

# Outcomes of asymptomatic malaria parasitaemia in neonates in a tertiary hospital, southeast Nigeria

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## ABSTRACT

**Background:** Malaria infestation during pregnancy is mostly asymptomatic and untreated especially in unbooked pregnancies. It presents with almost all the fetal complications of overt malaria in pregnancy. The aim of this study was to determine the effect of asymptomatic malaria parasitaemia on the neonates of unbooked parturients delivered at term at the Federal Teaching Hospital, Abakaliki. **Materials and Methods:** This study was conducted in the labour ward complex of the Federal Teaching Hospital, Abakaliki from March to May 2012. Unbooked pregnant women who fulfilled the inclusion criteria and gave consent were consecutively recruited. Cord blood and placenta tissue were collected for haemoglobin concentration determination and histology, respectively. Birth weights were determined with an electronic weighing machine. Statistical Analysis was done with 2008 Epi Info™ software and level of significant was set at  $P$ -value  $< 0.05$ . **Results:** A total of 250 unbooked parturients were recruited, of which 194 (77.6%) had asymptomatic malaria parasitaemia while 227 (90.8%) had placental parasitisation. The prevalence of low birth weight in the study was 16.4%. There was significant relationship between asymptomatic malaria parasitemia and birth weight ( $X^2 = 43.70$ ,  $P$ -value  $< 0.001$ ). There were no low-birth-weight deliveries among parturients without placental parasitemia. No neonate, however, had anaemia in the study. **Conclusion:** Asymptomatic malaria parasitemia and placental parasitisation by malaria parasites contribute to the outcome of the foetal birth weight. Asymptomatic malaria parasitaemia and placental parasitaemia did not result in a corresponding foetal anaemia on babies delivered.

**Key words:** Asymptomatic malaria parasitaemia, labour, neonatal outcome, Nigeria, pregnant women

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## INTRODUCTION

Malaria in pregnancy is an important cause of anaemia, miscarriages, intrauterine growth restriction, low birth weight, still birth and other pregnancy-related complications.<sup>1</sup> Several studies have reported that 125 million women worldwide and 25-50 million African women who get pregnant annually are at risk of malaria.<sup>2-5</sup> Ninety percent of global malaria burden occur in Africa south of Sahara where malaria in pregnancy has been most evaluated.<sup>6</sup> Seventy percent of pregnant women in Nigeria suffer malaria with maternal and foetal complications.<sup>1</sup>

Most pregnant women with malaria infestation are asymptomatic thus are undetected and untreated.<sup>7</sup>

Foetal complications of malaria in pregnancy include early pregnancy loss, intrauterine growth restriction, vertical transmission of malaria, intrauterine death, low birth weight and congenital malaria. Most of the foetal complications are attributed to placental malaria and reports have shown that the human placenta is an ideal site for the accumulation of *Plasmodium falciparum* and as a consequence, serious health problems arise for the mother and her baby.<sup>8</sup> Studies have shown that placental sequestration of *Plasmodium falciparum* result in accumulation of parasitised erythrocytes in the intervillous space, infiltration by inflammatory cells and release of pro-inflammatory mediators with malaria pigment and perivillous fibrinoid deposits, proliferation of cytotrophoblastic cells and thickening of basement membrane which impair materno- foetal exchange, often resulting in adverse pregnancy outcome.<sup>9-11</sup> There is a postulation that the biological basis for susceptibility to

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malaria in pregnancy is due to accumulation of erythrocytes infected by *Plasmodium falciparum* in the placenta through adhesion to molecules such as chondroitin sulphate A.<sup>12</sup> Studies have also shown that placental malaria induce pathological lesions that lead to intrauterine growth restriction and low birth weight and poses a great challenge to control as it occurs in asymptomatic malaria parasitaemia.<sup>13-18</sup>

This study was aimed at determining the prevalence of placental malaria and neonatal outcomes among unbooked pregnant women presenting in labour in the Federal Teaching Hospital, Abakaliki, Southeastern Nigeria. Those detected with the problem were given appropriate treatment in the Teaching Hospital.

## MATERIALS AND METHODS

This was a prospective cross-sectional study conducted in the labour ward complex of the Federal Teaching Hospital, Ebonyi state, Southeastern Nigeria, over a period of 3 months from March to May, 2012. This is the major tertiary hospital located in the state capital Abakaliki. About 75% of the population are rural dwellers with subsistence farming as the major occupation.<sup>19</sup> Malaria transmission is endemic in the state.

The study population included all the unbooked pregnant women presenting in labour at term within the period of study. The minimum sample size of 206 parturients was calculated using the formula proposed by Daniel, Lwanga and Lameshow.<sup>20-22</sup> The exclusion criteria were, those who refused consent, preterm delivery, multiple gestation, HIV in pregnancy, antepartum haemorrhage and symptomatic parturients. A structured data sheet was administered to the women by the authors after explaining to them the purpose of the study and informed consent obtained. The information obtained were of age, parity, gestational age and social class. Blood sample was collected from the antecubital vein under aseptic procedure. A thick and thin blood films were prepared immediately on two glass slides labeled for each parturient. At delivery, cord blood was collected and preserved in a labeled EDTA bottle for foetal haemoglobin estimation. A large placental biopsy was collected and fixed in 10% neutral-buffered formalin for histopathological studies.

The blood films were allowed to air dry and the thin films fixed with absolute methanol for 1 minute, then both were stained with 5% Giemsa stain for 20 minutes, rinsed with water, air dried and microscopic examination done under oil immersion at 100× magnification. The fixed placental tissue biopsies were processed in the histopathology unit, embedded in paraffin wax and sectioned onto slides by standard technique. The sections were stained with haematoxylin-eosin stain for detection of infection. The foetal haemoglobin estimation was done using the

hemocue system (hemocue AB, Angel-holm Sweden) which consist of a pre-calibrated, portable, battery or main operated photometer. No dilution was required as blood was run by capillary action directly into a curvette containing sodium nitrite and sodium azide that convert the haemoglobin to azide methaemoglobin. The absorbance was then measured at a wavelength of 565 and 880nm. This is reliable as high level of bilirubin, lipids or white blood cells do not affect measurements.<sup>23</sup> Anaemia was determined using the WHO standard of haemoglobin concentration less than 11.0g/dl. The neonates were weighed with an electronic weighing machine and the birth weight recorded. The coded data was fed into the computer using 2008 Epi Info™ statistical software (version 3.5.1, Centers for Disease Control, Atlanta, GA, USA) and analysis done. A *P*-value < 0.05 was considered significant. The Research and Ethics Committee of the Federal Teaching Hospital approved the study protocol.

## RESULTS

Two hundred and fifty women were screened for asymptomatic malaria parasitaemia and placental parasitisation during the study period. Out of this number, (194/250) 77.6% had peripheral parasitaemia while (227/250) 90.8% had placental parasitisation. Two hundred and fifty babies were delivered out of which 141/250 (56.4%) were male neonates while 109/250 (43.6%) were female neonates. The mean birth weight in the study was 2.9 ± 0.6 kg and ranged from 1.4 to 4.5 kg. The prevalence of low birth weight in the study was 41/250 (16.4%) neonates. One hundred and twenty-six (50.4%) neonates weighed ≥3.0 kg while 83/250 (33.2%) neonates weighed between 2.5 and 2.9 kg. A total of 39/41 (95.1%) of the low-birth-weight neonates were deliveries of parasitaemic mothers. The 2/41 (4.9%) low-birth-weight neonates delivered to parturients without peripheral parasitaemia, however, had placental parasitisation. All the neonates that weighed less than 1.5 kg were deliveries of parasitaemic mothers. This was statistically significant ( $X^2 = 43.70$ , *P*-value <0.001) [Table 1].

Sixty-six of the neonates that weighed 3.0 kg and above were males while 60 were females. Only three neonates weighed less than 1.5 kg, two males and one female.

**Table 1: Birth weight and asymptomatic malaria parasitaemia**

Birth weight	With parasitaemia		Without parasitaemia		$X^2$	<i>P</i> -value
	Number (194)	%	Number (56)	%		
≥3.0 kg	76	60.3	50	39.7	43.70	<0.001
2.5-2.9 kg	79	95.2	4	4.8		
1.5-2.4 kg	36	94.7	2	5.3		
<1.5 kg	3	100.0	—	—		

There was no statistically significant difference in birth weight between male and female neonates ( $X^2 = 3.19$ ,  $P$ -value = 0.3631)[Figure 1].

Table 2 showed birth weight distribution and placental parasitisation. All the low-birth-weight babies were deliveries of the parturients with placental parasitisation. No neonate among those without placental malaria weighed less than 3.0 kg. In the study, 190 parturients with placental parasitisation had peripheral parasitaemia while 37 did not have. Only four parturients had peripheral parasitaemia without placental parasitisation [Table 3]. The difference was statistically significant ( $X^2 = 49.08$ ,  $P$ -value < 0.001).

Among the parturients with peripheral parasitaemia, 21.1% had birth weight <2.5 kg while 18.0% of those with placental parasitisation had birth weight less than 2.5 kg. There was no neonate with haemoglobin concentration less than 11.0 g/dl. Logistic regression showed negative correlation

between malaria parasitaemia with foetal haemoglobin concentration. The only parasite specie identified during the study period was *Plasmodium falciparum*.

**DISCUSSION**

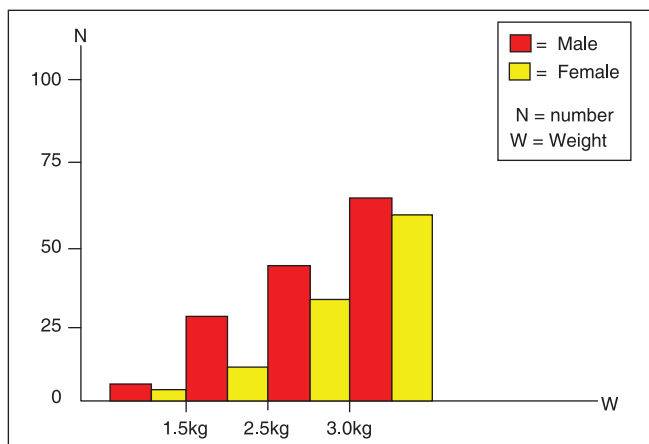
There was high prevalence of asymptomatic malaria parasitaemia and placental parasitisation in this study, 77.6% and 90.8%, respectively. This shows the superiority of placental histology over blood film in the diagnosis of asymptomatic malaria parasitaemia in pregnancy. This compares with the report by Rogerson *et al.*,<sup>18</sup> of 91% for placental histology, 63% for placental blood film and 47% for peripheral blood film. This was however slightly lower than the 97.4% reported by Achang – Kimbi JK *et al.*<sup>24</sup> There was no neonatal anaemia in the study but logistic regression showed negative correlation between neonatal haemoglobin concentration and asymptomatic malaria parasitaemia as shown by negative coefficient of -0.904 and correlation coefficient ( $r^2$ ) of 0.04. The prevalence of low birth weight in the study was 21.1% for peripheral parasitaemia and 18.0% for placental parasitisation. This compares with the report of 19.0% in previous studies in sub-Saharan Africa.<sup>25</sup> Low birth weight following asymptomatic malaria parasitaemia is an important contributor to neonatal and infant morbidity and mortality in malaria endemic areas like ours. This may be due to placental malaria and associated host response-induced changes in placental structure and function affecting pregnancy growth hormone and predisposing the offspring to hypertension and vascular dysfunction leading to foetal growth restriction, significantly lower birth weight and ponderal index.<sup>26-28</sup> No neonate, however, died or had features of malaria during the course of the study and none had any problem at 6-weeks postnatal clinic. There was no statistically significant difference in birth weight between male and female neonates in the study suggesting that foetal sex may not be a factor in the neonatal outcome of asymptomatic malaria parasitaemia in pregnancy.

**Table 2: Birth weight and placental malaria parasitisation**

Birth weight	Placental malaria parasitisation			
	Present		Absent	
	Number (227)	%	Number (23)	%
≥3.0 kg	103	45.4	23	100.0
2.5-2.9 kg	83	36.6	—	—
1.5-2.4 kg	38	16.7	—	—
<1.5 kg	3	1.3	—	—

**Table 3: Placental parasitisation and peripheral parasitaemia**

Placental parasitisation	Peripheral malaria parasitaemia		$X^2$	$P$ -value
	Present (194)	Absent (56)		
Present	190	37	49.08	<0.001
Absent	4	19		



**Figure 1: Foetal birth weight and sex distribution**

**CONCLUSIONS**

There was high prevalence of placental malaria parasitisation in the study group. Placental histology is a more sensitive method of diagnosing malaria in pregnancy at delivery when compared with peripheral blood film for malaria parasitaemia. Asymptomatic malaria parasitaemia and placental parasitaemia did not result in a corresponding foetal anaemia on the babies delivered. *Plasmodium falciparum* was the only type of *Plasmodium species* identified in the blood film.

**RECOMMENDATIONS**

There is need to scale up preventive measures (ITN, IPT and effective treatment) against malaria in pregnancy due

to the observed foetal impact of asymptomatic malaria parasitaemia. Vector control measures should also be put in place to reduce the level of malaria transmission in our environment. Adequate counselling and advocacy is required to encourage early antenatal booking and proper assessment of the services available during the antenatal care.

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