

Table S2. Treatment for each severe falciparum malaria sub-group group over time at the Shoklo Malaria Research Unit

| Start year | Treatment for pregnant women | |
|--|--|---|
| First-line initial treatment for severe malaria with organ dysfunction | | |
| 1989 | Q IV: 20mg/kg (H0) over 4 hours followed by 10mg/kg over 2 hours every 8 hours | |
| 1996 | Q IV: 20mg/kg (H0) over 4 hours followed by 10mg/kg over 2 hours every 8 hours Or AM IM: 3.2 mg/kg (H0) followed by 1.6 mg/kg every 24 hours (or 8-12 hourly in 1996–7) | |
| 2000 | AS IV + (from 2006) clindamycin p.o. H0: 2.4 mg, H12 and H24 1.2mg/kg and then 1.2 mg/kg every 24 hours Or AM IM or Q IV if AS IV is unavailable | |
| 2013 | AS IV + clindamycin p.o. 2.4mg/kg at H0, H12, H24 and then every 24 hours Suspected AS resistance or severe malaria with hyperparasitaemia ($\geq 4\%$ RBC) Q IV + AS IV | |
| First-line for severe malaria with hyperparasitaemia (>10% RBC) only (without other organ dysfunction) | | |
| 1986 | Q IV: 20mg/kg (H0) over 4 hours followed by 10mg/kg over 2 hours every 8 hours | |
| 1994 | AS p.o. 4mg/kg (D0), 2mg/kg (D1–2) and 1mg/kg (D3–6) (total dose 12 mg/kg)† | |
| 1997 | AS p.o. 4mg/kg (D0), 2mg/kg (D1–4) and 1mg/kg (D5–6) (total dose 16mg/kg) † | |
| 1999 | AS p.o. 4mg/kg (D0), 2mg/kg (D1–6) (total dose 16mg/kg) † | |
| 2012 | AS p.o. 4mg/kg (D0), 2mg/kg (D1–6) with DP p.o. 3 days (D4–6)* | |
| 2013 | AS p.o. 4 mg/kg (D0–2) followed by DP p.o. 5 days (D2–6)* | |
| 2016 | Any ACT extended regimen (for 7 days)* | |
| First-line for severe malaria with severe anaemia only (without other organ dysfunction) | | |
| 1989 | All trimesters | Q7 or or MFQ p.o. (25mg/kg single dose) |
| 1994 | 1st trimester | Q7 |
| | 2nd & 3rd trimester | 1st episode: Q7 or MFQ p.o. (25mg/kg single dose) ≥ 2 nd episode: AS p.o. (2mg/kg D0–D4, 1mg/kg D5–6) |
| 1996 | 1st trimester | Q7 |
| | 2nd & 3rd trimester | AS p.o. (2mg/kg D0–D4, 1mg/kg D5–6) or MAS |
| 2006 | 1st trimester | Q p.o. + clindamycin for 7 days |
| | 2nd & 3rd trimester | AS p.o. + clindamycin for 7 days |
| 2017 | All trimesters | Available ACT |

AS: artesunate. AC: artesunate+clindamycin. D: day. DP: dihydroartemisinin+piperazine. H: hour. IM: intramuscular. IV: intravenous. MAS:mefloquine+artesunate. MFQ: mefloquine. p.o.: per os. Q: quinine. RBC: red blood cells.

Clindamycin was given orally 5mg/kg three times a day for 7 days.

MAS: mefloquine 25 mg/kg in total + artesunate 4mg/kg for three days

Q7: Q p.o.: 10mg/kg three times a day for 7 days.

A rescue dose (1.2 mg/kg) was given if parasitaemia increased above the 95th percentile of the usual parasite clearance curve [19].

‡ Treatment was switched to oral drug once tolerated. Total 12–16 mg/kg (-2012) or 16–18 mg/kg (2013-) of AS or AM was given over 7 days.

† A partner drug (e.g. MFQ, or atovaquone-proguanil if available) was used whenever possible.

*First dose was given by IV or IM (2.4mg/kg) to those with asexual parasitaemia of >20% RBC.