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Pharmacokinetics and efficacy of dihydroartemisinin-piperaquine in the treatment of uncomplicated falciparum malaria in children in Burkina Faso

Submission date 11/09/2007	**Recruitment status** No longer recruiting	[_] Prospectively registered [_] Protocol [_] Statistical analysis plan [X] Results [_] Individual participant data
Registration date 05/02/2008	**Overall study status** Completed	
Last Edited 13/05/2015	**Condition category** Infections and Infestations	

Plain English summary of protocol
Not provided at time of registration

Contact information

Type(s)
Scientific

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Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers
N/A

Study information

Scientific Title
Assessing the efficacy of Dihydroartemisinin-piperaquine in African patients suffering of uncomplicated falciparum malaria and to determine the pharmacokinetics profile of piperaquine in children 2 - 10 years old presenting falciparum malaria

Study objectives
Preliminary results of pharmacokinetic (PK) studies indicate that the disposition of piperaquine is altered in children compared to adults.

Ethics approval required
Old ethics approval format

Ethics approval(s)
1. Institut de Recherche en Science de la Sante/Centre Muraz (IRSS/CM) (Burkina Faso) , 26/07 /2007, ref: 005-2007/CE-CM
2. University of California, San Francisco (USCF) committee on Human Research, 27/07/2007, ref: # H40380-31179-01

Study design
1. Treatment efficacy: open-label trial
2. Population kinetic studies will use sparse capillary sampling

Primary study design
Interventional

Secondary study design
Randomised controlled trial

Study setting(s)
Hospital

Study type(s)
Treatment

Participant information sheet

Health condition(s) or problem(s) studied
Malaria

Interventions
Patients are given dihydroartemisin piperaquine once daily for three days. Treatment is weight based and directly observed by the study nurse. The follow up duration is 42 days.

The study is a one arm study but there is a randomisation to determine the groups where the patient will be included for the PK purpose.

Intervention Type
Drug

Phase
Not Applicable

Drug/device/biological/vaccine name(s)
Dihydroartemisinin-piperaquine

Primary outcome measure
1. Determination of the pharmacokinetic profile of piperaquine in children with uncomplicated falciparum malaria
2. Assess the efficacy of dihydroartemisinin piperaquine

Secondary outcome measures
1. Risk of recurrent malaria*
2. Risk of clinical and parasitological treatment failure*
3. Prevalence of fever (defined as both subjective fever in the previous 24 hours and measured axillary temperature greater than 37.5°C) on follow-up days 1, 2, and 3
4. Prevalence of parasitaemia on follow-up days 2 and 3
5. Change in mean haemoglobin from day 0 to 42 (or day of rescue therapy for patients classified as late clinical failure [LCF] or late parasitological failure [LPF])
6. Prevalence of gametocytaemia on follow-up days 2, 3, 7, 14, 21 and 28
7. Change in the prevalence of molecular markers possibly associated with drug resistance on day 0 or the day of recurrent parasitaemia, including polymorphisms in Plasmodium falciparum chloroquine resistance transporter (Pfcrt) and Plasmodium falciparum multidrug-resistance (Pfmdr1) genes
8. In vitro sensitivity to antimalarial drugs

*Risks will be estimated using the Kaplan-Meier product limit formula based on a modified intention-to-treat analysis.

Overall study start date
06/08/2007

Completion date
31/01/2008

Eligibility

Key inclusion criteria
On day 0, patients with symptoms suggestive of malaria and a positive screening thick blood smear will be assessed for the following selection criteria by study physicians for appropriate care:
1. Not previously enrolled in this study
2. Aged greater than 6 months
3. Weight greater than 5 kg
4. Fever (greater than 37.5°C axillary) or history of fever in the previous 24 hours
5. Absence of any history of serious side effects to study medications
6. No evidence of a concomitant febrile illness in addition to malaria

7. Provision of informed consent and ability to participate in 42-day follow-up (patient has easy access to health unit)
8. No history of antimalarial use in the previous two weeks (except for chloroquine)
9. No danger signs or evidence of severe malaria defined as:
9.1. Unarousable coma (if after convulsion, greater than 30 minutes)
9.2. Recent febrile convulsions (within 24 hours)
9.3. Altered consciousness (confusion, delirium, psychosis, coma)
9.4. Lethargy
9.5. Unable to drink or breast feed
9.6. Vomiting everything
9.7. Unable to stand/sit due to weakness
9.8. Severe anaemia (haemoglobin [Hb] less than 5.0 gm/dL)
9.9. Respiratory distress (laboured breathing at rest)
9.10. Jaundice

After going to the laboratory, the subjects will be referred to the study nurse for treatment allocation and treatment with the study medications. Patients must also meet the following criterion:
10. Absence of repeated vomiting of study medications on day 0

Patients will return to the clinic on day 1 and will be excluded from the study if the following inclusion criteria are not met:
11. Plasmodium falciparum mono-infection
12. Parasite density 2000 - 200,000/ul

Participant type(s)
Patient

Age group
Child

Lower age limit
6 Months

Sex
Both

Target number of participants
330

Key exclusion criteria
1. Inhability to participate in 42 days follow up
2. Pregnant women
3. Severe malaria

Date of first enrolment
06/08/2007

Date of final enrolment
31/01/2008

Locations

Countries of recruitment
Burkina Faso

Thailand

Study participating centre
Shoklo Malaria Research Unit (SMRU)
Mae Sot
Thailand
63110

Sponsor information

Organisation
Beijing Holley-Cotec Pharmaceuticals Co. Ltd (China)

Sponsor details
Room 1602, Full Tower
No. 9, Dong San Huan Zhong Road
Chaoyang District
Beijing
China
100020

Sponsor type
Industry

Website
http://www.holleycotec.com

Funder(s)

Funder type
Charity

Funder Name
Doris Duke Charitable Foundation (USA)

Alternative Name(s)
Doris Duke Charitable Foundation, Inc., DDCF Trust, Doris Duke Foundation, DDCF

Funding Body Type
Private sector organisation

Funding Body Subtype
Trusts, charities, foundations (both public and private)

Location
United States of America

Funder Name
Beijing Holley-Cotec Pharmaceuticals Co. Ltd (China)

Funder Name
National Budget of Institut de Recherche en Science de la Sante (IRSS) (Burkina Faso)

Results and Publications

Publication and dissemination plan
Not provided at time of registration

Intention to publish date

Individual participant data (IPD) sharing plan

IPD sharing plan summary
Not provided at time of registration

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Results article	results	18/08/2014		Yes	No