage (exo-erythrocytic cycle as it happens outside RBCs).
- 3) the hepatocytes get infected, the parasite matures into “Schizont” ( multiply asexually to form **merozoites inside the schizont** (Exoerythrocytic schizont)
- 4) which grows up then ruptures released into blood, and infects the RBCs.
- 5) Human-Blood stages (erythrocytic cycle) begins, as the parasite is ingested by the RBCs it forms a ring (Ring stage) which is called “Trophozoite” and this is the **diagnostic stage**. then there is two pathways:



- 6) *First pathway:* the trophozoites mature and multiply to become “Schizonts” which grow and rupture the RBCs (**hemolysis**) releasing more parasite to the bloodstream, which infect more and more RBCs and cycle repeats. **Clinical attack** of malaria is due to this stage.

during the rupture (**hemolysis**) and release of the parasite, is when the patient presents with fever (intermittent fever) for example, this happens every 24h with *P. falciparum* so they have fever on a daily basis, while *P. ovale/vivax* it happens every 48h so they get fever every other day. while in *P. malariae* it happens every 72h, hence they get a fever every third day.

- 7) *Second pathway:* to complete the cycle some of the trophozoites, don't mature as a schizont, but into “Gametocytes” the sexual form of the parasite (male and female) that get ingested by the female mosquito. then the cycle of the gametocytes starts within the Mosquito to complete the “Sporogonic cycle” **Dr: which you don't need to know**

## Pathogenesis

- The pathology of malaria is related to **anaemia, cytokine release** and, in the case of *P. falciparum*, widespread organ damage due to **impaired microcirculation**. The anaemia seen in malaria is multifactorial . In *P. falciparum* malaria, red cells containing schizonts **adhere to the lining of capillaries in the brain, kidneys, gut, liver** and other organs. As well as causing **mechanical obstruction**, these schizonts rupture, releasing toxins and stimulating further cytokine release.
- **Causes of anemia** : Haemolysis of infected red cells ,Haemolysis of non-infected red cells (blackwater fever) , Dyserythropoiesis , Splenomegaly and sequestration ,Folate depletion.

## ◀ Modes of Transmission

- 1 Human to Human by **anopheles mosquito** (between dusk and dawn)
- 2 Blood transfusion
- 3 Contaminated needles
- 4 Congenital (vertical/mother to fetus)



## ◀ Predisposing factors:

- **non-immune individuals:**
  - children in any area
  - adults in areas of unstable transmission
  - visitors from a non-malarious region.
- People living **near airports** in Europe have acquired malaria from accidentally imported mosquitoes.

## ◀ Protective factors:

- **Certain genetic traits:**
  - P. falciparum : **sickle cell, thalassaemia, G6PD deficiency** and HLA-B53, does not grow well in red cells that contain haemoglobin F, C or especially S. Haemoglobin S heterozygotes (AS) are protected against the lethal complications of malaria.
  - P. vivax: People who **lack the Duffy antigen<sup>1</sup>** (a common finding in West Africa).

## ◀ Plasmodium species:

A protozoal infection is caused by one of the following organisms:

- **Plasmodium Falciparum** (most common and serious)
- **Plasmodium Ovale**
- **Plasmodium Vivax**
- **Plasmodium Malariae**
- **Plasmodium Knowlesi & Simium**

• P. vivax and P. ovale may persist in liver cells as dormant forms, hypnozoites, capable of developing into merozoites months or years later. Thus the first attack of clinical malaria may occur long after the patient has left the endemic area, and the disease may relapse after treatment if drugs that kill only the erythrocytic stage of the parasite are given.

• P. falciparum and P. malariae have no persistent exo-erythrocytic phase but recrudescence

1. The Duffy blood group antigen serves not only as blood group antigen, but also as a receptor for a family of proinflammatory cytokines termed chemokines, and as a **receptor for Plasmodium vivax malaria parasites.**

# Malaria

## Plasmodium Falciparum<sup>1</sup> Important

### General Information:

- Most **highly pathogenic** species, most serious and common
- **Resistant** to many antimalarial drugs.
- The species that causes the **most morbidity and mortality** worldwide.
- ★ Represents the major cause of malaria in **tropical** countries, Areas (Jazan)
- Responsible for the sporadic great regional **pandemics** that sometimes occur in **subtropics**.

### Pathogenesis:

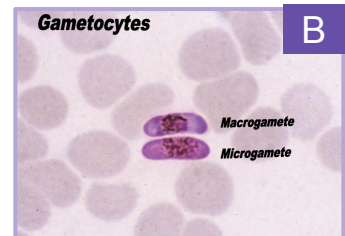
- 1) The pre-erythrocytic cycle starts immediately after injection of the **sporozoites** by the mosquito.
- 2) Infects **mature and young erythrocytes**. The surface of erythrocytes infected with late stage **trophozoites** or **schizonts** is altered so they **stick to endothelial cells in various tissues** (cytoadherence) causing **multi-organ failure**.
- 3) **Schizogony** (asexual reproduction) is particularly prolific in all stages (pre-erythrocytic, erythrocytic and sporogony) that may be the cause of its success as a species and its virulence.

### Characteristics:

- Infection in the peripheral blood is characterized by the presence of **ring forms and gametocytes**, whereas late trophozoites and schizonts are only seen exceptionally.
- The level of **parasitaemia may be high** and **multiple infection** in a single erythrocyte is common.
- The **gametocytes are characteristically crescent-shaped** (Pic B) and unlike the **gametocytes** of other species, are very slow to reach maturity (up to 10 days) and early forms of gametocytes are sequestered.

**Pic A: Trophozoite** (rings with chromatin dot) in thin blood film showing two **characteristic features of P. Falciparum:**

- 1) **Ring with double chromatin dots**
- 2) **Multiple parasites in one cell (RBC)**



## Plasmodium Vivax

### General Information:

- Occurs throughout most of the temperate zones as well as large areas of the tropics (but is mainly absent from tropical West Africa).
- It causes 'benign' **tertian malaria**.
- Polymorphic and the subspecies status proposed for some strains may be justified.

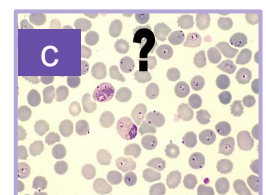
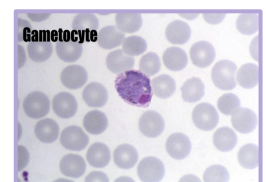
### Characteristics :

- Restriction of erythrocyte invasion to reticulocytes bearing Duffy blood group determinants (explains why **RBCs infected with trophozoites of P. vivax are sometimes described as larger than normal**).
- The presence of caveolar structures/ **vacuoles** on the surface of the infected erythrocyte membrane take up stain, and are described as **Schüffner's dots**.
- After invading the hepatocyte some of the sporozoites may transform into **hypnozoites**, then remain latent for months or years and be responsible for **subsequent relapses**.

**Pic C:** Co-infection with p.falciparum and p.vivax. ( there are features of both species: double chromatin dots and multiple parasite + large dysmorphic RBCs with vacuoles.



Dysmorphic-shaped large RBC, with **vacuoles**



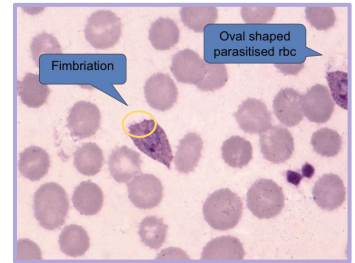
1. The fever has no particular pattern. **Jaundice is common** due to haemolysis and hepatic dysfunction. The liver and spleen enlarge and may become tender. **Anaemia develops rapidly, as does thrombocytopenia.**



## Plasmodium Ovale

### General Information:

- Distinct from *P. vivax* in minor morphological differences, antigenic and molecular differences. Most of the biological and **clinical features are identical**.
- The major biological difference is that *P. Ovale* **can infect Duffy negative reticulocytes**, whereas *P. Vivax* cannot.
- characteristic features: Oval shaped, **Fimbriation**



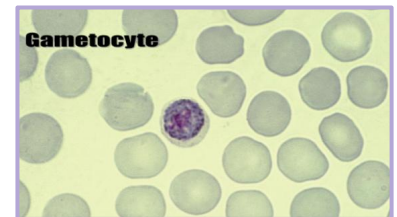
## Plasmodium Malariae<sup>1</sup>

### General Information:

- Differs from the other three human malaria parasites in its **slow development** and its **longer asexual cycle**.
- Development is slow in both the vector and the human host because of less efficient schizogony.
- The asexual cycle is **72 hours** instead of 48 hours – hence '**quartan malaria**' – because fever paroxysms occur every 4th day (according to the Roman custom of regarding day 0 as day 1).

### Characteristics:

- Infects **old erythrocytes**, explaining why infected cells are often described as '**smaller**' by microscopic.
- The presence of 'knobs' at the surface of infected erythrocytes, which are similar to *P. Falciparum*, but the cells **do not** exhibit any cytoadherence (and so no sequestration).
- The surface of infected erythrocytes **does not** exhibit any caveolar/vesicle complexes and **Schüffner's dots are absent**.
- **Sporozoites** of *P. Malariae* **do not transform into hypnozoites**, and so there are **no relapses**.
- *P. malariae* can survive for a very long time in the peripheral blood (**10 years or more**) at a very low level of parasitaemia occasionally producing detectable **peaks** with a recrudescence of clinical symptoms.



## Plasmodium Knowlesi & Simium

### General Information: *not common*

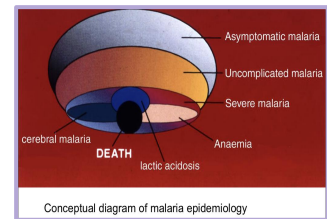
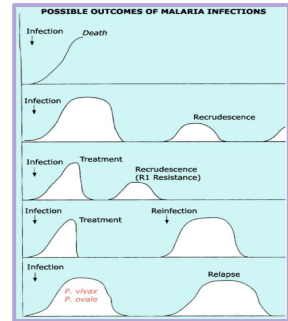
- Simian **Monkey to Human** transmission (zoonotic malaria)
- **P. Knowlesi**: in South East Asia, it looks like *P. Malariae*.
- **P. Simium**: in South America, it looks like *P. Vivax*.
- **Need PCR** for diagnosis.
- Can cause severe disease



1. In children, *P. malariae* infection is associated with glomerulonephritis and nephrotic syndrome.

## History

- **Travel to malaria endemic area**
  - **P. Falciparum**: up to 1 year from travel (usually first 3 months).
  - **P. Malariae**: up to 10 years.
  - The main symptom that is usually present is fever which is non specific.
- **Non specific: Fever**
- **Sweats, chills**
- **Myalgia**
- **Headache**
- **Diarrhea**
- **Cough**
- **Jaundice, dark urine**
- **Confusion, Seizures**



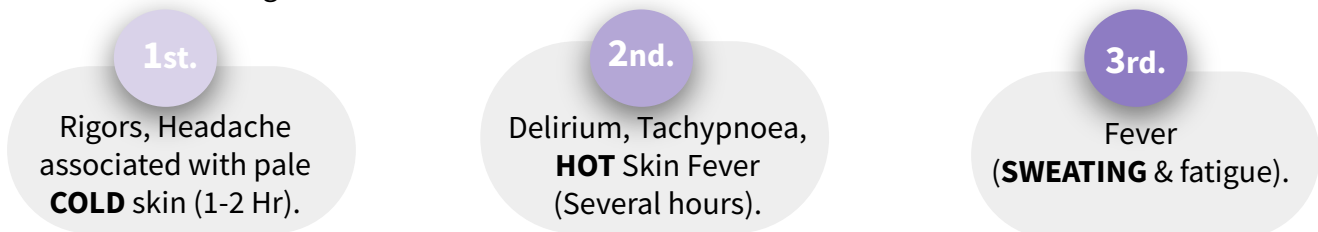
## Signs

- 01 Fever, HR tachycardia, low BP, low O2
- 02 Level of consciousness
- 03 Evidence of seizures
- 04 Splenomegaly the most common clinical sign
- 05 Jaundice
- 06 Anemia

Does this patient have malaria? <sup>1</sup>	
Acquired during travel:	Living in endemic region:
<ul style="list-style-type: none"> <li>• <b>Fever</b> +LR 5.1</li> <li>• <b>Splenomegaly</b> +LR 6.5</li> <li>• <b>Hyperbilirubinemia</b> +LR 7.3</li> <li>• <b>Thrombocytopenia</b> +LR 5.6</li> </ul>	<ul style="list-style-type: none"> <li>• Splenomegaly +LR 3.3</li> <li>• Hepatomegaly +LR 2.4</li> </ul>

## Malaria Paroxysm (attack) *437 slides*

- 3 successive stages:



### Notes Regarding Malaria Paroxysm:

- Paroxysms associated with **synchrony of merozoite release**.
- **Between paroxysms temperature is normal and patient feels well and Asymptomatic.**
- Falciparum may not exhibit classic paroxysms. (continuous fever)
- Classically the attacks (Periodicity):
  - ➔ Occur every **48** hours with (P. vivax, and P. ovale) “tertian parasites”.
  - ➔ Occur every **72** hours with (P.malariae) “quartan parasite”.
  - ➔ Whereas (P.falciparum) show **IRREGULAR** attacks , or hectic (especially in non-immune).

1) combination of these symptoms/ signs highly suggests malaria

# Malaria

## ◀ Malaria clinical spectrum:

- Most fever or flu-like illness
- Fatalities with (Pf) (occasionally with splenic rupture).
- **Cerebral malaria** (Pf) - may be focal or generalized symptoms.
  - ★ ○ risk factors for poor prognosis in cerebral malaria: high bilirubin, high creatinine, high lactase.
- **Hypoglycemia** - from disease and treatment (quinine leads to insulin release).
- **Risk Acute Respiratory Distress Syndrome** (Pf) due to capillary leak - need to limit fluids.
- **Acute renal failure** (Pf) - common complication of severe malaria.

## ◀ WHO Criteria – Severe Malaria

### Either

History of recent possible exposure and no other recognized pathology.

### Or

Asexual forms of (Pf) on blood smear.

### And one or more of the following 11 features:

- |  |   |
|--|---|
| <b>01</b>   Impaired consciousness or coma.                                | <b>07</b>   Spontaneous bleeding/disseminated intravascular coagulation.          |
| <b>02</b>   Severe normocytic anemia. Hb < 6                               | <b>08</b>   Repeated generalized convulsions.                                     |
| <b>03</b>   Renal failure.   | <b>09</b>   Acidemia/acidosis.  |
| <b>04</b>   Pulmonary edema or adult respiratory distress syndrome (ARDS). | <b>10</b>   Hemoglobinuria.   |
| <b>05</b>   Hypoglycemia.  | <b>11</b>   Parasitemia of > 5% (> 250 000/microlitre) in non-immune individuals. |
| <b>06</b>   Circulatory collapse, shock                                    |   |

## ◀ Pregnancy & Malaria *the extra from 437 slides*



- |           |  |
|-----------|--|
| <b>01</b> | Mortality, Abortion, Stillbirth.                             |
| <b>02</b> | Anemia, hypoglycemia, pulmonary edema: more common.          |
| <b>03</b> | Premature delivery high infant mortality.                    |
| <b>04</b> | Low birth weight ,Placental insufficiency.                   |
| <b>05</b> | High parasitemia? placenta favorable site for P. Falciparum. |
| <b>06</b> | Primigravidae , more susceptible to placental malaria        |



## ◀ Diagnosis

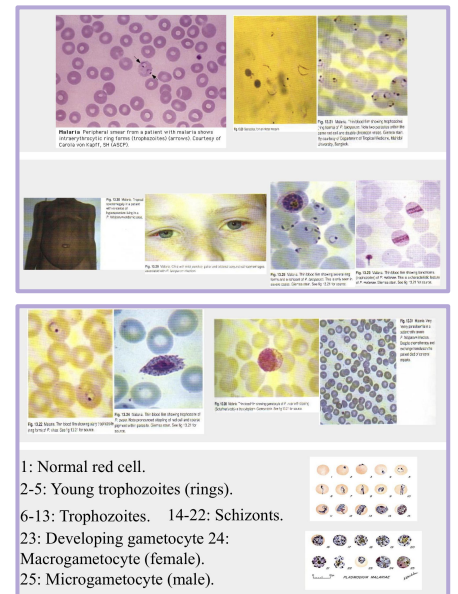
- Detailed targeted history including **travel history** and clinical examination together with High Index of Suspicion (HIS).

<b>Thin</b> (gold standard) <b>and Thick</b> (best initial) <b>films</b> <sup>1</sup>	<b>Urea and Creatinine</b>	<b>ABG</b>
<b>CBC, Coagulation profile</b>	<b>LFT, Bilirubin</b>	<b>CXR</b>
<b>Random Blood glucose</b>	<b>Lactic acid</b>	<b>Urine analysis</b>

### 01. Blood film

*Extra from 437 dr slides*

- Giemsa stain** or **wright's stain**.
- Correct identification of malarial Species is essential for treatment because of *P. Falciparum* is resistant to Chloroquine & others.
- **On Giemsa stain:**  
Cytoplasm: light blue, Nucleus: dark blue.
- **In *P. Falciparum*:**
  - Only ring stage asexual parasite and gametocytes seen in Peripheral Blood.**
  - While **RBC with Trophozoites or Schizonts** stage sequestered in peripheral microvasculature, and **NOT circulating Peripheral Blood.**
- **In *P. Vivax, Ovale & malariae*:**  
All asexual erythrocytic stages circulate in peripheral blood, thus seen on Blood Smear.
- ❖ **Acutely ill patients** | DDX: *P.F.* vs *P. Vivax*, because:
  - P. Ovale* – *Vivax* – clinical, morphological.
  - P. malariae* - ch. Infection



### Differential diagnosis of malaria in acutely ill patients based on peripheral blood smear: ★

Findings	<b><i>P. Falciparum</i></b>	<b><i>P. Vivax &amp; P. Ovale</i></b>
Multiple infected RBCs	Common	Rare
Mature (trophozoite & schizont) parasites	Absent	Common
RBC enlargement with later parasite stages	Absent	Common

- Mature (trophozoites & schizont) stage ***P. falciparum*** typically sequestered in the peripheral microvasculature.
- RBC enlargement in ***P. vivax*** typically occurs with later stage parasites that do NOT circulate in ***P. falciparum*** infection.

1. In the **thick** film, erythrocytes are lysed, releasing **all blood stages of the parasite**. This, as well as the fact that more blood is used in thick films, **facilitates the diagnosis of low-level parasitaemia**. A **thin** film is essential to **confirm** the diagnosis, to **identify the species** of parasite and, in *P. falciparum* infections, to quantify the **parasite load** (by counting the percentage of infected erythrocytes).

# Malaria

## 02. Serology

from 437 dr slides

- NOT useful in managing acutely ill patient.

## 03. DNA probe (PCR)<sup>1</sup>

- Limited utility for the diagnosis of acutely ill patients in the standard healthcare setting
- Most useful for confirming the species of malarial parasite after the diagnosis has been established by either smear microscopy or RDT (Rapid Diagnostic Test).

## ★ Management

### ◆ Basic Principles of Malaria Management:

- Think of the diagnosis and do **thick**(sensitivity) and **thin**(specificity) **blood films**.
- Ask lab about **species** of malaria – falciparum versus non-falciparum (vivax, ovale, malariae).
- Ask about percent parasitemia
  - If told **greater than 1% parasitemia** – think **PF!**.
- Verify no evidence of **severe or complicated malaria** (no need for parenteral therapy).
- Medical Emergency.
- Consider admission to hospital (especially for falciparum) at least observe tolerance of meds in ER.
- FOLLOW UP.
- **Drug of choice: Artesunate based therapy.**

Falciparum antimalarials	Essential features of general management	Falciparum antimalarials
<b>Uncomplicated:</b> a) Oral quinine 600mg/8r plus doxycycline 200mg daily (or clindamycin 450mg/8r) for 7 days OR b) Malarone® 4 standard tablets daily for 3 days OR c) Riamet® if weight >50kg, 4 tablets then 4 tablets at 0, 24, 36, 48 and 60 hours	<ul style="list-style-type: none"> <li>• Commence antimalarials immediately (see boxes)</li> <li>• <b>Severe malaria</b> <ul style="list-style-type: none"> <li>• Consider admission to high dependency/intensive care</li> <li>• Seek early expert advice from an infection or tropical unit</li> <li>• Oxygen therapy</li> <li>• Careful fluid balance (observe JVP, lying/standing BP and urine output). Avoid hyponatraemia. Over-hydration may induce pulmonary oedema; consider CVP monitoring</li> <li>• Monitor blood glucose regularly (especially during IV quinine)</li> <li>• ECG monitoring (especially during IV quinine)</li> <li>• 4-hourly observations until stable: i.e. pulse, temperature, BP, RR, SaO<sub>2</sub>, urine output &amp; GCS. Regular medical review until stable</li> <li>• Repeat FBC, clotting, U&amp;Es, LFTs and parasite count daily</li> <li>• In shock, treat for Gram negative bacteraemia</li> </ul> </li> </ul>	<b>Complicated or if patient is vomiting:</b> EITHER Quinine 20mg/kg loading dose (no loading dose if patient taking quinine or mefloquine already) as IV in 5% dextrose over 4hr and then 10mg/kg as IV over 4hr every 8hr plus oral doxycycline 200mg daily for 7 days (In pregnancy, use Woral clindamycin 450mg/8r). Max quinine dose 1.4 g OR if available, artesunate intravenously 2.4mg/kg at 0, 12, 24 hrs then daily to complete a course of seven days plus doxycycline or clindamycin as above When patient is stable & able to swallow, switch to oral quinine 600mg/8r plus doxycycline 200mg daily (or clindamycin 450mg/8r) to complete 7 days

### ● Oral therapy:

- Artemether-Lumefantrine<sup>2</sup>
- Atovaquone / Proguanil (Malarone)
- Quinine plus Doxycycline<sup>3</sup> / Clindamycin
- \* Avoid Mefloquine for treatment.



### ● Parenteral therapy

- Artesunate
- Quinine (or quinidine): needs telemetry.



**i** Box 20.57 Drug treatment of uncomplicated malaria

Parasite	Drugs	Regimen	Plus	Regimen
<i>Plasmodium vivax</i> <i>P. ovale</i> <i>P. malariae</i>	Chloroquine	600mg <sup>a</sup> 300mg <sup>b</sup> 6 h later 300mg 24 h later 300mg 24 h later	Primaquine	0.25–0.5 mg/kg per day for 2–3 weeks
	or (if known resistance to chloroquine, or dual infection with <i>P. falciparum</i> )			
	ACT (not artesunate + SP)	3 days	Primaquine	0.25–0.5 mg/kg per day for 2–3 weeks
<i>P. falciparum</i> (adults, endemic zone)	ACT	3 days	Primaquine	0.75 mg/kg single dose
	or (if not available)			
	Quinine + doxycycline	7 days	Primaquine	0.75 mg/kg single dose
<i>P. falciparum</i> (pregnant women)	All trimesters: Artesunate			
<i>P. falciparum</i> (infants)	ACT	3 days; appropriate dose for body weight	Primaquine	0.75 mg/kg single dose
<i>P. falciparum</i> (returning travellers)	Atovaquone–proguanil or quinine + doxycycline	7 days		

<sup>a</sup>10mg/kg in children. <sup>b</sup>5mg/kg in children. <sup>c</sup>Use ACT only if quinine is not available. ACT, artemisinin-based combination therapy; SP, sulfadoxine–pyrimethamine.

**+** Box 20.59 Drug treatment of severe *falciparum* malaria in adults and children

Drug/route	Immediate dose	Subsequent doses
Severe malaria is an emergency; after rapid assessment and confirmation of diagnosis, if possible, treatment should be started with whatever parenteral treatment is available. The options, in order of preference, are:		
1. Intravenous artesunate	2.4 mg/kg	2.4 mg/kg at 12 and 24 h, then daily (up to 7 days)
2. Intravenous quinine	20 mg/kg <sup>a</sup>	10 mg/kg 8-hourly (up to 7 days)
3. Intramuscular artesunate	2.4 mg/kg	2.4 mg/kg at 12 and 24 h, then daily (up to 7 days)
4. Intramuscular artemether	3.2 mg/kg	1.6 mg/kg daily
5. Rectal artesunate	10 mg/kg	Transfer to centre where parenteral therapy available
Continue parenteral treatment for at least 24 h, regardless of improvement in condition. After this, if the patient is improving, switch to oral therapy to complete 7 days with:		
ACT		
Or	Plus	Primaquine 0.75 mg/kg single dose
Quinine + doxycycline		

<sup>a</sup>10 mg/kg if patient has already received oral quinine or mefloquine. ACT, artemisinin-based combination therapy.

1. DNA detection (PCR) is used mainly in research and is useful for determining whether a patient has a recrudescence of the same malaria parasite or a re-infection with a new parasite.



**P. falciparum is now resistant to chloroquine and sulfadoxine-pyrimethamine (Fansidar) almost world-wide, so an artemisinin-based treatment is recommended.**

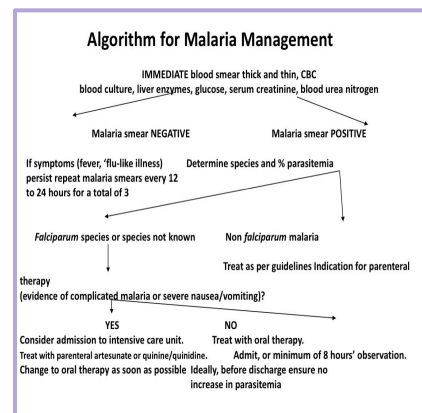
3. We usually **avoid doxy in pregnancy**, we don't use it in malaria as often as we used to. **Artesunate is the main therapy**

# Malaria

## Management cont.

### Non-P. Falciparum:

- **Chloroquine** (base) 600mg followed by 300mg at 6, 24 and 48 hours.
- **primaquine** (30mg base/day for vivax, 15 mg/day for ovale) for 14 days in **vivax and ovale** after treatment of acute infection use to **eradicate liver parasites; G6PD must be measured** before primaquine is given – seek expert advice if low.



13.59 Severe manifestations/complications of <i>falciparum</i> malaria and their immediate management	
<b>Coma (cerebral malaria)</b> <ul style="list-style-type: none"> <li>• Maintain airway</li> <li>• Nurse on side</li> <li>• Exclude other treatable causes of coma (e.g. hypoglycaemia, bacterial meningitis)</li> <li>• Avoid harmful ancillary treatments such as corticosteroids, heparin and adrenaline (epinephrine)</li> <li>• Intubate if necessary</li> </ul>	<b>Spontaneous bleeding and coagulopathy</b> <ul style="list-style-type: none"> <li>• Transfuse screened fresh whole blood (cryoprecipitate/fresh frozen plasma and platelets if available)</li> <li>• Vitamin K injection</li> </ul>
<b>Hyperpyrexia</b> <ul style="list-style-type: none"> <li>• Tepid sponging, fanning, cooling blanket</li> <li>• Antipyretic drug (paracetamol)</li> </ul>	<b>Metabolic acidosis</b> <ul style="list-style-type: none"> <li>• Exclude or treat hypoglycaemia, hypovolaemia and Gram-negative septicaemia</li> <li>• Fluid resuscitation</li> <li>• Give oxygen</li> </ul>
<b>Convulsions</b> <ul style="list-style-type: none"> <li>• Maintain airway</li> <li>• Treat promptly with diazepam or paraldehyde injection</li> </ul>	<b>Shock ('algid malaria')</b> <ul style="list-style-type: none"> <li>• Suspect Gram-negative septicaemia</li> <li>• Take blood cultures</li> <li>• Give parenteral antimicrobials</li> <li>• Correct haemodynamic disturbances</li> </ul>
<b>Hypoglycaemia</b> <ul style="list-style-type: none"> <li>• Measure blood glucose</li> <li>• Give 50% dextrose injection followed by 10% dextrose infusion (glucagon may be ineffective)</li> </ul>	<b>Aspiration pneumonia</b> <ul style="list-style-type: none"> <li>• Give parenteral antimicrobial drugs</li> <li>• Change position</li> <li>• Physiotherapy</li> <li>• Give oxygen</li> </ul>
<b>Severe anaemia (packed cell volume &lt; 15%)</b> <ul style="list-style-type: none"> <li>• Transfuse fresh whole blood or packed cells if pathogen screening of donor blood is available</li> </ul>	<b>Hyperparasitaemia</b> <ul style="list-style-type: none"> <li>• Consider exchange or partial exchange transfusion, manual or haemophoresis (e.g. &gt; 10% of circulating erythrocytes parasitised in non-immune patient with severe disease)</li> </ul>
<b>Acute pulmonary oedema</b> <ul style="list-style-type: none"> <li>• Nurse at 45°, give oxygen, venesect 250 mL of blood, give diuretic, stop intravenous fluids</li> <li>• Intubate and add PEEP/CPAP (p. 193) in life-threatening hypoxaemia</li> <li>• Haemofilter</li> </ul>	<b>Specific therapy</b> <ul style="list-style-type: none"> <li>• Intravenous artesunate</li> <li>• Mefloquine should be avoided due to increased risk of post-malaria neurological syndrome</li> </ul>
<b>Acute renal failure</b> <ul style="list-style-type: none"> <li>• Exclude pre-renal causes</li> <li>• Fluid resuscitation if appropriate</li> <li>• Peritoneal dialysis (haemofiltration or haemodialysis if available)</li> </ul>	

From WHO. Severe falciparum malaria. In: Severe and complicated malaria. 3rd edn. Trans Roy Soc Trop Med Hyg 2000; 94 (suppl. 1): S1–41.

## Drug toxicity

- **Quinine:**
  - Hypoglycemia, arrhythmias, bitter taste, GIT upset, nausea, vomiting, tinnitus, high tone deafness.
- **Doxycycline:**
  - GI upset, vaginal candidiasis.
- **Mefloquine<sup>1</sup>:**
  - Neuropsychiatric symptoms (mood changes), encephalopathy... transient.
- **Artemether-Lumefantrine:**
  - H/A, anorexia, dizziness, arthralgia and myalgia.

1. Contraindicated in the first trimester of pregnancy, lactation, cardiac conduction disorders, epilepsy, psychiatric disorders.

## Prevention

### When travelling to malaria endemic areas:

- Avoid mosquito bites .
- Wear long sleeved shirts and long trousers.
- Sleep in well-screened or air-conditioned rooms.
- Permethrin impregnated bed nets.
- Use bednetting of good quality with small mesh that is not damaged and impregnated with permethrin.
- Apply insect repellent containing no more than 30% DEET, or use 20% or greater Picaridin.
- At dusk, spray aerosolized insecticides (such as those containing pyrethrins) in living and sleeping areas.
- Chemoprophylaxis.

### Chemoprophylaxis<sup>1</sup>:

#### Start 2 days pre-travel, continue 7 days after return: (most effective)

→ Atovaquone/ Proguanil<sup>2</sup> (Malarone): 1 tab/d ( 250 mg atovaquone /100 mg proguanil)

#### One or 2 weeks pre-travel, continue 4 weeks after return: less preferred

- Mefloquine 250 mg once/wk.
- Doxycycline 100 mg daily.
- Primaquine 30 mg base daily.
- Chloroquine<sup>3</sup> sensitive areas: 500 mg (300 mg base) : once/wk.

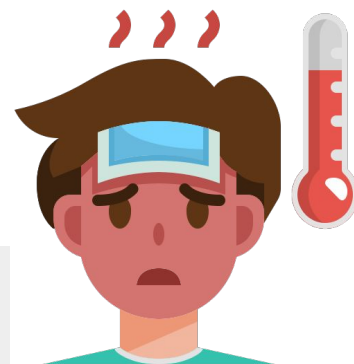
1. Choice of regimen is determined by area to be visited, length of stay, level of malaria transmission, level of drug resistance, presence of underlying disease in the traveler concomitant medication taken.
2. **Pregnant** and lactating women may take **proguanil or chloroquine** safely. Avoid Malarone in pregnancy.
3. **Chloroquine** should not be taken continuously as a prophylactic for more than 5 years without regular ophthalmic examination, as it may cause **irreversible retinopathy**.

### Malaria Vaccine:

- RTS,S/AS01 (Mosquirix).
- Engineered from T-cell epitope in the pre-erythrocytic circumsporozoite protein (CSP) of PF malaria parasite and the envelope protein of HBsAg with an adjuvant AS01.
- Efficacy of 25-50% in **infants and young children**. given to children of endemic areas

## ◀ Fever in Returning Travelers

❖ The **most common** etiologies of fever in returning travelers are listed first:



### 01 Dengue (flavivirus):

- **4-7 day** incubation, widespread in tropics (increasing in Africa), vector borne (Aedes).
- **Diagnosis:** PCR, NS1 antigen; IgM only after day 4.

### 02 Malaria (Plasmodia sp.):

- **7-30 day** incubation for P. falciparum, widespread but very high risk in Africa, vectorborne. (Anopheles).
- **Diagnosis:** blood smear, PCR, malaria RDT (rapid diagnostic test).

### 07 Leptospirosis:

- Incubation **2-29 days**, South and South Asia, South America, transmitted from urine of infected rodents.
- **Diagnosis:** IgM only after day 5; PCR earlier.

### 03 Typhoid fever (Salmonella sp.):

- Incubation **6-30 days**, mostly South and Southeast Asia, fecal-oral transmission.
- **Diagnosis:** blood culture. Avoid Widal test **can give a false -ve or false +ve.**

### 08 Rickettsial disease:

- **Scrub typhus:**
  - Incubation **6-20 days**, Asia and northern Australia, chigger mites. Murine typhus worldwide.
  - **Diagnosis:** PCR (blood or eschar), IgM.
- **Spotted fever group:**
  - Incubation **2-14 days**, widespread but highest risk with R. Africae in southern Africa, mostly ticks.
  - **Diagnosis:** PCR (blood or eschar), IgM/IgG.

### 04 Chikungunya virus:

- **4-7 day** incubation, widespread in tropics, vector borne (Aedes)
- **Diagnosis:** PCR; IgM only after day 4.

### 05 Zika virus:

- Incubation **3-14 days**, mostly Southeast Asia, little risk in Americas and Caribbean since 2016, vector borne (Aedes).
- **Diagnosis:** PCR of blood or urine, IgM (high false-positive rate), Viral Neutralization is definitive, no testing available in someone with previous flavivirus infection of any kind.

### 09 Rabies:

- Incubation **20-60 days** most common, widespread, animal or **bat bite**.
- **Diagnosis:** PCR of saliva or skin, serum antibody detection.

### 06 COVID-19 (SARS-CoV-2):

- Incubation most commonly **3-7 days**, present in almost all countries, can be acquired in-flight or after return.
- **Diagnosis:** PCR more sensitive than antigen, very sensitive during febrile phase.

### 10 East African trypanosomiasis:

- Incubation **7-21 days**, eastern and southern African game parks, **tsetse fly**.
- **Diagnosis:** microscopy of blood films, lymph node aspirate, or chancre.



## ◀ Fever in Returning Travelers

- ❖ Other **frequent diseases** in travelers:
  - influenza A (seasonal).
  - Acute HIV
  - Mononucleosis.
  - Measles.
  - Varicella.
  - Tuberculosis.
- ❖ **Infrequent diseases** in travelers to consider:
  - Avian influenza (H5N1 & H7N9).
  - African hemorrhagic fevers (Ebola incubation is 8-12 days).
  - CCHF (Crimean-Congo haemorrhagic fever).
  - Yellow fever.
  - Japanese encephalitis.
  - Monkeypox (Nigeria and neighbors).
  - Relapsing fever.
  - Acute toxoplasmosis.
  - Arboviruses including Ross River.
  - Tickborne encephalitis.
  - West Nile virus (emerging in Europe).
  - Hantavirus (both hantavirus cardiopulmonary syndrome and hemorrhagic fever with renal syndrome).



## ◀ Vaccines

### Required:

- Yellow Fever in Africa.
- Meningococcal, SA, required for Hajj
- COVID-19 for international travel.

### Recommended :

- Polio
- Influenza
- Hepatitis A/B
- Measles
- Tetanus/Diphtheria/Pertussis
- Typhoid
- Rabies
- Japanese Encephalitis
- Tick-borne Encephalitis

Journal of Nature and Science of Medicine  
(An International Peer-Reviewed Journal)

COVID-19 Vaccine in the Kingdom of Saudi Arabia: A True Operation Warp Speed  
Mazin Berry<sup>1</sup>, Ahmed S. Bahammam<sup>2\*</sup>

Manufacturer	Vaccine Name	Vaccine Platform	Completed Clinical Trial	Efficacy	Most common Adverse Effects	Countries Authorized/rolled out the vaccine
Pfizer/BioNTech	BNT162b2	Modified mRNA	Phase III	95%	Local site pain, swelling	UK (Dec 2) Bahrain (Dec 4) Canada (Dec 9) KSA (Dec 10)
Moderna	mRNA-1273	Modified mRNA	Phase III	94%	Local site pain, swelling	USA (Dec 18) Canada (Dec 23) Israel (Jan 4) EU (Jan 6)
Oxford/AstraZeneca	ChAdOx1 nCov-19	Adenoviral vector encoding Spike protein	Phase III	62%-90%	Pain and swelling	UK (Dec 30) Argentina (Dec 30) El Salvador (Dec 30) India (Jan 5) Mexico (Jan 4) Dom. Rep. (Jan 6) Morocco (Jan 6)

<https://www.insmonline.org/> ISSN 2589-6288



\* How Do You Know What Vaccines are Needed for Travel? The CDC's Health Information Website for Clinicians and Travelers (Published every 2 years, The "yellow book") Available online.

# Yellow Fever

for the next infections the dr only focused on the vaccines and skipped the remaining

## Yellow fever<sup>1</sup>

- Mosquito-borne hemorrhagic fever caused by a flavivirus.
- **Rare fatalities in travelers since vaccine introduction (very effective)**
- ❖ **Epidemiology:**

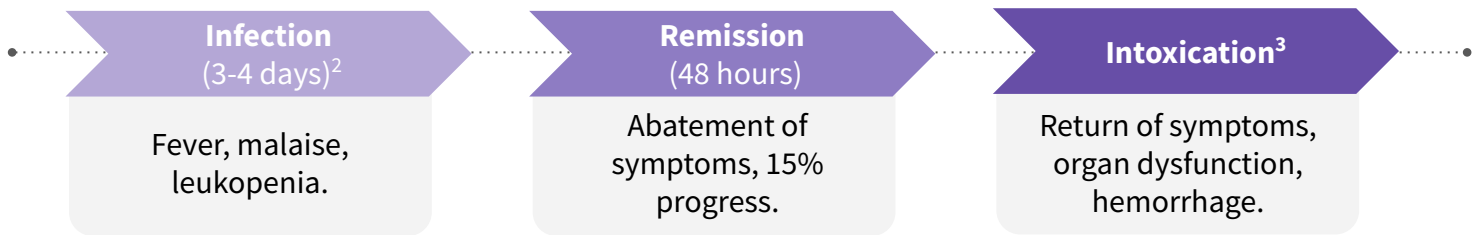


~200,000 cases per year, 90% in Africa

Indigenous case fatality rates vary **20-60%** (very high in endemic areas, however, the vaccine reduced fatality in travelers significantly)

## Stages

- Has **3** stages:



## Transmission

- 1 From primates or humans
- 2 Mosquito vector

## Disease Prevention

- ▶ Avoid mosquito bites: DEET, Clothing, Mosquito nets, Eliminate standing water.
- ▶ Vaccination

## Yellow Fever Vaccine

- Live-attenuated vaccine. Developed in 1936.
- Seroconversion >95%.
- Single 0.5ml subcutaneously.
- Revaccination at 10-year intervals required by World Health Organization.
  - Protection from one vaccine. However, may last 30 or more years.

### Contraindications



- Age <9 months old
  - (Can consider at 6-9 months old during outbreaks).
- Pregnant women
  - (Yellow fever can cross placenta).
- Severe egg allergies.
- Severe immunocompromise.
- Immunomodulatory drugs.

1. Diagnosis of yellow fever can be confirmed by viral isolation from blood in the first 24 days of illness, the presence of IgM or a fourfold rise in IgG antibody titre. Leucopenia is characteristic.
2. Humans are infectious during the viraemic phase, which starts 3-6 days after the bite of the infected mosquito and lasts for 4-5 days.
3. In more severe disease, fever recrudescence is associated with lower back pain, abdominal pain and somnolence, prominent nausea and vomiting, bradycardia and jaundice. Liver damage and DIC lead to bleeding with petechiae, mucosal haemorrhages and gastrointestinal bleeding. Shock, hepatic failure, renal failure, seizures and coma may ensue.

# Yellow Fever

## Yellow Fever Vaccine



### Side Effects

Adverse Reactions (10-30%):  
Local soreness, Mild fever, Headache, Myalgia.

### Rare Severe Reactions

- **Anaphylaxis:**
    - Risk 1/131,000.
  - **Yellow fever associated neurotropic disease (YEL-AND):**
    - Risk 1:150,000 - 200,000.
    - Multiple neurologic conditions:
      - Encephalitis (esp. infants <9 months), Guillian-Barre, Bell's Palsy.
    - Onset 2-28 days after vaccination.
    - Rarely fatal.
  - **Yellow fever associated viscerotropic disease (YEL-AVD):**
    - Mimics severe yellow fever infection.
    - Major organ system failure occurs:
      - Hepatic, renal, circulatory failure.
      - 50% or greater fatality rate.
    - Occurs 1-8 days (average 3 days) after initial vaccination.
    - Risk 1:200,000 - 300,000 (Greater risk if over age 60).
- Certification of vaccination required:
    - International Certificate of Vaccination or Prophylaxis for Yellow Fever form (ICVP).
    - Must be signed by licensed physician or designee.
  - Waiver form for medical contraindication to vaccine (e.g. pregnancy).

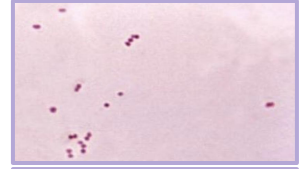
- |   |  |
|---|--|
| • Angola  | • French Guiana  |
| • Benin   | • Gabon  |
| • Bolivia (or signed affidavit at point of entry) | • Ghana  |
| • Burkina Faso                                    | • Liberia  |
| • Burundi   | • Mali   |
| • Cameroon  | • Niger  |
| • Central African Republic                        | • Rwanda   |
| • Congo, Republic of the                          | • São Tomé and Príncipe  |
| • Côte d'Ivoire                                   | • Sierra Leone   |
| • Democratic Republic of Congo                    | • Togo   |
|   | • Always check up to date list at <a href="http://www.cdc.gov/travel">www.cdc.gov/travel</a> |

**Yellow Fever Vaccination Proof Required for Entry**

# Meningococcal

## ◀ Meningococcal Disease

- 1st Picture: **Neisseria Meningitidis** (Gram negative diplococci).
- 2nd Picture: Sub-Saharan Africa.
- Youngest children = highest risk.
- ❖ **Epidemiology:**
  - Greatest risk: dry season (Dec. - June).
  - **Hajj pilgrimage to Saudi Arabia associated with outbreaks.**



0.5-10/100,000 in non-epidemic areas.

Risk of travelers: 0.4/100,000.

Up to 1,000/100,000 in epidemic areas.

## ◀ Meningococcal vaccine

- **Vaccine required to attend the Hajj (annual pilgrimage to Mecca)**
  - If under age 15, polio vaccination needed also.

### ❖ Available vaccines:

#### MCV4

(Menactra™)

- Quadrivalent meningococcal polysaccharide-protein conjugate vaccine.
- 2-55 years old.
- Preferred in <11 year olds.

#### MPVS4

(Menomune®)

- Quadrivalent meningococcal polysaccharide vaccine
- 2 years and older.
- Use for >55 years old.

#### MenACWY-CRM

(Menveo®)

- Quadrivalent meningococcal oligosaccharide diphtheria CRM197 Conjugate Vaccine.
- 11-55 years old.

- ❖ **Revaccination:** If high-risk (epidemic area or travel).

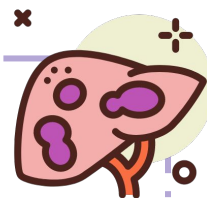
If vaccine given at **2-6 years** old:  
Repeat after **3** years, then every **5** years.

If vaccine given **>6 years** old:  
Repeat every **5** years.

*An area for your notes*

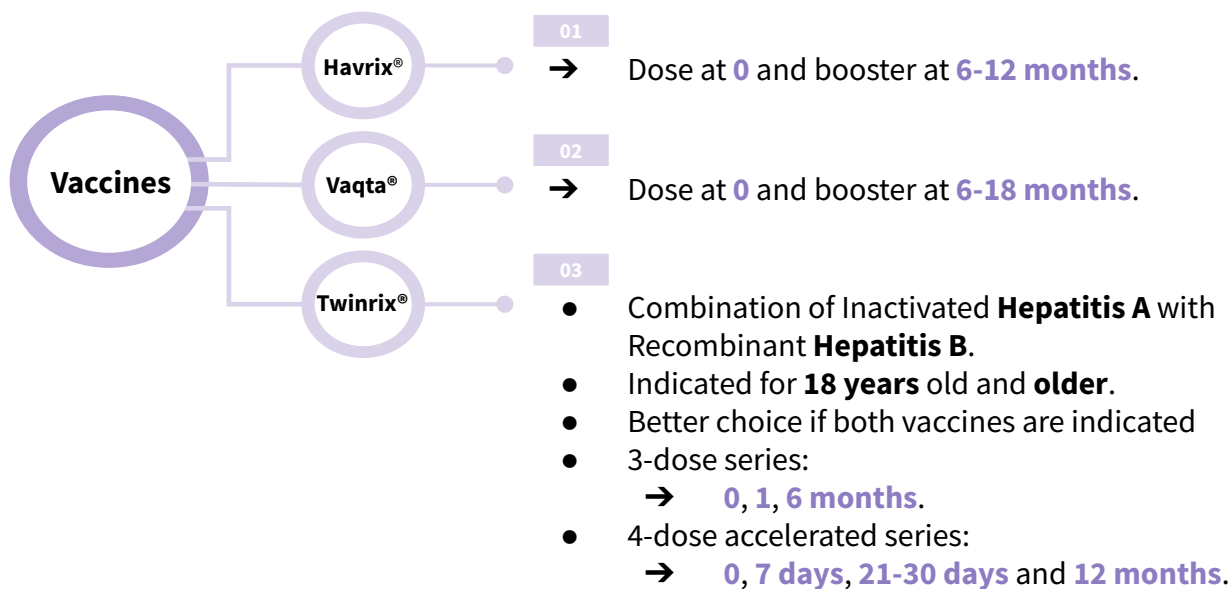
## ◀ Hepatitis A

- Worldwide prevalence.
- Adults often contract from asymptomatic children.
- Incubation 28 days (range 15-50 days).
- Viral shedding 2 weeks before to 1 week after symptoms.
- Usually self-limited disease.
- ❖ **Mode of transmission:**
  - Fecal/oral transmission ( Associated poor hygiene or sanitation).
- ❖ **Symptoms include:**
  - Jaundice
  - Fatigue
  - Abdominal pain
  - Anorexia
  - Nausea



## ◀ Hepatitis A Vaccine

- Inactivated Hep A virus (**Havrix®** or **Vaqta®**).
- Combined with Hepatitis B (**Twinrix®**).
- ❖ **Travel vaccine indications:**
  - Anyone **>1 year old** traveling anywhere other than to:
    - ◆ U.S. and Canada, Western Europe, Scandinavia, Japan, Australia and New Zealand.
  - For healthy patients **<40 years** old, one dose before travel confers adequate protection.



## ◀ Hepatitis A treatment

- ❖ **Consider immunoglobulin treatment for patients:**
  - Leaving in less than two weeks.
  - Older.
  - Immunocompromised.
  - Chronic medical conditions.
  - Under 12 months of age.



# Influenza

Skipped by the dr

## Risk factors

- ❖ **Risk depends on timing and destination:**
  - Tropics: year round risk.
  - Temperate climates: risk generally April-September.
- ❖ **Avian subtype risks:**
  - Visiting poultry farms.
  - Visiting open markets where live poultry are present.
  - Eating undercooked poultry products (eggs, meat, etc.).



## Preventative measures



Hygiene: washing hands.



Annual vaccination.

# Typhoid Fever

## Definition

Acute **life-threatening illness**. Caused by *Salmonella typhi*<sup>1</sup>.

## Epidemiology



**22,000,000** cases worldwide/year.  
**200,000** deaths.



### Risk area:

- Southeast Asia
  - 6-30 times more common.
  - Highest risk of FQ (Fluoroquinolones) drug resistance.
- Africa, Caribbean, Central and South America.
- Length of stay = increased risk.



## Mode of transmission

- Humans – only source:
  - Acquired through fecal contamination of food and water.

1. After a few days of bacteraemia, the bacilli localise, mainly in the lymphoid tissue of the small intestine, resulting in typical lesions in the Peyer's patches and follicles. These swell at first, then ulcerate and usually heal. After clinical recovery, about 5% of patients become chronic carriers (i.e. continue to excrete the bacteria after 1 year); the bacilli may live in the gallbladder for months or years and pass intermittently in the stool and, less commonly, in the urine.

# Typhoid Fever

## Signs & symptoms<sup>1</sup>

- Incubation period: **6-30 days**.
- Headache.
- Malaise.
- Fever.
- Increasing in severity.
- Low-grade septicemia.
- **“Rose spots” on trunk.**



**Rose spots on the chest in a patient with typhoid.**  
At the end of the first week, a rash may appear on the upper abdomen and on the back as sparse, slightly raised, rose-red spots, which fade on pressure.

## Complications

- Serious complications (**2-3 weeks**):
  - **Hepatosplenomegaly.**
  - **Intestinal hemorrhage/perforation.**

## Treatment

- Oral rehydration.
- Antibiotics:
  - **3rd generation cephalosporin** (10-14 days).
  - Azithromycin Ciprofloxacin only if no resistance (7-10 days).
- Steroids in severe cases.

## Prevention

- ▶ Avoid contaminated food and water.
- ▶ Hygiene.
- ▶ Local cuisine.

## Vaccines

- **2 available vaccines:**

### Vivotif®

- **Oral, live-attenuated.**
- Ages **6** and **older**.
- 50-80% protection.
- 4 pills – one every other day.
- Completed **1 week** before potential exposure.
- Revaccination **every 5 years**.

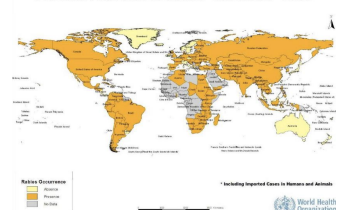
### Typhim Vi®

- **Capsular polysaccharide (IM).**
- Ages **2** and **older**
- 50-80% protection.
- Single 0.5ml injection.
- **2 weeks** before exposure.
- Booster **every 2 years**.

# Rabies<sup>2</sup>

- Found globally.
- ❖ **Consider vaccination:**
  - If potential exposure to wild animals (especially dogs).
  - Prolonged exposure where endemic.

Presence / Absence of Rabies\* Worldwide - 2005



## Pre-exposure prophylaxis<sup>3</sup>

- Series of 3 at **0, 7** and **21-28 days**.
- 2 vaccines available:
  - Imovax®
  - Rabavert®

## Post-exposure

- **Rabies Immunoglobulin (RIG) plus vaccine:**
  - RIG days **0, 4**.
  - Vaccine days **0, 3, 7, 14**.
- If had vaccine:
  - No RIG needed.
  - Vaccine days **0** and **3**.

1. The temperature rises in a stepladder fashion for 4 or 5 days with malaise, increasing headache, drowsiness and aching in the limbs. Constipation may be caused by swelling of lymphoid tissue around the ileocaecal junction, although in children diarrhoea and vomiting may be prominent early in the illness.

2. Rabies is caused by a rhabdovirus that infects the central nervous tissue and salivary glands of a wide range of mammals, and is usually conveyed by saliva through bites or licks on abrasions or on intact mucous membranes. Humans are most frequently infected from dogs and bats.

3. Pre-exposure prophylaxis is required by those who handle potentially infected animals professionally, those who work with rabies virus in laboratories and those who live at special risk in rabies-endemic areas.

# Japanese Encephalitis

## Japanese Encephalitis Virus (JEV)

- Most common cause of encephalitis in Southeast Asia.
- Carried by mosquitoes
- ❖ **Risk:**
  - Little risk in urban areas.
  - Mostly rural areas.
  - Not recommended for short-term travel to urban area.



Geographic distribution in Southeast Asia.

## Signs & symptoms<sup>1</sup>

- Incubation **5-15 days**.
- Most infections asymptomatic, <1% develop clinical disease:
  - Headache, fever, vomiting, diarrhea.
    - Most recover in 1 week.
- 1:300 severe symptoms with 30% fatality:
  - Mental status changes.
  - Focal neurological deficits.
  - Parkinsonian syndrome.
  - Seizures (especially children).

## vaccine

- Inactivated **Vero cell culture (JE-VC)**:
  - For people **over 17 years** old.
  - Duration of protection unknown.
  - Need for boosters undetermined.
  - Pregnancy Category B.

## Travelers' Diarrhea Self-treatment

### Antibiotic +/- loperamide

**Quinolones:** single dose, max 3 days.

**Azithromycin:** single dose of 500mg max 3 days or one time dose of 1000 mg.

**Rifaximin**

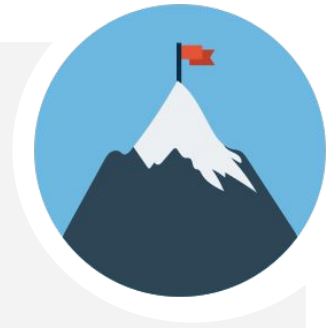
**Giardiasis (giardia duodenalis):** rotten eggs burping, diarrhea.

**Metronidazole:** 250 mg po TID 7 d.

**Nitazoxanide:** 500 mg po bid 3 days

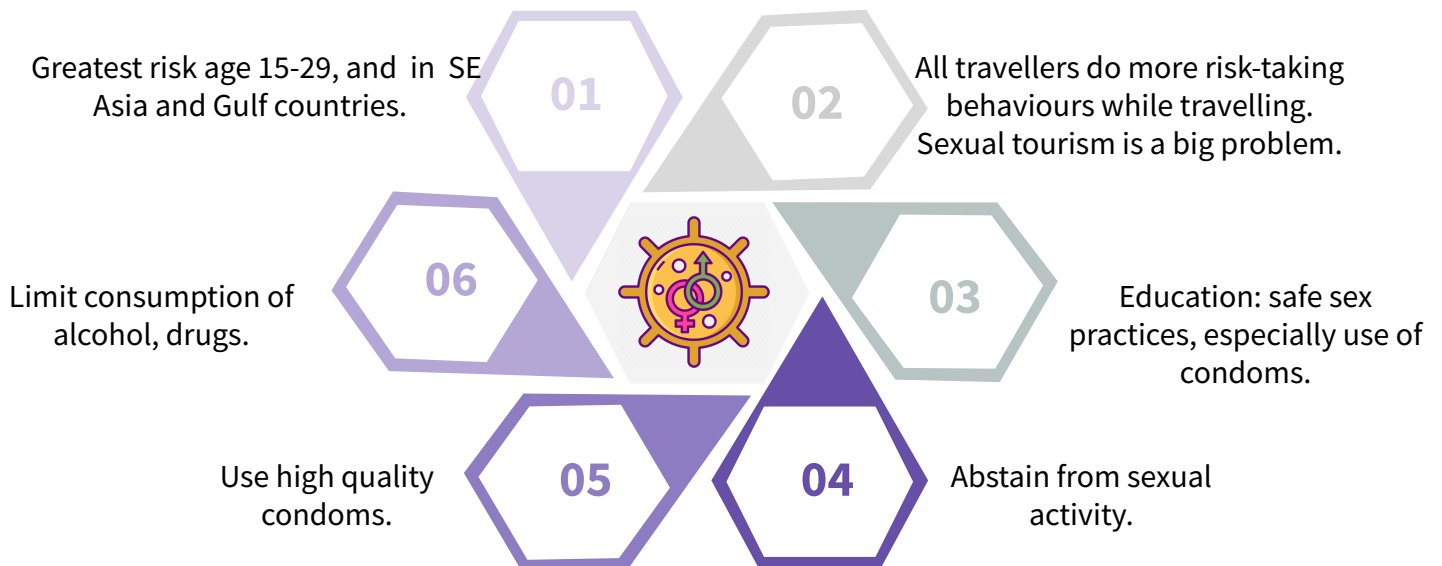
1. Initial systemic illness with fever, malaise and anorexia is followed by photophobia, vomiting, headache and changes in brainstem function. Neurological features other than encephalitis include meningitis, seizures, cranial nerve palsies, flaccid or spastic paralysis, and extrapyramidal features. Mortality with neurological disease is 25%.

# Advice for Acclimatization



- Tell climbers to avoid abrupt ascent to altitudes above 9850 ft.
- Spend 2-3 nights at 8200-9850 ft before further ascent.
- Add an extra night of acclimatization for every 2000-3000 ft of ascent.
- Make day trips to higher elevation with return to lower elevation for sleep.
- Avoid alcohol and sedatives for the first 2 nights at altitude.
- Moderate exercise (Extreme exercise at altitude may be harmful).
- **acetazolamide** 125-250 mg bid starting 24 hours before ascent and continue for 48 hrs at maximum altitude. to reduce the risk of pulmonary and vertebral edema
- Side effects: paresthesias, polyuria, nausea, drowsiness, impotence, myopia, bitter taste.

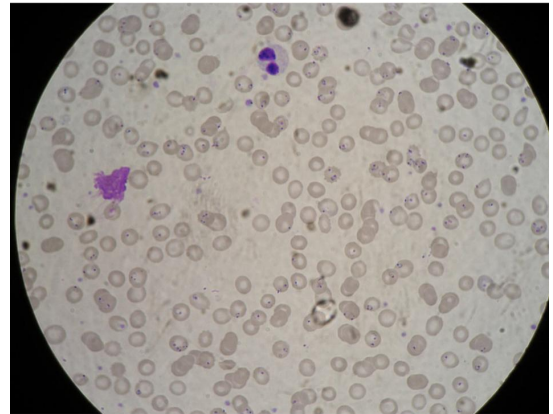
# Sexually transmitted illnesses and Travel



## Case study 1:

- ❖ 18 year old pregnant woman returned from **Jazan** one week ago. Fever and headache for 3 days.  
**Physical examination:** Pale, Temp. 39°C Spleen enlarged.  
**Investigations:** WBC 8, Hb 9.0, MCV 93, Plt 90, bilirubin 52

- **What is your diagnosis?**  
*P.falc malaria showing double chromatin dotted ring and multiple invasions in one RBC*
- **What are the risks in her condition?**
  - Severe disease.
  - Primigravida.
  - Risk of low birth & abortion.
  - Risk of low glucose , pulmonary oedema.



## Case study 2:

- ❖ 42 Saudi Man with continuous fever for one month Chills, rigors, weight loss 2 weeks of mid-back pain radiating to flanks 1 month prior to onset of fever traveled to Thailand, visited rice fields. He diagnosed with **Melioidosis**.

- **What do you see in the picture?**  
 Infrarenal abdominal aortic pseudoaneurysm.
- **What do you expect to see in the blood culture?**  
 Burkholderia pseudomallei causative agent of melioidosis.





# Summary

<p><b>Epidemiology</b></p>	<ul style="list-style-type: none"> <li>● <b>Stable transmission:</b> including much of <b>sub-Saharan Africa</b>, transmission occurs consistently year round. The bulk of the mortality is seen in <b>children</b>, while those who survive to adulthood acquire significant immunity; <b>low-grade parasitaemia</b> is still present but causes few symptoms.</li> <li>● <b>Unstable transmission:</b> <b>seasonal or low-level transmission</b> (e.g. in the <b>Sahel belt</b>, where mosquitoes feed only in the rainy season). <b>Little protective immunity</b> develops and symptomatic malaria occurs at <b>all ages</b>.</li> </ul>
<p><b>Pathogenesis</b></p>	<ul style="list-style-type: none"> <li>● RBCs invasion. , Microvascular pathology, Renal failure, Deep coma, Pulmonary edema , Immune-complex Nephrotic syndrome.</li> </ul>
<p><b>Plasmodium Falciparum</b></p>	<ul style="list-style-type: none"> <li>- <b>Resistant</b> to many antimalarial drugs.</li> <li>- <b>Most morbidity and mortality.</b></li> <li>- Infects <b>mature</b> and young erythrocytes.</li> <li>- Cytoadherence.</li> <li>- <b>Schizogony</b> is particularly prolific in all stages.</li> <li>- Infection in the peripheral blood &gt; <b>ring forms</b> and <b>gametocytes</b> “crescent-shape”.</li> <li>- Travel history up to 1 year (usually first 3 months), may present as continuous fever.</li> </ul>
<p><b>p.Vivax</b></p>	<ul style="list-style-type: none"> <li>- Benign’ <b>tertian malaria.</b></li> <li>- Restriction of erythrocyte invasion to <b>reticulocytes bearing Duffy blood antigen</b></li> <li>- <b>Schüffner’s dots.</b></li> <li>- <b>Hypnozoites</b> &gt; remain latent responsible for subsequent relapse.</li> </ul>
<p><b>p.Ovale</b></p>	<ul style="list-style-type: none"> <li>- Can <b>infect Duffy negative</b> reticulocytes.</li> <li>- <b>Late-onset vivax and ovale</b> malaria may occur despite effective prophylaxis.</li> <li>- <b>Hypnozoites</b> &gt;remain latent responsible for subsequent relapse.</li> </ul>
<p><b>P. Malariae</b></p>	<ul style="list-style-type: none"> <li>- <b>Slow development.</b></li> <li>- The asexual cycle is <b>72 hours</b> ‘<b>quartan malaria</b>’.</li> <li>- Infects old erythrocytes &gt;‘<b>smaller</b>’ infected cells.</li> <li>- <b>Do not transform into hypnozoites &gt; no relapses.</b></li> <li>- The peripheral blood (<b>10 years or more</b>) at a very low level of parasitaemia &gt;detectable peaks with a recrudescence of clinical symptoms.</li> <li>- Travel history up to 10 years.</li> </ul>
<p><b>Treatment</b></p>	<ul style="list-style-type: none"> <li>● <b>Oral therapy:</b> <ul style="list-style-type: none"> <li>→ Artemether-Lumefantrine.</li> <li>→ Atovaquone / Proguanil (Malarone).</li> <li>→ Quinine plus Doxycycline/Clindamycin</li> <li>-Avoid Mefloquine for treatment. -</li> </ul> </li> <li>● <b>Parenteral therapy:</b> <ul style="list-style-type: none"> <li>→ Artesunate.</li> <li>→ Quinine (or quinidine): needs telemetry.</li> </ul> </li> <li>● <b>Non-P. Falciparum:</b> <ul style="list-style-type: none"> <li>- <b>Chloroquine</b> at 6, 24 and 48 hours.</li> <li>- <b>Primaquine</b> for 14 days: In <b>vivax and ovale</b> after treatment of acute infection use to <b>eradicate liver parasites;</b> G6PD must be measured before primaquine.</li> </ul> </li> </ul>
<p><b>Chemoprophylaxis</b></p>	<ul style="list-style-type: none"> <li>● <b>start 2 days pre-travel, continue 7 days after return:</b> <ul style="list-style-type: none"> <li>→ Atovaquone/ Proguanil (Malarone): 1 tab/d.</li> </ul> </li> <li>● <b>One or 2 weeks pre-travel, continue 4 weeks after return:</b> <ul style="list-style-type: none"> <li>→ Mefloquine: once/wk.</li> <li>→ Doxycycline: 100 mg daily.</li> <li>→ Primaquine: base daily.</li> <li>→ Chloroquine: sensitive areas.</li> </ul> </li> </ul>

# Lecture Quiz

**Q1: A 55 year old male presented to you complaining of recurring fevers rigors vomiting diarrhea and confusion. Three weeks ago he came back from a 2 week trip to Kenya and Tanzania. On the physical exam his HR is 110 BP 105/64. His spleen is tender. HB: 10, WBC 3.5, Platelets 7 Blood smear shows 25% parasitemia. Only ring form is seen on microscopy Which of the following is the causative organism?**

- A. Plasmodium Falciparum
- B. Plasmodium Vivax
- C. Plasmodium Knowlesi
- D. Plasmodium Malariae

**Q2: 34-year-old pregnant female was recently diagnosed with malaria. The organism is P. ovale. What is the treatment of choice for hypnotic infection in this case?**

- A. Doxycycline
- B. Primaquine
- C. Chloroquine
- D. Mefloquine

**Q3: An old man who came back from his holiday in Africa, 2 days ago he presented with fever and 4% parasitemia. Falciparum malaria is suspected. What is the treatment of choice?**

- A. Artesunate
- B. Sulfadoxine-pyrimethamine
- C. Primaquine
- D. Chloroquine

**Q4: A 52 years old male suspected to have malaria. Peripheral blood film smear was done and showed mature trophozoite stage. What is the less likely causative organism?**

- A. P. Vivax
- B. P. Ovale
- C. P. Falciparum
- D. P. Malariae

**Q5: Which of the following can be used in case of uncomplicated falciparum infection**

- A. Atovaquone-proguanil
- B. Chloroquine
- C. Hydroxychloroquine
- D. Penicillin G

**Q6: A pregnant woman at her first trimester is planning to visit Nigeria she consulted you regarding safe prophylaxis against malaria. Which one of the following would be the best management in this patient ?**

- A. Chloroquine daily till she delivers
- B. Mefloquine weekly
- C. Quinine daily for 4 weeks
- D. Primaquine daily

# 438 Lecture Quiz

**Q1: The region with the greatest morbidity and mortality from malaria in the world is?**

- A. Africa.
- B. Southeast Asia.
- C. South America.
- D. Oceania.

**Q2: A young American adult consults with you before travel to Kenya. Appropriate regimens for the prevention of malaria in travelers from a developed country to areas with chloroquine-resistant *P. falciparum* malaria include**

- A. Malarone, mefloquine, or artemisinin.
- B. Malarone, mefloquine, or doxycycline.
- C. Coartem, Malarone, or doxycycline.
- D. quinine, artemisinin, or Fansidar.

**Q3: A child from Ghana is admitted with fever, altered consciousness, acute renal failure, and 7% parasitemia with *P. falciparum*. Severe malaria should be treated with:**

- A. Oral artemisinin-based combination therapy
- B. Malarone or mefloquine
- C. Quinine or quinidine
- D. Intravenous artesunate, if available, and as a back-up intravenous quinine or quinidine

**Q4: Reasons for the predilection for *P.falciparum* to cause severe malaria include all of the following except**

- A. cytoadherence of *P. falciparum*-infected erythrocytes to vascular endothelium.
- B. infection of erythrocytes of all ages.
- C. a high prevalence of resistance to available antimalarial drugs.
- D. increasing pathogenicity with increasing age of the patient.

**Q5: A 55-year-old woman presented in May with fever and headache of 1 day's duration. She returned 7 days earlier from a 3-week trip to game parks in Tanzania, East Africa. On examination, her temperature is 40° C, blood pressure is 120/85 mm Hg, heart rate is 120 beats per minute, and respiratory rate is 18 breaths per minute. She has no rash, her neck is supple, chest is clear to auscultation and percussion, and abdomen is soft and nontender with normal bowel sounds. A laboratory technician on call has never done a malaria smear, and the hospital does not offer rapid diagnostic tests. Results from a reference laboratory will be available in 24 hours. Which of the following would be the best course of action?**

- A. Treat with mefloquine
- B. Treat if parasites are identified at the reference laboratory
- C. Treat with chloroquine
- D. Treat with artemether/lumefantrine (Coartem)

**Q6: The region with the most highly resistant malaria parasites in the world is:**

- A. Africa
- B. Southeast Asia
- C. South America
- D. Oceania

# GOOD LUCK!

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