

Impact of Single Dose Praziquantel Treatment on *Schistosoma haematobium* Infection among School Children in an Endemic Nigerian Community

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Abstract: Schistosomiasis is prevalent in Nigeria, and the foremost pathogen is *Schistosoma haematobium*, which affects about 29 million people. Single dose of the drug praziquantel is often recommended for treatment but the efficacy has not been documented in certain regions. Therefore, this study was designed to assess the impact of single dose praziquantel treatment on *S. haematobium* infection among school children in an endemic community of South-Western Nigeria. Urine samples were collected from 434 school children and 10 ml was filtered through Nucleopore filter paper before examination for egg outputs by microscopy. The prevalence was 24.9% at pre-treatment. There was no statistically significant difference for the prevalence of infection between males (14.7%) and females (10.2%), although the mean egg count for the females (9.87) was significantly more ($P < 0.05$) than the males (6.06). At 6 and 12 months post-treatment there was 74.4% and 86.4% reduction in the mean egg count, respectively. Interestingly, an increased prevalence of infection from 2.1% at 6 months to 7.7% at 12 months post-treatment was observed, nonetheless the mean egg count was reduced to 0.27 at 12th month from 1.98 at 6 months post-treatment. Resurgence in the prevalence rate between 6 and 12 months post-treatment with praziquantel is herein reported and the need for a follow-up treatment in endemic areas for adequate impact on schistosomiasis control is discussed.

Key words: *Schistosoma haematobium*, school, children, praziquantel

INTRODUCTION

The presence of *Schistosoma haematobium* and *Schistosoma mansoni* in Nigeria has been documented since 1881 from accounts of the German traveler, Nachtigal who passed through the eastern region of Borno [1]. According to the report, it was introduced by Fulani herdsmen arriving from the Upper Nile valley [1]. Today, Schistosomiasis is recognized as one of the most devastating neglected tropical diseases (NTDs) in sub-Saharan Africa, and it is estimated that 779 million people are at-risk of infection while about 250 million people are currently infected [2,3] producing a disease burden that could exceed that

of malaria, which according to the Global Disease Burden study of 2010 was 3.31 million disability-adjusted life years (DALYs) and 11,700 deaths per year [3-6]. More worrisome is the fact that female genital schistosomiasis is a co-factor in the transmission of HIV [7,8], and Nigeria is categorized as a HIV endemic country. Therefore, schistosomiasis continues to be of great public health significance in Nigeria, and the epidemiology varies greatly within and between regions of the country [9]. Nigeria has the greatest number of cases of schistosomiasis worldwide, with both intestinal schistosomiasis caused by *S. mansoni* and the urogenital schistosomiasis caused by *S. haematobium* endemic in the country [10]. Currently, praziquantel is the major focus of preventive chemotherapy which is delivered by mass drug administration to those shown to be, or presumed, to be at-risk of infection and the disease [4]. Provision of praziquantel which cost just a few cents per tablet when given yearly to children could avert a level of morbidity in Africa that exceeds that of malaria or other disease scourges [11]. The realiza-

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tion of this goal can only be attainable with the identification of all endemic communities such as Ipogun community in Nigeria for mass treatment of schistosomiasis and other soil transmitted helminthic infections, especially that these diseases affect mainly the agrarian and rural populations [12]. As of 2006, a majority of Nigerians lack access to safe water coverage (53%) and sanitation (70%), a situation that has largely remain unchanged [13-15]. Praziquantel is the mainstay of the current strategy recommended by WHO against schistosomiasis for morbidity control through preventive chemotherapy [4,16,17]. Successful control programmes in China, Egypt, Brazil, Philipines, Japan and Uganda show that treating the total eligible population in areas that are highly endemic for schistosomiasis with praziquantel for a few years results in substantial reductions in the indices (e.g., prevalence, parasite intensity, and anaemia) of the disease and in the amount of maintenance drug required [18]. Despite WHO member states endorsement of the WHA 65.21 resolution on elimination of schistosomiasis [4] there is the need to identify pockets of infections as well as the effectiveness of praziquantel through the adoption of the right treatment strategy which may vary from one country to another depending on the community level of infrastructural development, health promotion and education. Yet over a decade after a World Health Assembly's resolution to provide complete drug coverage for schistosomiasis, less than 10% of school-aged children at risk received praziquantel [17,19]. This present study which was a longitudinal study evaluates the effectiveness of praziquantel in the treatment of schistosomiasis and also to monitor possible development of resistance in the use of praziquantel for the control of schistosomiasis.

MATERIALS AND METHODS

Study area

The study was conducted in Ipogun community in Ondo state of Nigeria. It is situated in the tropical rainforest belt of Western Nigeria. It is a rural community that enjoys electricity and it lies between latitude 5°6'N and 8°2'N and longitude 4°17'E and 6°17'E. There are 4 elementary schools and 1 secondary school in the community. Three of the four elementary schools are government owned schools while one is a private school. The inhabitants of the community are mainly farmers. There are very few water wells in the community therefore residents depend mainly on river Apogun which traverses the community for their water supply.

Consent and ethical approval

Prior to commencement of the study administrative approval was obtained from the State Ministry of Health through a letter and the Local Government Primary Health Care Department during advocacy visit. Ethical approval for the protocol was obtained from the Institutional Review Board of Nigerian Institute of Medical Research (IRB/14/237). The community head was duly informed and the parents were adequately enlightened on the purpose of the study to obtain their verbal consent and assent of the children. All the head teachers and teachers of the schools were adequately enlightened and mobilized for the study prior to commencement.

Study design

The inclusion criteria were: children who enrolled in the study school; child must assent to participate. The exclusion criteria were: children who did not enroll in the study school; child who did not assent to participate.

All the participants (age range 5-18 years old) were enrolled in the survey according to the school enrollment list in the class register. Each participant was given a survey code which included the school code, class and serial number which was the same as the code on the specimen containers for urine samples. Other demographic data such as sex and age of each child was recorded during the enrollment for the survey. Urine samples were collected from all the participants that consented to participate in the study between 10.00 and 14.00 hr [20]. The frequency of visible haematuria was also recorded. Filtration technique using nucleopore filter papers with swinnex filter holder for 10 ml of urine and microscopy was employed in examining the nucleopore filter papers for the presence and the intensity of *S. haematobium* eggs which was recorded as egg output per 10 ml of urine [20]. All the recruited participants were treated with single oral dose of praziquantel (600 mg, Batch No: M70116) at a dose of 40 mg/kg body weight immediately after submitting urine samples by directly observed treatment. Urine samples of the participants which were processed as described above were examined at pre-treatment, 6 and 12 months post-treatment intervals to determine changes in parasites intensities and prevalence with time.

Data analysis

Demographic data and results from the examined samples were analyzed using IBM SPSS version 20 (SPSS, Chicago, Illinois, USA). Results were presented using descriptive statistics

in frequencies, percentages and charts. Inferential statistics was also performed using chi square to test the significance of variation of infection between age group, sex and schools were tested by chi-square tests.

RESULTS

A total of 434 children (male 56.2%; female 43.8%) were examined for *S. haematobium* egg (Fig. 1) infection at baseline in the 4 elementary schools in the community with ages ranging from 5 to 18 years and mean age 9.16 ± 2.45 . At the end of 12 months we had 95.6% of the children still in the study. The drop-out rate of the children for the period of 12 months was 4.4% being the total number of children that were not present till the end of the survey at 12 months.

The prevalence of infection at pre-treatment was 24.9% (Males 14.7%; Females 10.2%) although there was no statistically significant difference ($P > 0.05$) in infection rate between male and female. The mean urine egg count of the samples col-

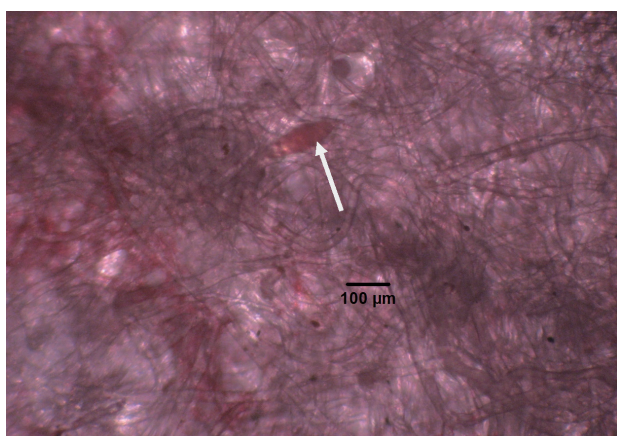


Fig. 1. Egg of *Schistosoma haematobium* (arrow), stained with eosin on Whatman filter paper. An estimated size is $175 \times 60 \mu\text{m}$.

lected from female students (9.87) was greater than that of the males (6.06). Age group 9-10 years had the highest prevalence of 30.5%, followed by age group 7-8 years with a prevalence of 29.5% while age group 13-14 years had the lowest prevalence of infection (Table 1). There was significant difference in the prevalence of infection among the age groups ($P < 0.05$).

Prevalence of infection by school ranged from 14% to 46.9% ($P < 0.001$). Prevalence of infection among the participants at 6 months post treatment was 2.1% (9 children) which showed a 91.6% reduction in prevalence. Among the 9 infected participants, 6 were new infections while the remaining 3 were part of those infected at pre-treatment. Age groups 7-8 and 9-10 had the highest prevalence of 0.7% each. Females had a higher prevalence of infection 6 (1.4%) at 6 months post treatment when compared to males 3 (0.7%) though the difference was not statistically significant ($P > 0.05$). The prevalence of infection among the participants at 12 months post treatment (7.7%) showed an increase in prevalence over the 6 months prevalence (2.1%) with a total of 32 (7.7%) participants infected at 12 months which was 69.1% reduction in prevalence. This difference was statistically significant (McNemar Test $P < 0.001$). Age group 9-10 years had the highest prevalence of infection 12 (2.9%). Males recorded a higher prevalence of infection 21 (5.1%) when compared to females 11 (2.7%) ($P > 0.05$). The mean urine egg count by sex at 6 months showed that males had a higher (1.99) mean urine egg count than females (1.97). The mean urine egg counts by sex at 12 months post treatment was 0.29 for males and 0.25 for females. There was 74.4% and 86.4% reduction in the mean egg count of the children at 6 and 12 months post treatment respectively while the mean urine egg count was reduced to 0.27 at 12 months from 1.98 at 6 months post treatment. At baseline, 4.4% of the participants had heavy egg infection (Fig. 2) which was reduced to 0.3% at 6 months post treat-

Table 1. Baseline prevalence of infection among the groups by sex and age

Age groups (year)	Male		Female		Both Sexes	
	No. of examined	No. of Infected (%)	No. of examined	No. of Infected (%)	No. of examined	No. of Infected (%)
5-6	44	7 (15.9)	31	4 (12.9)	75	11 (14.7)
7-8	75	27 (36.0)	54	11 (20.4)	129	38 (29.5)
9-10	57	17 (29.8)	48	15 (31.3)	105	32 (30.5)
11-12	39	10 (25.6)	45	14 (31.1)	84	24 (28.6)
13-14	17	1 (5.9)	12	0 (0.0)	29	1 (3.4)
≥15	12	2 (16.7)	0	0 (0.0)	12	2 (16.7)
Total	244	64 (26.2)	190	44 (23.2)	434	108 (24.9)

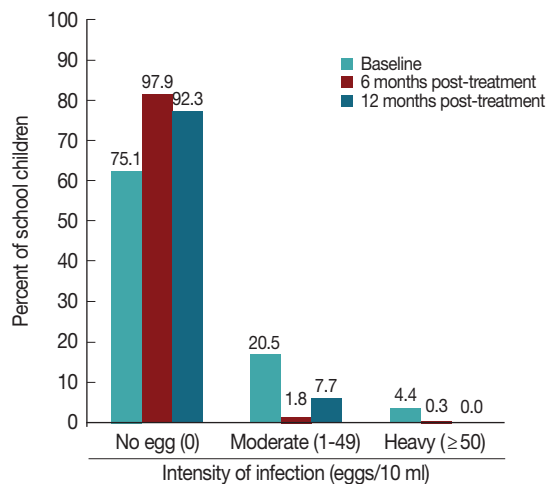


Fig. 2. Changes in *Schistosoma haematobium* egg intensity over a period of 12 months among pupils.

ment and eventually zero at 12 months post-treatment, though more participants (7.7%) had moderate infection at 12 months than at 6 months post-treatment (1.8%). The prevalence of visible haematuria was 1.8%.

DISCUSSION

The infection of *S. haematobium* cuts across all age groups among the children and it could be noted that the older age groups such as age group 13-14 years had lower prevalence than most of the younger age groups. The baseline result showed that *S. haematobium* was endemic in the community though the influence of seasons could not be ascertained despite the fact that samples were collected at different times in 12 months though Tian-Bi et al. [21] identified infected snails exclusively during the dry season. The school with the lowest prevalence was the only private school in the community which had the children of the most enlightened people in the community, whereas the school with highest infection rate is a public school and probably the use of stream water for religious purpose could be the contributory factor to the high infection rate which showed the influence of practice on transmission. Single dose of praziquantel at 40 mg/kg body weight was able to significantly reduce the prevalence and intensity of infection up till 6 months post-treatment. By the 12th month assessment of infection, there was an increased rate in the prevalence of infection among the children, some of whom were not found to be initially infected at baseline. This connotes new infection after treatment had started or some of the

children harbor the juvenile stages of *S. haematobium* which according to Sabah et al. [22] are less susceptible to chemotherapy and could start oviposition by the 5th week of infection. Therefore, there is the need to support treatment activities with health education to avoid either re-infection or new infection of children [23]. Though, there was a significant increase in prevalence at 12th month post-treatment above the recorded 6th month prevalence, the intensity of infection was lower than the 6th month intensity. According to Toure et al. [16], the reduction in the intensity is particularly of importance as high intensity of *S. haematobium* infection has been shown to contribute to morbidity seen in schistosomiasis infection which includes anaemia in children [24]. Contrary to findings of Nsowah-Nuamah et al. [25] in a study in Ghana where the prevalence of infection was reduced by 80-90% at 12 months after treatment, the prevalence in the current study was reduced by 91.6% only for 6 months which was reduced to 69.1% by the 12th month due to maturation of juvenile parasites that were not susceptible to the praziquantel [22] which have started oviposition and new infections. Therefore, our findings necessitate re-treatment with praziquantel after 12 months when prevalence increased contrary to the studies in Ghana [25] and Burkina Faso [16]. Although this is contrary to WHO [26] recommendation that communities with moderate risk ($\geq 10\%$ but $< 50\%$ prevalence) should treat once every 2 years. The uptrend of infection at the 12th month signified the need to constantly treat every year for achievement of elimination by year 2020. This will prevent a resurgence of infection to the pre-treatment level should there be an interruption in the drug distribution. Our present finding reveals the need for continuous efforts at monitoring the disease transmission and treatment coupled with other control measures such as snail control and health education. The current findings showed that WHO strategy of treatment once every 2 years could be adopted in some settings while there is need for some degree of flexibility in accordance to the epidemiological findings of other geographical areas.

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CONFLICT OF INTEREST

The authors have no conflict of interest to report.

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