



Research Article

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Nutritional Status and Asymptomatic Infection in Young Children in Two Different Endemic Areas after Seasonal Malaria Chemoprevention Campaign in Mali

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Abstract

Undernutrition represents an additional burden in Mali where malaria remains a major public health problem. A cross sectional survey was carried out in November 2018 and 2019 at the end of the Seasonal Malaria Chemoprevention (SMC) campaign to investigate the association between asymptomatic infection and nutritional status. A total of 541 children aged 6 to 59 months were included, 233 (43.0%) in Koila Bamanan (in an irrigated area with longer seasonal malaria transmission) and 308 (57.0%) in Kita. Data on parasitemia and nutritional status were collected and analyzed using logistic regression. Overall, underweight prevalence was 16.9% in Kita vs. 14.6% in Koila ($p = 0.4$); 25.6% vs. 40.8% ($p = 0.001$) for stunting and 14.9% vs. 10.7% ($p = 0.001$) for wasting in Kita and Koila, respectively. The prevalence of asymptomatic infection was 17.9% in Kita vs. 6.2% in Koila among children 6-23 months ($p = 0.045$). In multivariate analysis, anemia was associated with underweight (adjusted OR = 2.92; 95%IC [1.59-5.38]), and stunting (adjusted OR = 3.17, 95%IC [2.02-4.98]). Asymptomatic infection was associated with underweight (adjusted OR = 2.41, 95%IC [1.21-4.80]). Stunting was higher in children living in Koila (adjusted OR = 2.44, 95%IC [1.60-3.71]). However, the reduction in stunting was observed in children 24-59 months (adjusted OR = 0.44 95%CI [0.28-0.96]). A significant correlation was observed between parasite density and underweight in Kita (0.049). Undernutrition remains high among children in both areas and was associated with asymptomatic malaria infection.

Keywords: Nutritional status, Asymptomatic infection, Young children, Mali

Abbreviations: ACF/AAH: Action Contre la Faim/Action Against Hunger; ENA: Emergency Nutrition Assessment; GIS/UCRC: Geographic Information System/ University Clinical Research Center; ICEMR: International Center for Excellence in Malaria Research; MPV: Millennium Promise Village; NGO: Non-Governmental Organization; NMCP: National Malaria Control Program; SMART: Standardized Monitoring and Assessment of Relief and Transitions; WHO: World Health Organization

Background

Undernutrition represents an additional burden in Sub-Saharan Africa countries where malaria remains a major public health problem [1]. Most of deaths in children under five years linked to undernutrition occur in low-income countries [2]. National

prevalence of undernutrition varies according to residence areas in Mali. The various forms of undernutrition represent respectively 24.1%, 18.6% and 10% for stunting, underweight and wasting in children under five years [3].



Among factors associated to undernutrition occurrence in children under five years, infectious disease, especially malaria and geohelminth were mainly reported in sub-Saharan Africa countries [4-6]. Undernutrition and malaria are common in children under 5 years old, mutually worsen each other especially in malaria endemic areas. Their coexistence has a negative impact on child's learning, cognitive and physical development and responsible of most death in target population [5,7,8].

World Health of Organization (WHO) recommends Seasonal Malaria Chemoprevention (SMC) to prevent malaria in children under five years during malaria transmission in the sub-Saharan Africa region. It consists to monthly treatment of Sulfadoxine-Pyrimethamine (SP) plus Amodiaquine (AQ) during 4 months in high transmission season. Children received SP the 1st day and AQ for 3 days [9]. The authors reported that SMC has effective, safe and cost-effective to prevent malaria in target population [10-12] and could improve children's nutritional status in malaria endemic areas during its implementation period. This study was carried out in two endemic areas: Koila village, endemic malaria area Northeast from Bamako and Kita, another endemic area West from Bamako; to investigate the association between nutritional status and malaria after the last round of SMC and the possible effect of SMC in nutritional status in perspective to combine the two diseases control interventions ongoing.

Materials and Methods

Study Site and Participants

This study is part of pilot study on SMC implementation in Kita funded by Save the Children and President's Malaria Initiative (PMI), and International Center for Excellence in Malaria Research (ICEMR) on malaria control interventions impact in Koila, both in collaboration with Malian National Malaria Control Program (NMCP). The study was carried out in two eco-climatic areas:

- a) Koila Village, at 45km from Markala district health and 385km Northeast of Bamako. It is an irrigated area with longer seasonal malaria transmission. Malaria control interventions ongoing are seasonal malaria chemoprevention, malaria cases management after rapid test diagnosis, use of insecticide treated nets and intermittent preventive treatment in pregnant women. Koila was a millennium challenge experimental site until 2016 and currently ICEMR project, UNICEF and Terre des Hommes International non-governmental organization doing research on malaria and malnutrition, respectively.
- b) Kita, another endemic area at 186km West of Bamako with many rivers. Malaria transmission is also intense during rainy season. President's Malaria Initiative (PMI) and Action Against Hunger has undertaken different activities in children since 2014 in collaboration with NMCP.

Study Design & Data Collection

The cross-sectional survey was carried out in November 2018 in Kita and 2019 in Koila, one month after the last round of SMC campaign to investigate association of children's nutritional status and malaria infection. The study protocol has already been approved by Ethical Committee of Faculties of Medicine, Pharmacy and Odontostomatology of University of Sciences, Techniques and Technologies of Bamako. After obtaining written informed consent from the parents/guardians, temperature, age, weight and height were measured in each child during physical examination, blood sample were collected to perform blood smear and measure hemoglobin level.

Blood smear was used to determine parasitemia by microscopy. The thick blood films were stained with 10% Giemsa and examined under the $\times 100$ oil immersion objective lens of a light microscope. The number of asexual parasites was counted on 200 leucocytes and sexual parasites on 1000 leucocytes. Hemoglobin level was measured from whole blood using Hemocue 301. Electronic thermometer was used to determine axillary temperature. Malaria infection was defined as having a positive blood smear with Plasmodium, anemia by Hemoglobin (Hb) $< 11\text{g/dl}$ and the fever for any temperature $> 37.5^\circ\text{C}$. Each case of malaria or undernutrition has been referred to community health center for best management.

Sample Calculation & Data Analysis

Sample size has been estimated from the national prevalence of undernutrition among children under five years (23.1%) in Mali, assuming a confidence level at 95%, cluster effect at 2 and 10% of lost to follow up, a sample size was 542 participants. Data on parasitemia and nutrition were collected by using electronic data capture and exported in Excel 2010 and analyzed with STATA 14. Anthropometric indices were calculated with ENA for SMART. Logistic regression was used to determine association between nutritional status of children and malaria infection. Pearson correlation was used to determine the correlation between parasite density and z-score of anthropometric indices with a significant p value at 0.05.

Asymptomatic malaria infection was defined as having Plasmodium falciparum in blood smear without fever, and anemia was defined as Hemoglobin (Hb) $< 11\text{g/dl}$. The undernutrition was defined by z-score < -2 standard deviation for height/age (stunting), weight/age (underweight) and weight/height (wasting) [13].

Results

A total of 541 children under five years were enrolled in this study. Age group 24-59 months represents 81.8% in Kita vs. 72.1% in Koila, and 18.2% vs. 27.9% for those 6-23 months. The prevalence of asymptomatic malaria infection was 17.9% vs. 6.2% among

children 6-23 months in Kita compared to Koila ($p=0.045$); 15.5% vs. 14.9% for those 24-59 months ($p=0.8$). Anemia prevalence was 63.9% among 24-59 months in Kita against 48.2% in Koila ($p = 0.001$); and similar between those 6-23 months in both areas. Underweight prevalence was 23.2% vs. 18.5% among 6-23 months in Kita compared to Koila ($p = 0.5$); and 15.5 vs. 13.1% in 24-59

months ($p = 0.4$). Among children 6-23 months, stunting was high in Koila (73.9%) compared to Kita (19.6%) ($p = 0.001$); and similar in those 24-59 months. The prevalence of wasting was 28.6% in Kita vs. 1.5% in Koila among children 6-23 months ($p = 0.001$); but similar in 24-59 months (Table 1).

Table 1: Characteristics of participants in Kita and Koila.

Characteristics	6-23 months			24-59 months		
	Kita n (%)	Koila n (%)	p	Kita n (%)	Koila n (%)	p
Parasitemia	10(17.9)	4(6.2)	0.045	39(15.5)	25(14.9)	0.8
Anemia	39(69.6)	42(64.6)	0.5	161(63.9)	81(48.2)	0.001
Gametocyte carriage	4(7.1)	3(4.6)	0.5	12(4.8)	11(6.6)	0.6
Underweight	13(23.2)	12(18.5)	0.5	39(15.5)	22(13.1)	0.4
Stunting	11(19.6)	48(73.9)	0.001	68(27)	47(28)	0.8
Wasting	16(28.6)	1(1.5)	0.001	30(11.9)	24(14.3)	0.4

For afebrile children, multivariate analysis revealed that anemia was associated with underweight (AOR = 2.92; 95% CI [1.59-5.38]) and stunting (AOR = 3.17, 95% CI [2.02-4.98]). Asymptomatic malaria infection was associated with underweight (AOR = 2.41, 95% CI [1.21-4.80]). Children living in Koila had a

strong risk to develop stunting (AOR = 2.44, 95% CI [1.60-3.71]). However, age group 24-59 months was associated with stunting reduction in the both areas (AOR = 0.44 CI 95% [0.28-0.96]). Any significant association between wasting and parasitemia, anemia and gametocyte carriage was found (Table 2).

Table 2: Associated factors with nutritional status among afebrile children in both sites.

Independents Variables	Underweight			Stunting			Wasting		
	AOR	CI 95%	P	AOR	CI 95%	P	AOR	CI 95%	P
Study site									
Kita	Ref.			Ref.			Ref.		
Koila	0.9	[0.53-1.53]	0.7	2.44	[1.60-3.71]	0.001	0.75	[0.42-1.33]	0.3
Age group									
6-23 months	Ref.			Ref.			Ref.		
24-59 months	0.66	[0.38-1.17]	0.1	0.44	[0.28-0.96]	0.001	0.79	[0.42-1.48]	0.4
Asymptomatic parasitemia	2.41	[1.21-4.80]	0.01	1.43	[0.75-2.71]	0.2	1.88	[0.87-4.03]	0.1
Gametocyte carriage	0.41	[0.12-1.36]	0.1	0.56	[0.21-1.45]	0.2	0.53	[0.14-2.01]	0.3
Anemia	2.92	[1.59-5.38]	0.001	3.17	[2.02-4.98]	0.001	1.02	[0.58-1.80]	0.9

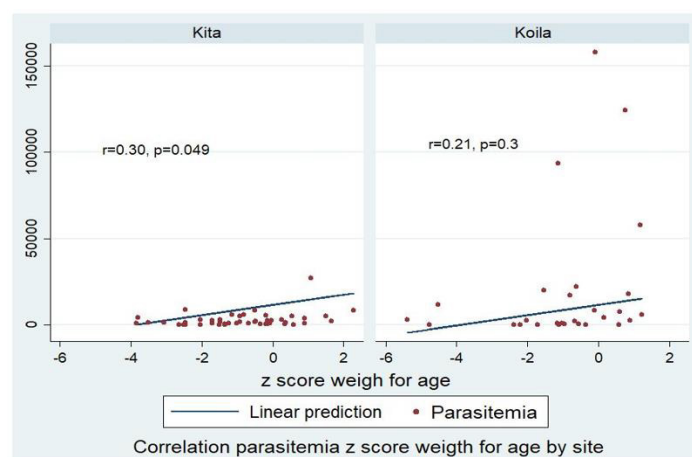


Figure 1: Correlation between parasite density and anthropometric indices weight for age.

A significant correlation was found between parasite density and z-score of underweight children in Kita ($r=0.30$, $p=0.049$) (Figure 1). This correlation was not significant in wasting children

(Figure 2), but a trend of small decrease in parasite density in children having stunting in both study sites was observed (Figure 3).

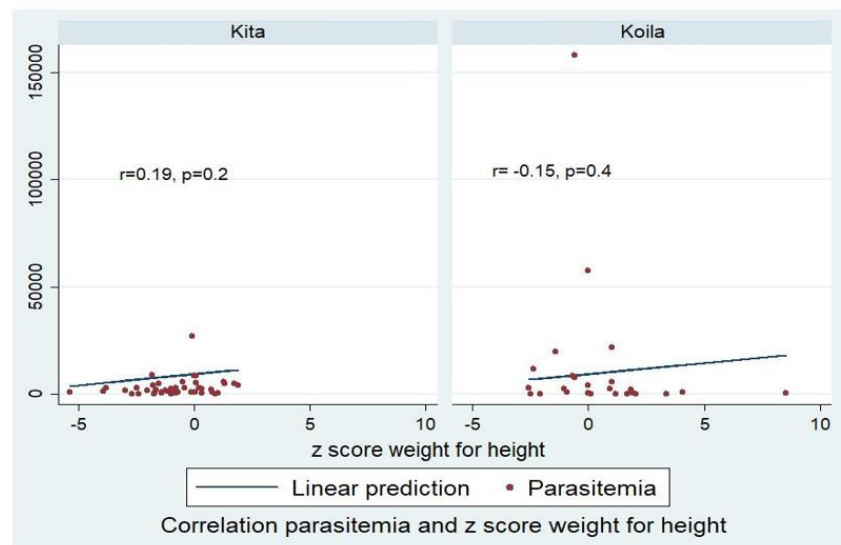


Figure 2: Correlation between parasite density and anthropometric indices weight of height.

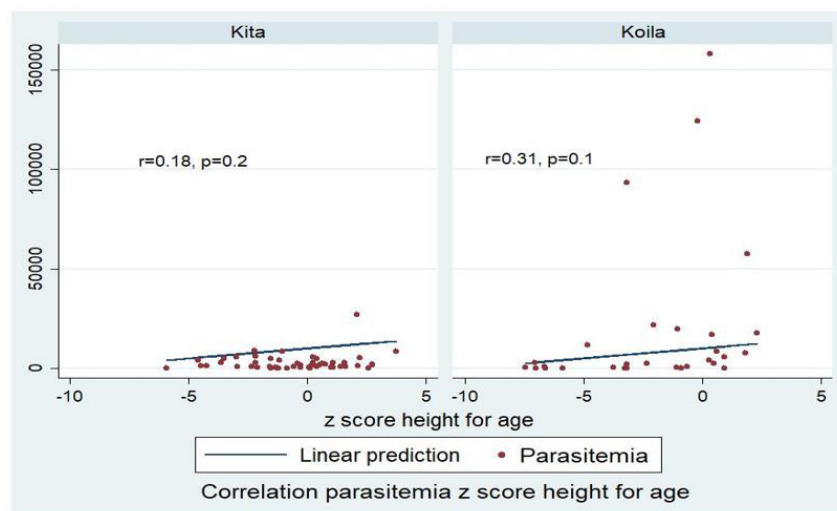


Figure 3: Correlation between parasite density and anthropometric indices height for age.

Discussion

Malaria and undernutrition are frequently associated Malian children under five years and are a major cause of their mortality [14]. This study was carried out to investigate undernutrition and their association with malaria infection among children living in two endemic areas after SMC campaign. Children aged 24 to 59 months were the most represented in both sites (Table 1) as reported generally in Mali [3,15]. Malaria infection was frequent among children aged 6 to 23 months in Kita ($p = 0.045$), but similar in those 24-59 months in both sites ($p = 0.8$). Otherwise, the prevalence of anemia was higher among 24-59 months in Kita

($p=0.001$). Anthropometric indices of children nutritional status varied according to residence areas and age groups. Underweight and wasting were more frequent in Kita and stunting in Koila among children 6-23 months, but it was similar among children aged 24 to 59 months in both sites (Table 1).

Undernutrition situation remains a major concern in terms of public health in both sites [3], despite the efforts of the Non-Governmental Organization (NGO) as Action Against Hunger (AAH) in Kita and Millennium Promise Village in Koila to ensure safe water, good health and food self-sufficiency. Since 2007, AAH was deployed in Kita District and conduct activities in different areas

as health, nutrition, education and safe water. Millennium Promise Village from 2006 to 2016 worked on the same field in Koila, and currently Terre des Hommes carry out activities in Dioro area on food security.

Using logistic regression, anemia was associated with underweight (AOR = 2.92; 95% CI [1.59-5.38]) and stunting (AOR = 3.17, 95% CI [2.02-4.98]). An association was found between asymptomatic malaria infection and underweight (AOR = 2.41, 95% CI [1.21-4.80]). The risk to develop stunting was higher among children living in Koila after adjusting on confounding factors (AOR = 2.44, 95% CI [1.60-3.71]). However, a decrease in stunting was observed in children aged 24 to 59 months (AOR = 0.44, 95% CI [0.28-0.96]) in the two sites. Any significant association was found between wasting and asymptomatic malaria infection, anemia and gametocyte carriage (Table 2). The role of infectious diseases, particularly *Plasmodium falciparum* infection in undernutrition occurrence among children has been already demonstrated in sub-African malaria endemic areas [16,17].

Children living in a pathogenic environment where they are exposed to repeat infectious diseases contribute to nutrient deficit, leads food imbalance and thereafter undernutrition [7,14]. Also, a repetitive infectious disease quickly induces malnutrition among children already underweight in those areas [18-20]. A small significant correlation was found between parasite density and underweight in Kita ($r=0.3$, $p=0.049$) (Figure 1). Any significant correlation was not found between parasite density and others stunting and wasting according to anthropometric indices, but a small trend in parasite density decrease in children having stunting was observed in both sites (Figure 3). In *Plasmodium falciparum* endemic areas, children having stunting and underweight were more infected by *Plasmodium*, but parasite density remains lower [21,22].

Seasonal malaria chemoprevention with SP/AQ decrease malaria infection and reduce *Plasmodium falciparum* parasite density in children. All children in this study received four rounds of SMC during malaria season. Combine nutritional interventions to seasonal malaria chemoprevention in malaria endemic areas could reduce parasite density and improve nutritional status in children [23]. Other parameters as family income, mother's characteristics, other pathologies were not investigated and could be influenced children nutritional status [24-26] are the limit of this study.

Conclusion

Undernutrition remains high among children in both sites and was associated with asymptomatic malaria infection. More studies are required to assess the treatment compliance during SMC and investigate another factor associated with nutritional status in children.

Declaration

Ethic approval and Consent to participate. This study as part of the ICEMR protocol was already approved by ethical committee of Faculty of Medicine, Pharmacy and Odontostomatology, University of Sciences, Techniques and technology of Bamako under N°2019/04/FMPOS revised. All participants given their agreement before data collection.

Consent for Publication

- i. All authors read and approved the final version of the manuscript before submission.
- ii. Availability of data and materials
- iii. Contact the corresponding author for any information about data availability.

Conflict of Interest

The authors declare any competing interests.

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Author's contributions

- MD, SD, SD: Study design
 LD and BK: data acquisition
 DK, SD and MT: Data analysis and interpretation
 DK, SD and MT: wrote main manuscript text.
 DK, SD, MT, IS, NS, FK: reviewed the manuscript.
 Final version: MD, SD

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