

## List L1. List of included studies

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| 1 | Kaneko. A. An uncontrolled before and after study to examine the effect of a short-term mass drug administration (MDA) on malaria parasite prevalence on Ngodhe Island, Homa Bay County, Kenya [unpublished]   |
| 2 | Okebe J, Bousema T, Affara M, Di Tanna GL, Dabira E, Gaye A, Sanya-Isijola F, Badji H, Correa S, Nwakanma D, Van Geertruyden JP, Drakeley C, D'Alessandro U. The Gametocytocidal Efficacy of Different Single Doses of Primaquine with Dihydroartemisinin-piperaquine in Asymptomatic Parasite Carriers in The Gambia: A Randomized Controlled Trial. <i>EBioMedicine</i> . 2016 Nov;13:348-355.                                       |
| 3 | Mwaiswelo R, Ngasala BE, Jovel I, Gosling R, Premji Z, Poirot E, Mmbando BP, Björkman A, Mårtensson A. Safety of a single low-dose of primaquine in addition to standard artemether-lumefantrine regimen for treatment of acute uncomplicated Plasmodium falciparum malaria in Tanzania. <i>Malar J</i> . 2016 Jun 10;15:316.  |
| 4 | Hamid MMA, Thriemer K, Elobied ME, Mahgoub NS, Boshara SA, Elsafi HMH, Gumaa SA, Hamid T, Abdelbagi H, Basheir HM, Marfurt J, Chen I, Gosling R, Price RN, Ley B. Low risk of recurrence following artesunate-Sulphadoxine-pyrimethamine plus primaquine for uncomplicated Plasmodium falciparum and Plasmodium vivax infections in the Republic of the Sudan. <i>Malar J</i> . 2018 Mar 16;17(1):117.                                 |
| 5 | Gonçalves BP, Tiono AB, Ouédraogo A, Guelbéogo WM, Bradley J, Nebie I, Siaka D, Lanke K, Eziefula AC, Diarra A, Pett H, Bougouma EC, Sirima SB, Drakeley C, Bousema T. Single low dose primaquine to reduce gametocyte carriage and Plasmodium falciparum transmission after artemether-lumefantrine in children with asymptomatic infection: a randomised, double-blind, placebo-controlled trial. <i>BMC Med</i> . 2016 Mar 8;14:40. |
| 6 | El-Sayed B, El-Zaki SE, Babiker H, Gadalla N, Ageep T, Mansour F, Baraka O, Milligan P, Babiker A. A randomized open-label trial of artesunate- sulfadoxine-pyrimethamine with or without primaquine for elimination of sub-microscopic P. falciparum parasitaemia and gametocyte carriage in eastern Sudan. <i>PLoS One</i> . 2007 Dec 12;2(12):e1311   |
| 7 | Samuels A, Kwambai T, Desai M, Kariuki S, Oneko M, Ochomo E, Otieno K, Smit M, Ter Kuile F, Gimnig J, ping Shi Y, Gosling R, Hwang J. Determining a safe and maximally efficacious dosing range of primaquine in combination with standard weight-based dosing of Dihydroartemisinin-piperaquine against Plasmodium falciparum gametocytes in G6PD-normal and -deficient patients with uncomplicated malaria. unpublished              |
| 8 | Eziefula AC, Bousema T, Yeung S, Kamya M, Owaraganise A, Gabagaya G, Bradley J, Grignard L, Lanke KH, Wanzira H, Mpimbaza A, Nsoya S, White NJ, Webb EL, Staedke SG, Drakeley C. Single dose primaquine for clearance of Plasmodium falciparum gametocytes in children with uncomplicated malaria in Uganda: a randomised, controlled, double-blind, dose-ranging trial. <i>Lancet Infect Dis</i> . 2014 Feb;14(2):130-9.              |

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| 9&10 | Bastiaens GJH, Tiono AB, Okebe J, Pett HE, Coulibaly SA, Gonçalves BP, Affara M, Ouédraogo A, Bougouma EC, Sanou GS, Nébié I, Bradley J, Lanke KHW, Niemi M, Sirima SB, d'Alessandro U, Bousema T, Drakeley C. Safety of single low-dose primaquine in glucose-6-phosphate dehydrogenase deficient falciparum-infected African males: Two open-label, randomized, safety trials. PLoS One. 2018 Jan 11;13(1):e0190272.  |
| 11   | Tine RC, Sylla K, Faye BT, Poirot E, Fall FB, Sow D, Wang D, Ndiaye M, Ndiaye JL, Faye B, Greenwood B, Gaye O, Milligan P. Safety and efficacy of adding a single low dose of primaquine to the treatment of adult patients with Plasmodium falciparum malaria in Senegal, to reduce gametocyte carriage: a randomized controlled trial. Clin Infect Dis. 2017 Aug 15;65(4):535-543.  |
| 12   | Poirot E, Soble A, Ntshalintshali N, Mwandemele A, Mkhonta N, Malambe C, Vilakati S, Pan S, Darteh S, Maphalala G, Brown J, Hwang J, Pace C, Stergachis A, Vittinghoff E, Kunene S, Gosling R. Development of a pharmacovigilance safety monitoring tool for the rollout of single low-dose primaquine and artemether-lumefantrine to treat Plasmodium falciparum infections in Swaziland: a pilot study. Malar J. 2016 Jul 22;15(1):384.                         |
| 13   | Shekalaghe S, Drakeley C, Gosling R, Ndaro A, van Meegeren M, et al (2007) Primaquine Clears Submicroscopic Plasmodium falciparum Gametocytes that Persist after Treatment with Sulphadoxine-Pyrimethamine and Artesunate. PLoS ONE 2(10): e1023.   |
| 14   | Shekalaghe SA, ter Braak R, Daou M, Kavishe R, van den Bijllaardt W, van den Bosch S, Koenderink JB, Luty AJ, Whitty CJ, Drakeley C, Sauerwein RW, Bousema T. In Tanzania, hemolysis after a single dose of primaquine coadministered with an artemisinin is not restricted to glucose-6-phosphate dehydrogenase-deficient (G6PD A-) individuals. Antimicrob Agents Chemother. 2010 May;54(5):1762-8.   |
| 15   | Sutanto I, Suprijanto S, Kosasih A, Dahlan MS, Syafruddin D, Kusriastuti R, Hawley WA, Lobo NF, Ter Kuile FO. The effect of primaquine on gametocyte development and clearance in the treatment of uncomplicated falciparum malaria with dihydroartemisinin-piperaquine in South Sumatra, Western Indonesia: an open-label, randomized, controlled trial. Clin Infect Dis. 2013 Mar;56(5):685-93.   |
| 16   | Raman J, Allen E, Workman L, Mabuza A, Swanepoel H, Malatje G, Frean J, Wiesner L, Barnes KI. Safety and tolerability of single low-dose primaquine in a low-intensity transmission area in South Africa: an open-label, randomized controlled trial. Malar J. 2019 Jun 24;18(1):209  |
| 17   | Stone W, Sawa P, Lanke K, Rijpma S, Oriango R, Nyaurah M, Osodo P, Osoti V, Mahamar A, Diawara H, Woestenenk R, Graumans W, van de Vegte-Bolmer M, Bradley J, Chen I, Brown J, Siciliano G, Alano P, Gosling R, Dicko A, Drakeley C, Bousema T. A Molecular Assay to Quantify Male and Female Plasmodium falciparum Gametocytes: Results From 2 Randomized Controlled Trials Using Primaquine for Gametocyte Clearance. J Infect Dis. 2017 Aug 15;216(4):457-467. |
| 18   | Ley B, Alam MS, Thriemer K, Hossain MS, Kibria MG, Auburn S, Poirot E, Price RN, Khan WA. G6PD Deficiency and antimalarial efficacy for uncomplicated malaria in Bangladesh: a prospective observational study. PLoS One. 2016 Apr 29;11(4):e0154015.   |

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| 19 | Dicko A, Brown JM, Diawara H, Baber I, Mahamar A, Soumare HM, Sanogo K, Koita F, Keita S, Traore SF, Chen I, Poirot E, Hwang J, McCulloch C, Lanke K, Pett H, Niemi M, Nosten F, Bousema T, Gosling R. Primaquine to reduce transmission of Plasmodium falciparum malaria in Mali: a single-blind, dose-ranging, adaptive randomised phase 2 trial. Lancet Infect Dis. 2016 Jun;16(6):674-684. doi: 10.1016/S1473-3099(15)00479-X. Epub 2016 Feb 20 |
| 20 | Bancone G, Chowwwiat N, Somsakchaicharoen R, Poodpanya L, Moo PK, Gornsawun G, Kajeechiwa L, Thwin MM, Rakthinthong S, Nosten S, Thinraow S, Nyo SN, Ling CL, Wiladphaingern J, Kiricharoen NL, Moore KA, White NJ, Nosten F. Single low dose primaquine (0.25 mg/kg) does not cause clinically significant haemolysis in G6PD-deficient subjects. PLoS One. 2016 Mar 24;11(3):e0151898.  |

Supplementary Table 1. Summary of characteristics of included studies

| Study ID | PubMed ID   | Study Design | Location     | Year      | ACT        | FU days | Population at baseline |                      |               |                       |                         |                            |
|----------|-------------|--------------|--------------|-----------|------------|---------|------------------------|----------------------|---------------|-----------------------|-------------------------|----------------------------|
|          |             |              |              |           |            |         | N                      | Malaria              | Age           | Pregnant              | G6PD normal / deficient | Inclusion Hb cutoff (g/dL) |
| 1        | Unpublished | MDA          | Kenya        | 2016      | AP*        | 7       | 291                    | Either               | All           | Include               | Either                  | ≥ 7                        |
| 2        | 27825738    | RCT          | The Gambia   | 2013-2015 | DP         | 42      | 689                    | Pf asymptomatic      | > 1 year      | Exclude               | Normal                  | ≥ 8                        |
| 3        | 27287612    | RCT          | Tanzania     | 2014      | AL         | 28      | 220                    | Pf uncomplicated     | ≥ 1 year      | Exclude               | Either                  | ≥ 8                        |
| 4        | 29548285    | RCT          | Sudan        | 2015      | ASSP       | 42      | 231                    | Pf uncomplicated     | ≥ 1 year      | Exclude               | Either**                | ≥ 8                        |
| 5        | 26952094    | RCT          | Burkina Faso | 2013-2014 | AL         | 14      | 360                    | Pf asymptomatic      | 2 - 15 years  | Not tested            | Normal                  | ≥ 8                        |
| 6        | 18074034    | RCT          | Sudan        | 2004      | ASSP       | 14      | 100                    | Pf asymptomatic      | ≥ 6 months    | Exclude               | Not tested              | ≥ 8                        |
| 7        | Unpublished | RCT          | Kenya        | 2014-2015 | DP         | 42      | 54                     | Pf uncomplicated     | 1 ≤ 12 years  | Not tested            | Both**                  | ≥ 8                        |
| 8        | 24239324    | RCT          | Uganda       | 2011      | AL         | 28      | 454                    | Pf uncomplicated     | 1 - 10 years  | Not tested            | Normal                  | ≥ 8                        |
| 9        | 29324864    | RCT & cohort | Burkina Faso | 2014-2015 | AL         | 28      | 78                     | Pf asymptomatic      | 18 - 45 years | Males only            | Normal                  | ≥ 11                       |
| 10       | 29324864    | RCT          | The Gambia   | 2015-2016 | DP         | 28      | 61                     | Pf asymptomatic      | ≥ 10 years    | Males only            | Either                  | ≥ 11                       |
| 11       | 28605472    | RCT          | Senegal      | 2014-2016 | AL ASAQ DP | 28      | 267                    | Pf uncomplicated     | > 18 years    | Exclude               | Either                  | ≥ 8                        |
| 12       | 27450652    | Cohort       | eSwatini     | 2014-2015 | AL         | 7       | 94                     | Pf uncomplicated     | > 1 year      | Exclude               | Either                  | > 8                        |
| 13       | 17925871    | RCT          | Tanzania     | 2006      | ASSP       | 42      | 107                    | Pf uncomplicated     | 3 - 15 years  | Not tested            | Either                  | > 8                        |
| 14       | 20194698    | MDA          | Tanzania     | 2008      | ASSP       | 7       | 840                    | Not tested           | 1 - 12 years  | Exclude               | Either                  | > 8                        |
| 15       | 23175563    | RCT          | Indonesia    | 2008-2010 | DP         | 42      | 373                    | Pf uncomplicated     | ≥ 5 years     | Exclude               | Normal                  | ≥ 8                        |
| 16       | 31234865    | RCT          | South Africa | 2016-2018 | AL         | 42      | 140                    | Pf uncomplicated     | > 1 year      | Exclude               | Either                  | ≥ 7                        |
| 17       | 28931236    | RCT          | Kenya        | 2014-2015 | DP         | 14      | 114                    | Pf uncomplicated *** | 5 - 15 years  | Not tested            | Not tested              | > 9.5                      |
| 18       | 27128675    | Cohort       | Bangladesh   | 2014-2015 | AL         | 28      | 115                    | Pf uncomplicated     | > 1 year      | Males only            | Either                  | ≥ 8                        |
| 19       | 26906747    | RCT          | Mali         | 2013-2014 | DP         | 28      | 81                     | Pf uncomplicated *** | 5 - 50 years  | Males only            | Normal                  | ≥ 8                        |
| 20       | 27010542    | MDA          | Thailand     | 2014      | DP         | 7       | 1737                   | Not tested           | > 6 mths      | Exclude 1st trimester | Either                  | none                       |

ACT = artemisinin-based combination therapy, AE = adverse event, AL = artemether-lumefantrine, ASAQ = artesunate-amodiaquine, ASSP = artesunate sulfadoxine-pyrimethamine, DP = dihydroartemisinin piperaquine, FST = fluorescent spot test, Hb = haemoglobin, MDA = mass drug administration, PCR = polymerase chain reaction, PQ = primaquine, RCT = randomised controlled trial \*piperaquine if pregnant \*\*normal if vivax \*\*\*gametocyte carriers

Supplementary Table 1. Summary of characteristics of included studies (continued)

| Study ID | Methods  |                          | PQ treatment |       |     |                          | Included in IPD meta-analysis |    |                 |
|----------|--|--------------------------|--------------|-------|-----|--------------------------|-------------------------------|----|-----------------|
|          | G6PD testing method used for IPD meta-analysis | Malaria diagnosis method | %            | N     | Day | Target dose              | Hb                            | AE | Haemoglobinuria |
| 1        | Dojindo WST                                    | Microscopy               | 98           | 286   | 0   | 0.144, 0.208             | Y                             |    |                 |
| 2        | FST (Dimopolous)                               | RDT                      | 75           | 515   | 2   | 0.2, 0.4, 0.75           | Y                             | Y  | Y               |
| 3        | CareStart RDT (Access Bio)                     | Microscopy               | 50           | 110   | 0   | 0.25                     | Y                             | Y  | Y               |
| 4        | Carestart RDT (Access Bio)                     | Microscopy               | 52           | 119   | 2   | 0.25                     | Y                             |    |                 |
| 5        | FST  | Microscopy               | 69           | 247   | 2   | 0.25, 0.4                | Y                             | Y  | y               |
| 6        | Not applicable                                 | Microscopy               | 49           | 49    | 3   | 0.75                     | Y                             |    |                 |
| 7        | FST (Trinity BioTech)                          | Microscopy               | 78           | 42    | 0   | 0.125, 0.25, 0.4, 0.75   | Y                             | Y  | y               |
| 8        | FST (R&D Diagnostics)                          | Microscopy               | 75           | 339   | 2   | 0.1 0.4 0.75             | Y                             |    | y               |
| 9        | CareStart RDT (Access Bio)                     | Microscopy               | 87           | 68    | 0   | 0.25, 0.4                | Y                             | Y  | y               |
| 10       | CareStart RDT (Access Bio)                     | Microscopy               | 82           | 50    | 0   | 0.25, 0.4                | Y                             | Y  |                 |
| 11       | CareStart RDT (Access Bio)                     | Microscopy               | 50           | 133   | 0   | 0.25                     | Y                             |    | y               |
| 12       | CareStart RDT (Access Bio)                     | Not known                | 100          | 94    | 0   | 0.25                     | Y                             | Y  |                 |
| 13       | PCR  | Microscopy               | 50           | 54    | 2   | 0.75                     | Y                             |    |                 |
| 14       | PCR  | Not known                | 93           | 784   | 2   | 0.75                     | Y                             |    |                 |
| 15       | FST (Trinity Biotech)                          | Microscopy               | 52           | 194   | 3   | 0.75                     | Y                             |    |                 |
| 16       | CareStart RDT (Access Bio)                     | RDT, microscopy          | 50           | 70    | 3   | 0.25                     | Y                             | Y  |                 |
| 17       | Not applicable                                 | Microscopy               | 51           | 58    | 2   | 0.25                     | Y                             |    |                 |
| 18       | FST (Randox UK)                                | Microscopy               | 100          | 115   | 2   | 0.75                     |                               | Y  |                 |
| 19       | Colorimetric quantification (R&D Diagnostics)  | Microscopy               | 80           | 65    | 0   | 0.0625, 0.125, 0.25, 0.5 | Y                             | Y  | y               |
| 20       | FST (R&D Diagnostics)                          | Not known                | 100          | 1,737 | 1   | 0.25                     |                               | Y  |                 |

ACT = artemisinin-based combination therapy, AE = adverse event, AL = artemether-lumefantrine, ASAQ = artesunate-amodiaquine, ASSP = artesunate sulfadoxine-pyrimethamine, DP = dihydroartemisinin piperaquine, FST = fluorescent spot test, Hb = haemoglobin, MDA = mass drug administration, PCR = polymerase chain reaction  
PQ = primaquine, RCT = randomised controlled trial \*piperaquine if pregnant \*\*normal if vivax \*\*\*gametocyte carriers