Malaria 1 Clinical Tropical Medicine FACTM (Clinical) Pt 1

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Malaria series

Malaria 1 Clinical Tropical MedicineMalaria 2 Clinical ParasitologyMalaria 3 Public Health, Travel & Expedition MedicineMalaria 4 Clinical Entomology



Study materials

Oxford Handbook of Tropical Medicine. 3rd Edition, Eddlestone M *et al.* 2008. OUP.

- Recommended bookshelf
- FACTM study notes
- Specialist review articles
- Self-assessment questions
- Blog <u>http://micrognome.priobe.net</u>
- Web <u>www.priobe.net</u>

FACTM <u>http://lifeinthefastlane.com/exams/actm-fellowship/</u>



Clinical features

- Classical presentation:
 - COLD initial shaking/rigor; then
 - **HOT** fever (may be >40°C), restlessness, vomiting & convulsions; then final
 - **SWEATING** temperature returning to normal & possibly sleep.
- **Setting**: history of travel to or residence in malaria-endemic area
- **Prodrome**: aching, lethargy
- **Timing**: 6-10hr overall with interval of 38-42hr for *P.vivax* or *P.ovale* and 62-66hr for *P. malariae*. *P.falciparum* timing is less predictable, temperature may not return to normal between paroxysms
- **Exceptions**: less clear cut in children
- **Misleading features**: cough, headache, myalgia, diarrhoea, jaundice may all be present in acute malaria



Severe malaria

WHO criteria:

- Clinical. prostration, impaired consciousness, respiratory distress, multiple convulsions, circulatory collapse, pulmonary oedema, abnormal bleeding, jaundice, haemoglobinuria
- Laboratory. severe anaemia, hypoglycaemia, acidosis, renal impairment, hyperlactataemia, hyperparasitaemia
- **Blackwater fever**: massive haemoglobinuria in malaria. After use of quinine or primaquine. Commoner in patients with G6PD deficiency
- Cerebral malaria: "unrousable coma in the presence of peripheral parasitaemia when other causes of encephalopathy have been excluded" 20% mortality. Children and non-immune adults. Kernig's NEG, neck rigidity & photophobia usually not present.



Severe malaria ii

- **Respiratory distress**: due to compensation for metabolic acidosis, pulmonary capillary damage by parasite, 2° pneumonia, severe anaemia
- Severe anaemia: haematocrit <15% in presence of parasitaemia. pallor, gallop rhythm, pulmonary oedema, neuro signs
- **Jaundice**: signs of liver failure uncommon unless also has hepatitis
- **Renal impairment**: raised Cr and urea. Oliguric, anuric, occasionally polyuria. Acute failure in malaria has poor prognosis, approx 45% mortality.
- Hypoglycaemia: blood glucose < 2.2 mmol/L. commoner in pregnancy, after quinine or due to liver impairment. clinical features easy to miss if reduced conscious level.



Investigations

Key questions:

- Does the patient have malaria?
- Does the patient have *P. falciparum* malaria?
- Does the patient have another infection?
- Blood films: at least 2, preferably 3 at intervals by 2 methods (thin & thick), for parasite detection, density, determination of species & stages present
- **Rapid tests**: dipstick for *P. falciparum* histidine-rich protein, quantitative buffy coat, and PCR assays; mainly for *P. falciparum* infection, not useful for parasite density
- Other infections: blood culture, arbovirus serology, PCR assays
- Blood glucose, U&Es, liver function tests, FBC
- Others, as indicated by severity of infection



Antimalarial treatment

WHO guidelines:

- Artemisinin-based combination therapy (ACT) for uncomplicated malaria
- Artesunate for parenteral treatment in low transmission area & later pregnancy

General rules:

- If signs of *P.falciparum* malaria, weigh patient & start immediately
- Avoid discharging patients with mild symptoms but high parasitaemia (≥ 100,000 parasites/µL or ≥ 2% RBC infected)
- If benign malaria, await results of blood film
- Uncomplicated malaria can be treated on outpatient basis
- If outpatient treatment, advise return if worsens or no improvement in 48hr
- For Chemoprophylaxis, see Malaria 3.



Antimalarial agents

- **ACTs**: e.g. artemether 20mg/lumefantrine 120mg fixed combination
 - Rapid effect against schizont stage of *P.falciparum* infection
 - 6 doses in 3d, orally. Taken with milk or fatty food
- Artesunate 2.4mg/kg IV
 - In severe malaria, 3 doses in 24hr, then once daily
 - More effective than quinine
- Chloroquine: 25mg/kg base in divided doses over 3d
 - For benign malaria only
 - Primaquine needed for liver schizont stage of *P.vivax* & *P.ovale* (beware G6PD def.)
- Quinine: 10mg/kg salt 8hourly
 - Tolerated poorly due to cinchonism, requires additional tetracycline
 - Risk of hypoglycaemia, prolonged QT interval
 - Use in relapse within 14d of ACT
 - Loading dose of 20mg/kg, especially in severe or complicated malaria



Managing the patient

- <u>All</u> need antimalarial chemotherapy
- ABCs, including venous access
- Deal with hypoglycaemia
- Weigh patient, assess hydration
- If diminished conscious level, consider LP
- If convulsions, use Diazepam by slow IV
 - GCS, mannitol etc have no clear benefit in cerebral malaria
- Monitor urine output & renal function
- If severe anaemia (Hct <15%), consider transfusing
- If shocked, consider possibility of bacterial infection, give IV antibacterial



Emerging issues

- 1. Choice of antimalarial therapy after failed chemoprophylaxis
- 2. Potential for resistance to new artemisinin agents
- 3. Range of new fixed combination ACTs
- 4. The role of antipyretics e.g. in children with malaria
- 5. The role of exchange transfusion in severe malaria
- 6. Treatment of parasitaemia in refugee clinics
- 7. Treatment of malaria in pregnancy
- 8. Emergency treatment of malaria in remote places

