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Control and elimination of *Schistosoma mansoni* infection in adult individuals on Ukerewe island, northwestern Tanzania: baseline results before implementation of intervention measures

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Abstract

Background Communities living along the shoreline and on the islands of Lake Victoria in northwestern Tanzania remain endemic for schistosomiasis and suffer from the life-threatening morbidities associated with the disease.

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Background

Schistosomiasis remains endemic in Tanzania and the country is the second country after Nigeria with the highest number of schistosomiasis cases [1, 2]. The country is endemic for parasites *Schistosoma mansoni* and *Schistosoma haematobium* [2]. The current study focused on *S. mansoni*, the causative agent of the intestinal form of schistosomiasis, which is highly endemic in the northwestern regions of Tanzania, particularly the regions bordering the Lake Victoria shoreline [3, 4]. Communities living along the shoreline of Lake Victoria bears the highest burden of the disease and its associated morbidities [4–8]. Previous studies have found a high prevalence of hepatomegaly, splenomegaly and periportal fibrosis in school aged children and the adult population [4, 5, 9, 10]. These are chronic manifestations of an *S. mansoni* infection that, if left untreated lead to the development of portal hypertension, oesophageal varices, portal-systematic venous shunts (collateral veins) and haematemesis [11, 12]. In northwestern Tanzania, these *S. mansoni*



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Nevertheless, the control measures particularly the mass drug administration do not cover the adult population. The current project on Ukerewe island aims to close this gap by involving adult community members in the control program. Here we report the baseline results of *S. mansoni* infection and associated hepatosplenic morbidities and factors before implementing the project activities.

Methods A cross-sectional analytical study was conducted with 4,043 participants aged ≥ 18 years living in 20 villages on Ukerewe island, northwestern Tanzania. Individual stool and urine samples were collected and examined using the Kato-Katz (KK) technique and point-of-care circulating cathodic antigen testing (POC-CCA) to identify *S. mansoni* eggs and antigens, respectively. All study participants underwent ultrasound evaluation of *S. mansoni* hepatosplenic morbidities using the Niamey protocol. Rapid diagnostic tests were used to diagnose HIV infection, hepatitis C and chronic hepatitis B. A questionnaire was used to collect demographic data and reported clinical symptoms of study participants.

Results A total of 4,043 participants took part in the study, of which 49.7% ($n=2,009$) and 50.3% ($n=2,035$) were male and female, respectively. The overall prevalence of *S. mansoni* infection was 30.4% (95%CI:29.0–31.9%) and 84.7% (95%CI:83.3–85.9%), respectively, based on the KK technique and the POC-CCA test. The geometrical mean eggs per gram of faeces (GMepg) was 105.3 (95%CI:98.7–112.3% GMepg) with 53.9%, 32.4% and 13.7% of the participants having mild, had moderate and severe intensity of infection. The prevalence of hepatitis C, HIV, and hepatitis B was 0.4%, 2.2% and 4.7%, with 0.2%, 2.2% and 5.4% of the infected individuals coexisting with *S. mansoni* infection. The prevalence of splenomegaly, periportal fibrosis, hepatomegaly, and portal vein dilatation was 40.5% (95%CI: 38.8–42.1%), 48.1% (95%CI:64.4–49.7%), 66.2% (95%CI:4.6–67.7%) and 67.7% (95%CI:66.2–69.2%), with their prevalence varying depending on the demographic information and infection status of the participants. Other detectable ultrasound-related morbidities included ascites (1.7%), collateral veins (18.3%) and gall bladder wall thickness (40.4%). Age groups, gender, reported clinical characteristics, reported non-use of the drug praziquantel, liver imaging pattern, and place of residence remained independently associated with hepatosplenic morbidities.

Conclusion The current study setting is endemic for *S. mansoni* infection and the population has a high prevalence of the disease associated hepatosplenic morbidities characterized by hepatomegaly, splenomegaly, ascites, gall bladder wall thickening, periportal fibrosis and portal vein dilatation. Several demographic, clinical and epidemiological circumstances remained independently associated with *S. mansoni* infection and associated morbidities. These findings call for integrative intervention efforts, starting with whole community MDA that includes all out of schools community members.

Keywords *Schistosoma mansoni*, Hepatosplenomegaly, Risk factors, Adult, Ukerewe, Tanzania

morbidities which are detected using an ultrasound are recorded during communities' surveys [9, 13] and in hospital settings [14]). In the hospital settings, these morbidities are associated with high mortalities in the adult population [14].

The control of schistosomiasis follows the recommendations of the World Health Organization [15]. Currently, the common control measure is mass preventive chemotherapy with praziquantel drugs, with the main target group being the school aged children (SAC), who receive medication within the school environment [15]. Repeated mass preventive chemotherapy regimens against schistosomiasis targeting SAC over the past 15 years have resulted in a decrease in the prevalence and intensity of the infection [16, 17] but have not lead to the elimination of schistosomiasis as a public health problem [16]. Among the challenges that are currently facing the control programme, is the lack of inclusion of the adult population and children under five years of age. The untreated population maintain the transmission of the parasite within the communities and serves as a source

of re-infection to the treated population. Furthermore, untreated adult, or community members outside of the school setting live with an untreated infection for many years, resulting in high morbidity and mortality [5, 18]. For example, in a tertiary hospital in northwestern Tanzania, 50% of adult cases with upper gastrointestinal bleeding are caused by chronic *S. mansoni* morbidities [14] and 25% of these cases die within six months of diagnosis [14]. In light of these challenges and in response to the sustainable Development Goals, that call for "leaving no one behind," the WHO has published a new guideline to combat schistosomiasis. They call for the inclusion of all age groups (≥ 2 years) [15], especially in areas where the prevalence of schistosomiasis is $>10\%$ [15]. In order to reach the stage of elimination, the WHO recommendation emphasizes the need to integrate other intervention measures such as the provision of clean water, the use of improved sanitation facilities and improved hygiene in endemic communities [15]. This should be accompanied by the implementation of snail control strategies to reduce transmission [19].

Planning and implementation of schistosomiasis interventions requires understanding of the disease ecology by identifying areas with remaining sustained transmission, categorizing and characterizing these areas as recommended by WHO and finally, deciding on the type of interventions to implement [15]. Furthermore, for future monitoring of the impact of the intervention on the disease prevalence and associated morbidities, it is important to collect baseline information to serve as measurement indicators during the process monitoring and impact assessment. In that context, here we present the baseline characteristics of the adult (≥ 18 years) community members associated with *S. mansoni* infection, associated morbidities, and their associated factors in 20 villages of Ukerewe district, north-western Tanzania before the implementation of the five-years community-based mass preventive chemotherapy project. It combines adult treatment of the adult individuals, test-and-treat campaigns, public health education and capacity building of the healthcare workers, health facilities and affected communities to expand the provision of praziquantel drug to remote populations.

Methods

Study area

The project activities are being implemented at Ukerewe district, in the Mwanza region of northwestern Tanzania. Ukerewe District is an island in the Lake Victoria located at 2.0299° S and 33.0339° E. The districts have a tropical climate with temperatures between 17.0 °C and 27.0 °C, and the average annual rainfall is 1,090 mm. Rainfall is bimodal and is characterised by a short rainy season between September and January and a long rain season between March and June.

In total, the district consists of 38 islands, of which 15 islands have permanent residents and 23 are fishing camps. Ukara and Nansio are the biggest islands. The district consists of a total of 76 villages, 54 sub-villages and 25 wards spread across four divisions, including Mumbuga, Mumlambo, Ilangala and Ukara. The district has a district hospital located at Nansio (capital of the district), four (4) health centres and 32 dispensaries located on various islands in Lake Victoria (district health report). According to 2022 census, the district had a population of 387,815 [20]. The main economic activities of the inhabitants are fishing, subsistence agriculture (cultivating casava, corn, banana, sweet potatoes, and various fruits), livestock farming and small business.

Intestinal schistosomiasis is highly endemic in the district and the prevalence is high both in children and the adult population [6, 7, 18]. Hepatosplenomegaly and associated sequelae are common in school aged children and the adult population [9]. To date, the control of *S. mansoni* infection in the study area has included mass chemotherapy with praziquantel drug, primarily, focused

on school children to reduce the long-term morbidities associated with the infection. Adult individuals and non-school going children were not included. In contrast, the current study covers the adult population, approximately 55,000 adult in 20 villages. Project activities are implemented in 20 villages (Fig. 1) of the 76 villages, selected based on hospital data and result of previous studies [9, 21], which suggested that these villages were highly endemic for intestinal schistosomiasis and needed to be prioritized for interventions. The villages included in the study are Bugorola, Bugula, Bukiko, Bukindo, Bukondo, Bukungu, Busumba, Bwisya, Chibasi, Chifule, Gallu, Kamea, Kome, Kweru, Muriti, Muritilima, Musozi, Nebuye, Nyamanga and Nyang'ombe (Fig. 1).

Implemented intervention against intestinal schistosomiasis

A five-year community-based health care project is being implemented in 20 villages, in which the following measures are being taken:

- (i) Capacity building of health workers with a focus on the detection of schistosomiasis cases, treatment, and referral of cases of severe hepatosplenic disease associated with hematemesis to nearest health facilities.
- (ii) Improving early detection of patients with chronic manifestations of intestinal schistosomiasis at the community level through training of community healthcare workers/community drug distributors and targeted community and referral to health facilities.
- (iii) Improve community awareness of intestinal schistosomiasis (disease epidemiology, risk factors, modes of transmission, signs, and symptoms), preventive measures and behaviours involving various community stakeholders such as beach management units, village and sub-village administration, community healthcare workers and community drug distributors, owners of fishing camps and cooperatives and various economic groups in the targeted communities.
- (iv) Treatment of chronic diseases caused by intestinal schistosomiasis - establishment of an ultrasound and endoscopy department at the district hospital for the diagnosis and treatment of oesophageal varices and other hepatosplenic complications (ascites, haematemesis, portal hypertension, etc.) and capacity building of healthcare workers in the district hospital through knowledge transfer within primary and tertiary hospitals.
- (v) Annual community-based mass drug administration, preceded by community awareness rising, test-and-treat campaigns, and training of community drug distributors.

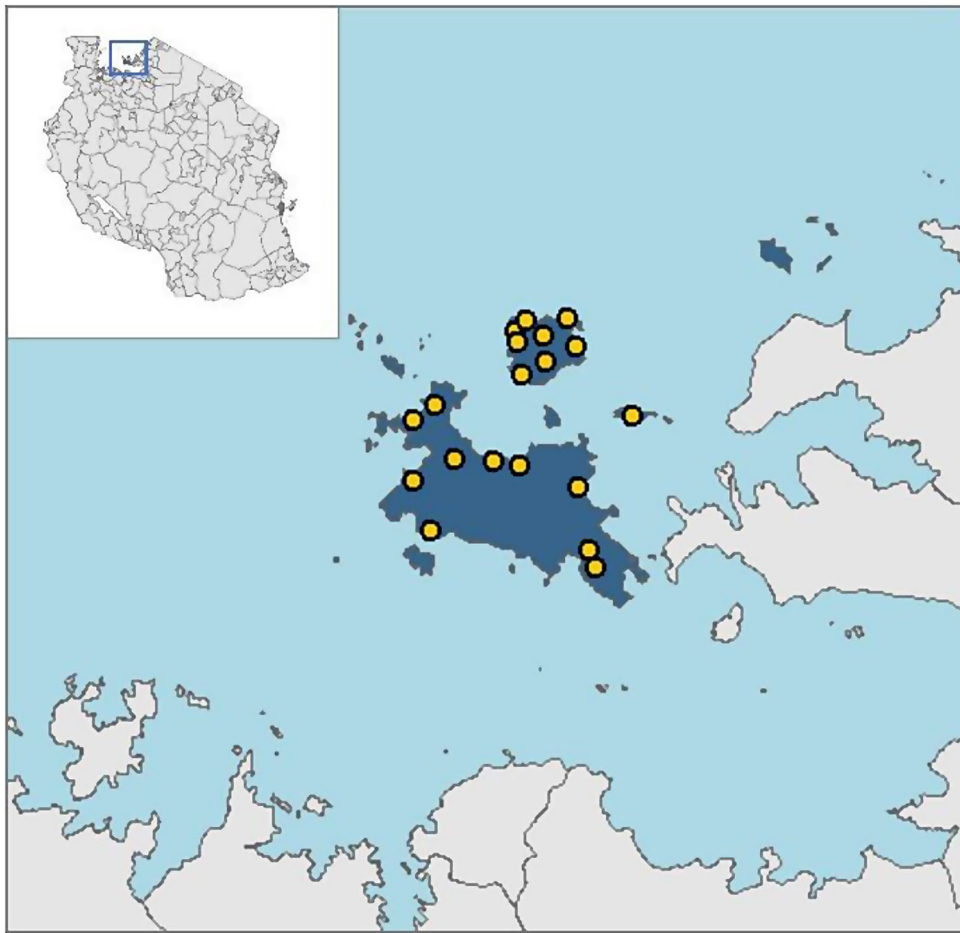


Fig. 1 Location of Ukerewe district and the 20 villages involved in the current study

(vi) Supporting healthcare facilities to ensure regular availability and distribution of praziquantel to community members.

health facilities (ii) refused/declined to complete all study procedures and (iii) reported using praziquantel in the past six months.

Study design, inclusion, and exclusion criteria

This was a cross-sectional analytical study conducted among randomly selected community members in their households. Before starting the study, a population census was conducted to determine the population size of the targeted villages. This allowed participants to be randomly selected from each study village.

Participants were included in the study if they (i) were permanent residents of the target villages of the study and had lived in the village for at least two years, (ii) were ≥ 18 years old, both male and female, (iii) agreed to provide written consent, (iv) participants agreed to provide individual stool and urine samples for testing for *S. mansoni* infections, and (v) had not received treatment with the drug praziquantel in the past six months. On the other hand, participants were excluded from the study if they (i) suffered from a clinical illness requiring treatment in

Sample size and sampling procedures

A single proportion sample size formula for cross-sectional studies was used to calculate the sample size [22]. With a confidence interval of 95%, with the proportion of infected people from the previous study of 0.86 and a tolerable error of 1.1%, a total of 3,823 people had to be included in the study. With a refusal rate to participate of 5%, a total of 191 participants were added. The total sample size required was 4,014. However, a total of 4,043 participants agreed to participate in the study and were enrolled. For the study, 200 participants per villages were recruited and enrolled from 100 randomly selected households of each village.

Data collections

Questionnaire

Each participants was administered a structured questionnaire from the previous study in Tanzania [23] to

collect demographic information (gender, age, village, and sub-village of residence). In addition, the questionnaire collected information on signs and symptoms of intestinal schistosomiasis including history of blood in faeces, history of abdominal pain, history of vomiting blood, current vomiting blood, abdominal distension, and the previous use of praziquantel.

Parasitological examination of *Schistosoma mansoni* infection

A single stool sample was collected in a labelled container. Duplicate thick Kato Katz (KK) smears were prepared from each sample after mixing the stool sample to obtain a homogenous mixture [24]. The thick KK smears were prepared using a 41.7 mg template and were examined for the presence of *S. mansoni* eggs after 24 h of clearance using a light microscope in the Parasitology laboratory of the Catholic University of Health and Allied Sciences, Mwanza, Tanzania. To adhere to quality assurance procedures, 20% of the positive and negative KK slides were re-examined by a third laboratory technician who was not involved in the initial examination of the slides.

Examination of *Schistosoma mansoni* related circulating Cathodic Antigen (CCA) using a point of care test

A single urine sample was collected from a randomly selected sub-sample of the study participants (73% of the total study participants) and tested for the presence of circulating cathodic antigens (CCA) using rapid point-of-care circulating cathodic (POC-CCA) test (<http://www.rapid-diagnostic.com/>). Trace results were considered positive. The use of the rapid test was preceded by refresher training of the laboratory technicians involved in our previous field work with the same diagnostic test. The team using the POC-CCA test to diagnose *Schistosoma* infection was not involved in the preparation or examination of the KK thick smears.

Examination of hepatitis-C and hepatitis B viral infections

A fingerpick blood sample was collected from each participant and examined for the presence of hepatitis C antibodies in whole blood using a rapid qualitative test (ACON Laboratories, Inc., San Diego, CA) [25]. The same fingerpick blood sample was used to detect chronic hepatitis B virus infection using a rapid test for hepatitis B surface antigen (HBsAg) test [25].

Examination of human immunodeficiency Virus-1 infection

A blood sample from fingerprick was used to diagnose HIV infection. According to the Tanzanian national algorithm, two types of rapid HIV antibody rapid tests were used: the SD Bioline and Uni-GOLD (Trinity Biotech PLV, Bray, Ireland). All participants were counselled

before and after HIV testing according to Tanzanian guidelines [26].

Ultrasonographical examination for presence of *S. Mansoni* hepatosplenic morbidities

Height of each study participant was measured with a stadiometer (height was recorded in centimetre) and each of them was clinically examined for liver and spleen consistency before the ultrasonographical examination following established procedures described elsewhere [13]. A pair of ultrasonographers with experience in the Niamey protocol [13] examined all study participants.

To ensure quality, a sub-population of the study participants were examined by both Ultrasonographers and in case of any difference, the study participants were re-examined by both. All identified hepatosplenic morbidities (spleen size, peripheral portal vein branches (PPBs), periportal fibrosis, liver texture patterns, thickness of the peripheral portal vein branches) detected were classified according to the Niamey Protocol [13]. Periportal fibrosis was qualitatively classified based on the liver imaging patterns (A, B, C, D, E and F) [13], with liver imaging pattern A and B being classified as normal.

The size of the left liver lobe was measured along the right para-sternal line and that of the spleen was measured along the craniocaudal axis of it from the left flank side. The portal vein diameter (PVD) was measured at a halfway between the confluence of the splenic vein and the superior mesenteric veins and their bifurcation within the liver [13]. A portal vein diameter of 2 to ≤ 4 SD was considered dilated and >4 SD was considered significantly marked dilated, as described in the Niamey protocol [27].

Data analysis and management

Data were entered using Excel and transferred to Stata Version 17 (StataCorp, statistical software, College Station, TX: StataCorp LP, Texas, USA). All continuous variables were summarized using mean, median and their standard deviations or interquartile ranges depending on data distributions. *S. mansoni* eggs counts were obtained from the two slides and the mean was multiplied by 24 to obtain eggs per gram of faeces. *S. mansoni* geometrical mean eggs per gram of faeces (GMepg) was calculated using the non-logarithmically transformed means. The comparison of mean *S. mansoni* egg counts were done using either student t-test (for two groups) or ANOVA (for more than two groups). Intensity of infection was classified according to WHO criteria in which individuals with 1–99 epg, 100–399 epg and ≥ 400 epg had low, moderate, and heavy intensity of infection [15]. Hepatosplenic morbidities were analysed according to the Niamey protocol after adjustment to the height of the reference population [13]. The same protocol was used to define

Table 1 Reported symptoms and signs associated with *S. mansoni* among study participants

| Variable | Response | Sex | | Statistical test |
|--|----------|---------------|---------------|--------------------------------|
| | | Male | Female | |
| Reported diarrhoea in the past two weeks | No | 1,388 (69.1%) | 1,415 (69.5%) | $\chi^2=0.0937$ $P=0.8$ |
| | Yes | 621 (30.9%) | 620 (30.5%) | |
| Blood in faeces | No | 1664 (82.8%) | 1,756 (86.3%) | $\chi^2=9.3724$ $P=0.002$ |
| | Yes | 345 (17.2%) | 279 (13.7%) | |
| Abdominal pain | No | 435 (21.7%) | 240 (11.8%) | $\chi^2=70.6663$ $P=0.001$ |
| | Yes | 1,574 (78.3%) | 1,795 (88.2%) | |
| Ever vomited blood | No | 1961 (97.6%) | 2011 (98.8%) | $\chi^2=8.4626$ $P=0.004$ |
| | Yes | 48 (2.4%) | 24 (1.2%) | |
| Currently vomiting blood | No | 1,998 (99.5%) | 2030 (99.7%) | $\chi^2=2.3402$ $P=0.1$ |
| | Yes | 11 (0.5%) | 5 (0.3%) | |
| Ever had abdominal distension | No | 1,922 (95.7%) | 1,961 (96.4%) | $\chi^2=1.2743$ $P=0.3$ |
| | Yes | 87 (4.3%) | 74 (3.6%) | |
| Ever received praziquantel (not in the past six month) | No | 894 (44.5%) | 1,284(63.1%) | $\chi^2=139.9255$ $P=0.001$ |
| | Yes | 1,115 (55.5%) | 752 (36.9%) | |

cut-off points between normal and enlarged spleen, left liver lobe and dilatation of the portal vein diameter [13]. Generalized Linear Model (GLM) was used to investigate which risk factors were independently associated with *S. mansoni* infections, periportal fibrosis, left liver lobe and portal vein dilatation (PVD). At bivariate analysis, explanatory variables with P -value < 0.2 were transferred to multivariate analysis. The Prevalence ratio with their 95%CI were generated and presented in the final tables. For all analysis, a P -value of < 0.05 was considered significant.

Treatment

All participants received praziquantel drug (40 mg/kgBWT) regardless of their infection status. The medication was administered under direct observation at the study site within the village. After treatment, participants were asked to remain at the treatment site for one hour to observe any side effects of the medication and to be monitored by the project team.

Results

Demographic characteristics of the study participants

A total of 4,043 participants were enrolled in the study, of which 50.3% ($n=2,034$) were female and 49.7% ($n=2,009$) were male. The overall mean age of the participants was

Table 2 Prevalence of *Schistosoma mansoni* based on Kato Katz technique categorised by age and sex

| Variable | N | Schistosoma mansoni infection | | Statistical test |
|------------------------------|-------|-------------------------------|--------------|----------------------------------|
| | | Positive | Negative | |
| Sex | | | | |
| Female | 2,034 | 570 (28%) | 1465 (71.9%) | $\chi^2(1)=11.2005$ $P=0.001$ |
| Male | 2,009 | 660 (32.9%) | 1349 (67.2%) | |
| Age groups (in years) | | | | |
| 18–25 | 503 | 209 (41.6%) | 294 (58.4%) | $\chi^2(5)=73.4663$ $P=0.001$ |
| 26–35 | 798 | 293 (36.7%) | 505 (63.3%) | |
| 36–45 | 888 | 262 (29.5%) | 626 (70.5%) | |
| 46–55 | 767 | 198 (25.8%) | 569 (74.2%) | |
| 56–65 | 561 | 153 (27.3%) | 408 (72.7%) | |
| ≥ 66 | 527 | 115 (21.8%) | 412 (78.2%) | |

45.3±16.7 years. At enrolment, participants reported various symptoms and signs as presented in Table 1. In relation to sex, blood in faeces ($P<0.004$), ever received PZQ ($P<0.001$) and ever vomited blood ($P<0.004$) were reported more frequently among males' participants. There was no significant difference in relation to age for the following reported symptoms: (i) diarrhoea ($\chi^2(5)=1.4938$, $P=0.9$), (ii) blood in stool in the last two weeks ($\chi^2(5)=1.7392$, $P=0.9$) and (iii) abdominal pain ($\chi^2(5)=7.9583$, $P=0.2$). Having ever vomited blood was reported more frequently in the age group 36–45 years (1.6%), 46–55 years (3%) and 56–65 years (2.1%) (Exact=0.1), currently vomiting blood (Exact=0.9) and ever had abdominal distension problems ($\chi^2(5)=5.1942$, $P=0.4$). In the older age groups ≥ 36 years, the proportion of individuals who had ever taken PZQ treatment was higher compared to the younger age groups ($\chi^2(1)=164.2949$, $P=0.001$).

Prevalence and intensity of *Schistosoma mansoni* infection

The overall prevalence of *S. mansoni* infection based on Kato-Katz-Technique was 30.4% (95%CI:29.0-31.9%), with male participants having a higher prevalence than female participants (32.2% versus 28%, $\chi^2(1)=11.2005$, $P=0.001$). The youngest age groups 18–25 years and 26–35 years had the highest prevalence compared to the older age groups ≥ 36 years (Table 2). Figure 2 shows the prevalence of *S. mansoni* categorised by age groups.

The geometrical mean of eggs per gram of faeces (GMepg) was 105.3 (95%CI:98.7-112.3 GMepg), with males having the highest GMepg value of 113.8 (95%CI:104.0–124.4 GMepg) while the GMepg value of females was 96.3 (87.8-105.6 GMepg) ($t=3.6076$, $P=0.001$). There was a significant difference in the intensity of infection between the age groups, with the youngest age groups 18–25 years (126.6–95%CI:100.3-131.9 GMepg) and 25–35 years (115.04-95%CI:100.3-131.9 GMepg) having the highest intensity of infection ($F=9.61$, $P=0.001$). In relation to infection intensity

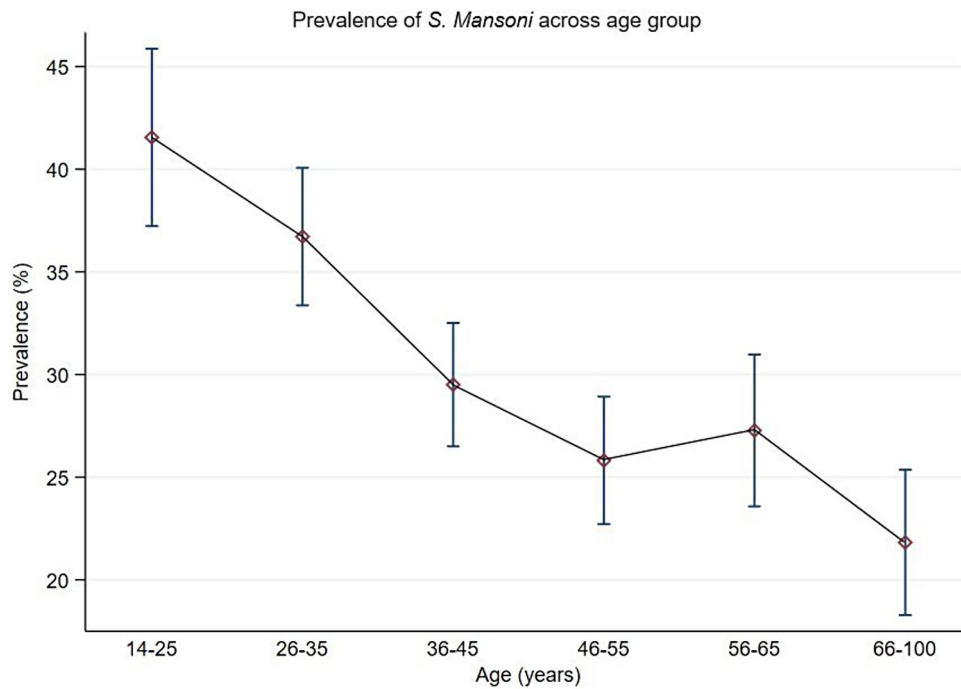


Fig. 2 Prevalence of *Schistosoma mansoni* infection categorized by age based on Kato Katz technique among adult community members on Ukerewe district

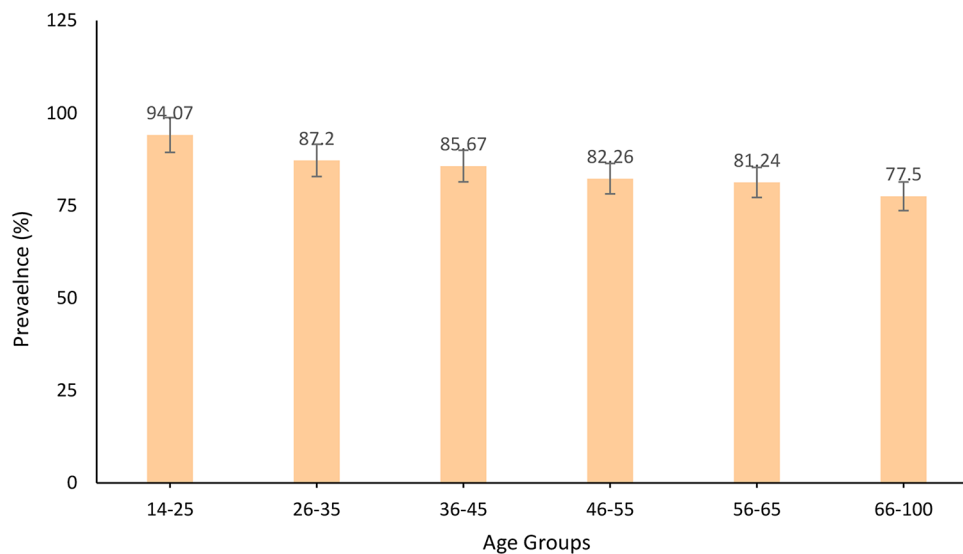


Fig. 3 Prevalence of *Schistosoma mansoni* infection categorized by age and sex based on Point-of-Care Circulating Cathodic Antigen Test among adult community members on Ukerewe district

categories, 53.9%, 32.4% and 13.7% had light, moderate and heavy intensity of infection, respectively. No significant difference was observed in all infection categories ($\chi^2(2)=6.2142, P=0.045$) in relation to sex. The youngest age groups 18–25 years and 26–35 years had a higher proportion of people with heavy infection intensity (19% and 16.4%) than other age groups ($\chi^2(10)=21;2487, P=0.02$).

A limited number of POC-CCA tests were available, both stool and urine samples could therefore only be analysed from a random sample of participants ($n=2,945$). Overall, the prevalence of *Schistosoma* based on the POC-CCA test was 84.7% (95%CI:83.3–85.9%), with no significant difference between male and female participants (84.9% versus 84.4%, $\chi^2(1)=0.1387, P=0.7$). The youngest age groups (Figs. 3), 18–26 years (94.1%) and 26–35 years (87.2%) had the highest prevalences

compared to the oldest age groups ($\chi^2(5)=48.1767$, $P=0.001$). Of the 2494/2945 (84.7%) who had POC-CCA positive results, 803/2,995 (27.3%) had *S. mansoni* egg positive slides.

Based on KK-technique, factors associated with *S. mansoni* infections were being male (aPR=1.2,95%CI:1.0-1.1, $P=0.001$), age groups, 18–25 years (aPR=1.2,95%CI:1.1–1.3, $P<0.001$), 26–35 years (aPR=1.2,95%CI:1.1–1.2, $P<0.001$), 36–45 years (aPR=1.1,95%CI:1.0-1.1, $P<0.001$), 46–55 years (aPR=1.1,95%CI:1.0-1.1, $P<0.04$) and 56–65 years (aPR=1.1,95%CI:1.0-1.1, $P<0.03$), reported blood in stool (aPR=1.1,95%CI:1.0-1.1, $P<0.002$), no history of PZQ use (aPR=1.1,95%CI:1.0-1.1, $P<0.001$) and village of residence. Table 3 present risk factors associated with *S. mansoni* infection among the adult population.

Co-infection of *S. mansoni* with Hepatitis C, B and HIV-1

A total of 3,939 participants had complete results for hepatitis C antibodies, 3, 933 for hepatitis B surface antigen and 3,895 for HIV-1 antibodies. Overall, the prevalences of hepatitis C antibodies, HIV and hepatitis B surface antigen were 0.4%, 2.2% and 4.7%. In general, 0.3%, 2.2% and 5.4% of the participants were co-infected with hepatitis C and *S. mansoni*, HIV and *S. mansoni* and hepatitis B and *S. mansoni*. A possible triple infection with hepatitis B, C and *S. mansoni* infections was observed in one participant whereas triple infection with HIV, hepatitis B and *S. mansoni* infection was observed in two participants.

Schistosoma mansoni related hepatosplenic morbidities and their associated factors

Complete data for abdominal ultrasound data were available for 3,630 participants from 20 villages.

Periportal fibrosis and its associated factors

A total of 3,447 participants had complete data for hepatosplenomegaly. The overall prevalence of periportal fibrosis (PPF) was 48.1% (1,657/3,447,95%CI:64.4–49.7%) and 47.3% of them had a detectable active *S. mansoni* infection as indicated by *S. mansoni* egg positive slides. Male participants had a higher prevalence of PPF than female participants (55.8%,95%CI:56.1–60.7% versus 37.3%,95%CI:34.9–39.6%, $\chi^2(1)=154.0461$, $P=0.001$). The older age group (≥ 36 years) had the highest prevalence of PPF compared with the youngest age group < 36 years ($\chi^2(2)=21.4302$, $P=0.001$). For all the participants who had PPF, had liver image pattern C and Cb (15.2%), D (7.8%), E and F (2.9%).

Factors associated with PPF are presented in Table 4. At univariate analysis, increased liver image pattern was associated with increased risk of having PPF. At multivariable analysis, being male (aPR=1.2,

95%CI:1.1–1.2, $P<0.001$), older age groups, 36–45 years (aPR=1.2,95%CI:1.0-1.2, $P<0.02$), 46–55 years (aPR=1.1,95%CI:1.1–1.2, $P<0.001$), 56–65 years (aPR=1.2,95%CI:1.0-1.1, $P<0.02$) and ≥ 66 years (aPR=1.2,95%CI:1.1–1.2, $P<0.001$), reported ever vomited blood (aPR=1.1,95%CI:1.0-1.3, $P<0.03$), reported abdominal distension (aPR=1.1,95%CI:1.1–1.3, $P<0.001$) and villages of residences were associated with increased risk of having PPF.

Prevalence of splenomegaly

The overall prevalence of splenomegaly was 40.5% (95%CI: 38.8–42.1%) with significant difference observed between sex. Male participants had a higher prevalence than female participants (45.7% versus 35.1%, $\chi^2(1)=41.9573$, $P=0.001$). There was no significant difference in prevalence of splenomegaly between age groups ($\chi^2(5)=1.7894$, $P=0.9$). Of those with splenomegaly, 57.9% ($n=851$) and 42.1% ($n=618$) had enlarged and severely enlarged spleens.

Left liver lobe hepatomegaly and its associated factors

The overall prevalence of hepatomegaly (enlargement of the left liver lobe) was 66.2% (95%CI:4.6–67.7%). Female participants recorded a higher prevalence of hepatomegaly than male participants (70.4% versus 62.1%, $\chi^2(4)=28.2256$, $P=0.001$). By age groups, the older age groups ≥ 36 years recorded a higher prevalence of hepatomegaly than the youngest age groups ($\chi^2(5)=28.6018$, $P=0.001$). Of those who had hepatomegaly, 66.8% and 33.2% had enlarged and severely enlarged left liver.

Table 5 illustrates factors which remained independently associated with left liver lobe hepatomegaly were being female (aPR=1.1,95%CI:1.0-1.1, $P<0.001$), age groups, reported history of non-use of praziquantel drug (aPR=1.1,95%CI:1.0-1.1, $P<0.002$), village of residences and liver image pattern (C-D).

Portal vein dilatation and its associated factors

The overall prevalence of portal vein dilatation (PVD) was 67.7% (95%CI: 66.2–69.2%). Male participants had a higher prevalence of PVD than female participants (72.6% versus 62.7%, $\chi^2(1)=40.2610$, $P=0.001$) (Fig. 4). In relation to age groups, the older age groups (> 40 years), had the highest prevalence of PVD and the prevalence was observed to increase with increasing age ($\chi^2(5)=28.4994$, $P=0.001$) (Fig. 5). Of the participants with PVD, 51.2% and 48.8% had a dilated and severely dilated portal vein.

Factors associated with portal vein dilatation included age groups, 46–55 years (aPR=1.1,95%CI:1.0-1.1, $P<0.03$), 56–65 years (aPR=1.1, 95%CI:1.0-1.1, $P<0.02$) and (aPR=1.1,95%CI:1.0-1.2, $P<0.001$), as well as liver image pattern C (aPR=1.5,95%CI:1.4–1.5, $P<0.001$),

Table 3 Factors associated with *S. mansoni* infections among adult individuals in 20 selected villages of Ukerewe district

| Variable | PR | 95%CI | P-values | aPR | 95%CI | P-values |
|---|-----|---------|----------|-----|---------|----------|
| Sex | | | | | | |
| Female | 1 | | | 1 | | |
| Male | 1.0 | 1.0–1.1 | 0.001 | 1.2 | 1.0–1.1 | 0.001 |
| Age groups | | | | | | |
| 14–25 | 1.2 | 1.2–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| 26–35 | 1.2 | 1.1–1.2 | 0.001 | 1.2 | 1.1–1.2 | 0.001 |
| 36–45 | 1.1 | 1.0–1.3 | 0.002 | 1.1 | 1.0–1.1 | 0.001 |
| 46–55 | 1.0 | 0.9–1.1 | 0.1 | 1.1 | 1.0–1.1 | 0.04 |
| 56–65 | 1.1 | 1.0–1.1 | 0.04 | 1.1 | 1.0–1.1 | 0.03 |
| ≥ 66 | 1 | | | | | |
| Reported diarrhoea in the past two weeks | | | | | | |
| No | 1 | | | 1 | | |
| yes | 0.9 | 0.9–1.1 | 0.2 | 0.9 | 0.9–1.1 | 0.04 |
| Reported blood in faeces | | | | | | |
| No | 1 | | | 1 | | |
| Yes | 1.1 | 1.0–1.1 | 0.001 | 1.1 | 1.0–1.1 | 0.002 |
| Reported abdominal pain | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.9–1.0 | 0.004 | 0.9 | 0.9–1.0 | 0.03 |
| Reported history of blood vomiting | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.9–1.1 | 0.3 | --- | --- | --- |
| Currently vomiting blood | | | | | | |
| No | 1 | | | | | |
| Yes | 0.8 | 0.7–1.1 | 0.3 | --- | --- | --- |
| History of abdominal distension | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.9–1.0 | 0.4 | --- | --- | --- |
| History of praziquantel use | | | | | | |
| No | 1.1 | 1.0–1.1 | 0.001 | 1.1 | 1.0–1.1 | 0.001 |
| Yes | 1 | | | 1 | | |
| Hepatitis C | | | | | | |
| No | 1 | | | | | |
| Yes | 0.8 | 0.7–1.1 | 0.3 | --- | --- | --- |
| Hepatitis B | | | | | | |
| No | 1 | | | | | |
| Yes | 1.1 | 0.9–1.1 | 0.3 | --- | --- | --- |
| HIV infection | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.8–1.0 | 0.3 | --- | --- | --- |
| Village of residence | | | | | | |
| Nebuye | 1 | | | 1 | | |
| Bugorola | 1.1 | 0.9–1.1 | 0.2 | 1.0 | 0.9–1.1 | 0.3 |
| Bugula | 1.2 | 1.1–1.3 | 0.01 | 1.2 | 1.1–1.3 | 0.01 |
| Bukiko | 1.1 | 1.0–1.2 | 0.03 | 1.1 | 1.0–1.2 | 0.02 |
| Bukindo | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Bukondo | 1.0 | 0.9–1.1 | 0.2 | 1.1 | 0.9–1.2 | 0.1 |
| Bukungu | 1.4 | 1.2–1.5 | 0.001 | 1.4 | 1.2–1.5 | 0.001 |
| Busumba | 1.1 | 1.0–1.2 | 0.03 | 1.1 | 1.0–1.2 | 0.04 |
| Bwisya | 1.1 | 0.9–1.2 | 0.06 | 1.1 | 1.0–1.2 | 0.04 |
| Chibasi | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Chifule | 1.3 | 1.2–1.5 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |
| Galu | 1.1 | 1.0–1.3 | 0.003 | 1.1 | 1.1–1.3 | 0.002 |

Table 3 (continued)

| Variable | PR | 95%CI | P-values | aPR | 95%CI | P-values |
|------------|-----|---------|----------|-----|---------|----------|
| Kamea | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.002 |
| Kome | 1.3 | 1.2–1.4 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |
| Kweru | 1.3 | 1.2–1.4 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |
| Muriti | 1.1 | 0.9–1.2 | 0.1 | 1.1 | 0.9–1.2 | 0.1 |
| Murutilima | 1.1 | 1.1–1.2 | 0.1 | 1.1 | 0.9–1.2 | 0.2 |
| Musozi | 1.3 | 1.2–1.4 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |
| Nyang'ombe | 1.2 | 0.9–1.2 | 0.2 | 1.1 | 0.9–1.2 | 0.2 |
| Nyamanga | 1.3 | 1.2–1.4 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |

pattern D (aPR=1.4,95%CI:1.4–1.5, $P<0.001$), pattern E (aPR=1.6,95%CI:1.5–1.7, $P<0.001$) and pattern F (aPR=1.6,95%CI:1.5–1.8, $P<0.001$). Other factors are shown in Table 6.

Prevalence of both splenomegaly, hepatomegaly, and portal vein dilatation

A proportion of participants who had hepatomegaly (66.2%) and splenomegaly (40.5%) had an enlargement of both organs (hepatosplenomegaly). A total of 46.6% of those who had enlargement of either the liver or spleen had hepatosplenomegaly. In relation to the portal vein dilatation, a total of 50.9% of those who had splenomegaly, had also a portal vein dilatation. Conversely, 71.9% of those who had hepatomegaly had also a portal vein dilatation. Lastly, 56% of the participants who had hepatosplenomegaly, had portal vein dilatation. Figures 6 and 7 summarizes the prevalence of hepatosplenic morbidities categorised by age and sex among the adult participants on Ukerewe island.

Other ultrasound detectable morbidities related to *S. mansoni* infection

Other detectable morbidities included gall bladder wall thickening (≥ 4 mm) with a prevalence of 40.4% (95%CI:38.5–42.3%). The prevalence was higher in males than in females ($\chi^2(1)=98.8089$, $P<0.001$), as well as in the older age groups (≥ 26 years, $\geq 40\%$). Ascites was found in 1.7% and collateral veins in 18.3% of the study participants.

Discussion

The result of the current study confirm that Ukerewe District remains endemic for *S. mansoni* infection and that its prevalence varies depending on the age, gender, and geographical locations of the study villages. Ultrasound detectable *S. mansoni* related hepatosplenic morbidities characterised by hepatomegaly of the left liver lobe, splenomegaly, hepatosplenomegaly, ascites, collateral veins, gall bladder wall thickening and portal vein dilatation were common in the adult population involved in the current study. Hepatitis B, C and HIV have been found to occur in the same population which resulting

into a proportion of adult individuals being co-infected with *S. mansoni* and these viral diseases simultaneously. We identified several of demographic, clinical, and epidemiological factors that partially explain the occurrence of *S. mansoni* related hepatosplenic morbidities in the current study population.

The results of the current study confirm the results of the previous studies on Ukerewe island, which suggested that the area is endemic only for *S. mansoni* infection [6, 10] and the affected groups have high infection intensity. Previous studies in pre- and school-aged children [6, 7, 10] and adults [4, 9] found a high prevalence and intensity of infections on Ukerewe island, suggesting that these areas are hotspots for *S. mansoni* infections for a long time. The current study found a lower prevalence of *S. mansoni* infection based on the KK-technique than 78% [4] and 86.3% [9] reported in previous studies in adults on the same island. The recorded lower prevalence observed in the current study may be explained by the changes in the epidemiology of *S. mansoni* infection in northwestern Tanzania, where accumulative decline in the prevalence of the disease was observed due to repeated school-based mass drug administration in this area [2, 3, 17]. Despite this positive observation, there are still areas of high *S. mansoni* prevalence and infection intensity in communities living along the Lake Victoria shoreline, particularly on Ukerewe island [3]. This shows that an intervention that focuses only on a part of the population, mainly school-age children, and excludes the other population groups, namely adults and preschool-age children, from control efforts will not lead to disease control. An integrative MDA will be used in the current study population, which include all community members. While the Tanzanian government offers annual MDA to the SAC through the primary school platform, the current project activities are focusing on adult community members. They will receive annual MDA within their villages and the plan is to provide five rounds of treatment during the five-year project period. Previous studies have shown that a combination of school-and-community based MDA had a major impact on the disease prevalence and infection intensity [28, 29].

Table 4 Risk factors associated with periportal fibrosis among adult participants on Ukerewe Island

| Variable | cPR | 95%CI | P-value | aPR | 95%CI | P-value |
|---|-----|---------|---------|-----|---------|---------|
| Sex | | | | | | |
| Female | 1 | | | | | |
| Male | 1.2 | 1.2–1.3 | 0.001 | 1.2 | 1.1–1.2 | 0.001 |
| Age groups in years | | | | | | |
| 15–25 | 1 | | | 1 | | |
| 26–35 | 1.1 | 1.0–1.2 | 0.002 | 1.1 | 1.0–1.1 | 0.2 |
| 36–45 | 1.1 | 1.1–1.2 | 0.001 | 1.2 | 1.1–1.2 | 0.02 |
| 46–55 | 1.1 | 1.1–1.2 | 0.001 | 1.1 | 1.1–1.2 | 0.001 |
| 56–65 | 1.1 | 1.0–1.2 | 0.002 | 1.2 | 1.0–1.1 | 0.02 |
| ≥ 66 | 1.1 | 1.1–1.2 | 0.001 | 1.2 | 1.1–1.2 | 0.001 |
| Reported ever vomited blood | | | | | | |
| No | 1 | | | | | |
| Yes | 1.2 | 1.1–1.4 | 0.004 | 1.1 | 1.0–1.3 | 0.03 |
| HIV serostatus | | | | | | |
| Negative | 1 | | | | | |
| Positive | 0.9 | 0.8–1.1 | 0.8 | -- | -- | -- |
| Hepatitis C | | | | | | |
| Negative | 1 | | | | | |
| Positive | 1.1 | 0.8–1.3 | 0.6 | -- | -- | -- |
| Hepatitis B | | | | | | |
| Negative | 1 | | | | | |
| Positive | 0.9 | 0.9–1.1 | 0.5 | -- | -- | -- |
| Reported current vomiting blood | | | | | | |
| No | 1 | | | | | |
| Yes | 1.2 | 0.9–1.6 | 0.1 | - | - | - |
| Reported abdominal distension | | | | | | |
| No | 1 | | | | | |
| Yes | 1.2 | 1.1–1.3 | 0.001 | 1.9 | 1.1–1.3 | 0.001 |
| Liver image pattern | | | | | | |
| A | 1 | | | | | |
| B | 1.2 | 1.0–1.3 | 0.01 | -- | -- | -- |
| C (Cb) | 1.5 | 1.4–1.6 | 0.001 | -- | -- | -- |
| D(Dc, Db, DcB) | 1.5 | 1.4–1.5 | 0.001 | -- | -- | -- |
| E(Ec, Eb, Ecb) | 1.7 | 1.6–1.8 | 0.001 | -- | -- | -- |
| F (Fc) | 1.7 | 1.6–1.8 | 0.001 | -- | -- | -- |
| Intensity of <i>S. mansoni</i> infection | | | | | | |
| Low | 1 | | | | | |
| Moderate | 1.0 | 0.9–1.1 | 0.2 | -- | -- | -- |
| Heavy | 1.0 | 0.9–1.1 | 0.5 | -- | -- | -- |
| <i>S. mansoni</i> infection | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.9–1.0 | 0.5 | -- | -- | -- |
| Villages of residence | | | | | | |
| Muriti | 1 | | | | | |
| Bugorola | 1.4 | 1.3–1.6 | 0.001 | 1.4 | 1.3–1.5 | 0.001 |
| Bugula | 1.1 | 0.9–1.2 | 0.2 | 1.1 | 0.9–1.3 | 0.2 |
| Bukiko | 1.5 | 1.3–1.6 | 0.001 | 1.5 | 1.3–1.6 | 0.001 |
| Bukindo | 1.3 | 1.2–1.5 | 0.001 | 1.3 | 1.2–1.5 | 0.001 |
| Bukondo | 1.1 | 0.9–1.2 | 0.1 | 1.0 | 0.9–1.2 | 0.4 |
| Bukungu | 1.1 | 1.0–1.3 | 0.01 | 1.1 | 0.9–1.2 | 0.1 |
| Busumba | 1.5 | 1.4–1.7 | 0.001 | 1.5 | 1.3–1.7 | 0.001 |
| Bwisya | 1.5 | 1.3–1.7 | 0.001 | 1.5 | 1.3–1.7 | 0.001 |
| Chibasi | 1.3 | 1.2–1.5 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |

Table 4 (continued)

| Variable | cPR | 95%CI | P-value | aPR | 95%CI | P-value |
|------------|-----|---------|---------|-----|---------|---------|
| Chifule | 1.5 | 1.3–1.7 | 0.001 | 1.5 | 1.3–1.6 | 0.001 |
| Gallu | 1.1 | 0.9–1.2 | 0.2 | 1.1 | 0.9–1.2 | 0.2 |
| Kamea | 1.4 | 1.2–1.4 | 0.001 | 1.4 | 1.2–1.5 | 0.001 |
| Kome | 1.3 | 1.2–1.4 | 0.001 | 1.3 | 1.1–1.4 | 0.001 |
| Kweru | 1.1 | 1.0–1.2 | 0.04 | 1.1 | 0.9–1.2 | 0.1 |
| Murutilima | 1.1 | 1.3–1.7 | 0.001 | 1.5 | 1.3–1.6 | 0.001 |
| Musozi | 1.2 | 1.1–1.3 | 0.002 | 1.2 | 1.0–1.3 | 0.01 |
| Nebuye | 1.1 | 1.0–1.3 | 0.01 | 1.1 | 1.0–1.3 | 0.02 |
| Nyamanga | 1.3 | 1.1–1.4 | 0.001 | 1.3 | 1.1–1.4 | 0.001 |
| Nyang'ombe | 1.3 | 1.2–1.5 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |

In the current study population, a significant difference in sensitivity between the KK technique and the POC-CCA test in detecting *S. mansoni* cases was observed in the subpopulation of the participants screened with both tests. The POC-CCA test detected more cases of the disease in the adult population compared to the KK technique in the adult population. Studies along the Lake Victoria have shown similar results in the adult population [7, 21, 30, 31] and in pre-school children and school aged children [7, 30]. The POC-CCA test remain highly sensitive than the KK technique and is recommended for the rapid assessment of schistosomiasis in endemic areas [32–35]. However, the test has several limitations including reduced specificity, inability to quantify eggs and is still an expensive test [32]. As intensity of infection continue to be an indicator for measuring the elimination of schistosomiasis as a public health challenge [19] and the decline in intensities of infection due to repeated rounds of MDA impacting the performance of the KK technique [36]. It will be important to continue combining the multiple diagnostic as countries approach the elimination phase.

In the current study, some of the participants had hepatitis B, C, and HIV infections that resulted in co-infection with *S. mansoni* parasite. Due to their different pathogenic mechanisms, these co-infections can lead to an exacerbation of hepatosplenic pathologies, which in turn can result in liver fibrosis [37]. We have found similar results in our previous studies in the adult population in villages around the Lake Victoria shoreline [38]. However, the co-infections of these viral infections with *S. mansoni* parasite occurs at lower rate than in other endemic areas of Africa [39], particularly in Egypt, where the prevalence of HCV is high [39, 40]. It is worth noting that in the present study, the prevalence of the hepatitis viruses HCV and hepatitis B is low, therefore, the prevalence of co-infection patterns is also low. Regarding HIV infections, the fishing villages on the southern shoreline of Lake Victoria have lower prevalence of HIV than comparable villages in western Kenya and Uganda [41]. A number of behavioural, cultural, and socio-economic

conditions may partially explain the observed differences. Overall, our results demonstrate the need of integrating health services rather than implement parallel programs when targeting communities living in tropical environments where multiple diseases transmission occurs. This can reduce the costs in the program, as in the present survey, which included the district's HIV program and offered HIV screening services.

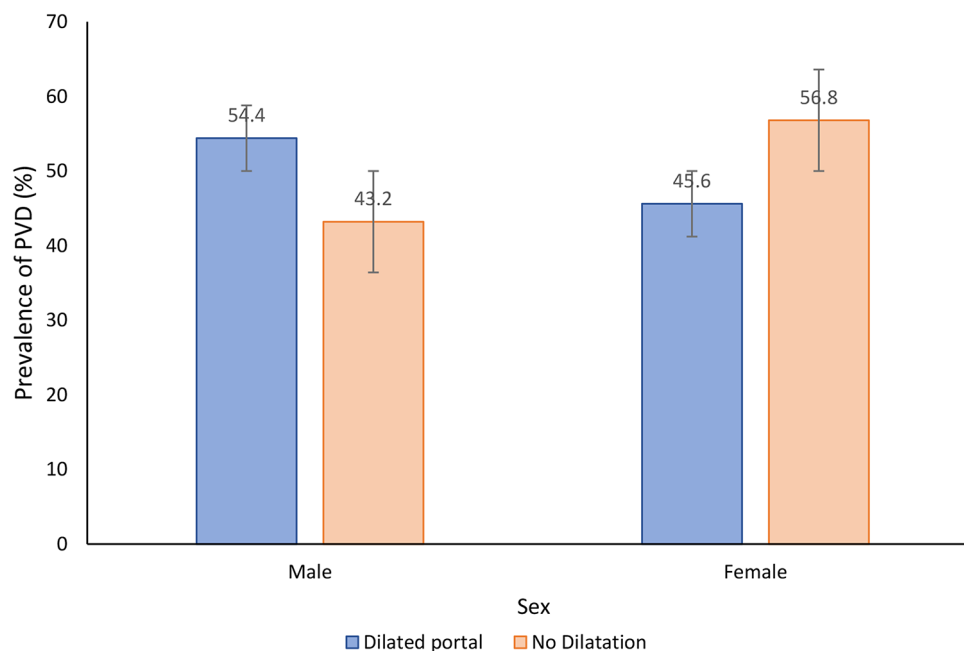
It has been repeatedly reported that communities living on the shoreline of Lake Victoria and on its islands where *S. mansoni* is highly endemic have the highest burden of hepatosplenic disease. The pathology results mainly from Th-2 immune responses due to *S. mansoni* eggs shed at various sites of the body [11]. The results of the current study and previous studies [9, 18] on Ukerewe island have found a high prevalence of hepatosplenic disease in the adult population. Previous studies reported a prevalence of 35% [18] and 80% of hepatomegaly and splenomegaly [9]. Similarly, a high prevalence of PPF was observed among adult at Kome (42%) and Msozi (42%) villages of Ukerewe island [18]. In the current study, similar trends were noted in the high prevalence of hepatosplenic diseases characterized by splenomegaly (40.5%), PPF (48.1%), left hepatic lobe hepatomegaly (66.2%) and portal vein dilatation (67.7%). We observed that the prevalence of these chronic *S. mansoni* chronic morbidities increases with age and that they occur frequently after the age of 40 years. Overall, these results suggest that *S. mansoni* infection reaches its chronic morbidities with increasing age in this population. Our previous study of pre-school children from the same area shows that, in this population, *S. mansoni* infection begins at two years of age [7] and we have recorded visible signs of hepatosplenic diseases by ultrasound in children aged 5 to 6 years [21]. In the hospital setting in northwestern Tanzania, our studies have found a high prevalence of hepatosplenic disorders, characterized by upper gastrointestinal bleeding. In adults with oesophageal varices, 50% of cases are associated with *S. mansoni* and almost 25% of these patients die within six months [14]. These results suggest that the adult population suffers from *S.*

Table 5 Factors associated with left liver lobe among adult participants in 20 selected villages of Ukerewe district

| Variable | cPR | 95%CI | P-value | aPR | 95%CI | P-value |
|--|-----|---------|---------|-----|---------|---------|
| Sex | | | | | | |
| Female | 1.1 | 1.0–1.1 | 0.001 | 1.1 | 1.0–1.1 | 0.001 |
| Male | 1 | | | 1 | | |
| Age groups (in years) | | | | | | |
| 14–25 | 1 | | | 1 | | |
| 26–35 | 0.9 | 0.9–1.0 | 0.6 | 0.9 | 0.9–1.0 | 0.6 |
| 36–45 | 1.1 | 1.0–1.1 | 0.02 | 1.1 | 1.0–1.1 | 0.02 |
| 46–55 | 1.1 | 0.9–1.1 | 0.1 | 1.1 | 1.0–1.1 | 0.02 |
| 56–65 | 1.1 | 1.1–1.2 | 0.001 | 1.1 | 1.1–1.2 | 0.01 |
| ≥ 66 | 1.1 | 1.0–1.1 | 0.04 | 1.1 | 0.9–1.1 | 0.05 |
| History of blood vomiting | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.8–1.1 | 0.3 | --- | --- | --- |
| History of blood in stool | | | | | | |
| No | 1 | | | | | |
| Yes | 1.0 | 0.9–1.1 | 0.3 | --- | --- | --- |
| History abdominal pain | | | | | | |
| No | 1 | | | | | |
| Yes | 1.0 | 1.0–1.1 | 0.4 | --- | --- | --- |
| History of abdominal distension | | | | | | |
| No | 1 | | | 1 | | |
| Yes | 1.1 | 0.9–1.1 | 0.2 | 1.0 | 0.9–1.1 | 0.8 |
| History of praziquantel treatment | | | | | | |
| No | 1.0 | 1.0–1.1 | 0.01 | 1.1 | 1.0–1.1 | 0.002 |
| Yes | 1 | | | | | |
| HIV serostatus | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.8–1.2 | 0.9 | --- | --- | --- |
| Hepatitis C | | | | | | |
| No | 1 | | | | | |
| Yes | 1.1 | 0.8–1.4 | 0.5 | --- | --- | --- |
| Hepatitis B | | | | | | |
| No | 1 | | | | | |
| Yes | 1.0 | 0.9–1.1 | 0.9 | --- | --- | --- |
| S. mansoni infection status | | | | | | |
| No | 1 | | | 1 | | |
| Yes | 1.0 | 0.9–1.1 | 0.2 | 1.0 | 0.9–1.1 | 0.2 |
| Intensity of S. mansoni infection | | | | | | |
| Low | 1 | | | | | |
| Moderate | 1.0 | 0.9–1.1 | 0.7 | --- | --- | --- |
| Heavy | 1.1 | 0.9–1.1 | 0.2 | --- | --- | --- |
| Village of residence | | | | | | |
| Muriti | 1 | | | 1 | | |
| Bugorola | 1.1 | 0.9–1.2 | 0.08 | 1.1 | 0.9–1.2 | 0.2 |
| Bugula | 1.0 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Bukiko | 1.4 | 1.3–1.5 | 0.001 | 1.4 | 1.2–1.5 | 0.001 |
| Bukindo | 1.3 | 1.2–1.4 | 0.001 | 1.3 | 1.1–1.4 | 0.001 |
| Bukondo | 1.3 | 1.1–1.4 | 0.001 | 1.3 | 1.1–1.4 | 0.001 |
| Bukungu | 1.3 | 1.2–1.4 | 0.001 | 1.3 | 1.1–1.4 | 0.001 |
| Busumba | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Bwisya | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Chibasi | 1.3 | 1.1–1.4 | 0.001 | 1.3 | 1.1–1.4 | 0.001 |
| Chifule | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.002 |

Table 5 (continued)

| Variable | cPR | 95%CI | P-value | aPR | 95%CI | P-value |
|----------------------------|-----|---------|---------|-----|---------|---------|
| Gallu | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.003 |
| Kamea | 1.3 | 1.3–1.6 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |
| Kome | 1.4 | 1.3–1.6 | 0.001 | 1.3 | 1.2–1.5 | 0.001 |
| Kweru | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Murutilima | 1.2 | 1.0–1.3 | 0.004 | 1.2 | 1.0–1.3 | 0.01 |
| Musozi | 1.4 | 1.3–1.5 | 0.001 | 1.4 | 1.2–1.5 | 0.001 |
| Nebuye | 1.2 | 1.1–1.4 | 0.001 | 1.2 | 1.1–1.4 | 0.001 |
| Nyamanga | 1.4 | 1.3–1.6 | 0.001 | 1.3 | 1.2–1.5 | 0.001 |
| Nyang'ombe | 1.2 | 1.1–1.4 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Liver image pattern | | | | | | |
| A and B | 1 | | | 1 | | |
| C | 1.1 | 1.0–1.1 | 0.02 | 1.1 | 1.0–1.1 | 0.03 |
| D | 1.0 | 1.0–1.1 | 0.04 | 1.1 | 1.0–1.1 | 0.002 |
| E | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| F | 1.1 | 1.0–1.2 | 0.02 | 1.1 | 1.0–1.2 | 0.01 |

**Fig. 4** Prevalence of portal vein dilatation categorised by sex among adult population on Ukerewe island, north-western Tanzania

mansoni infection for a long time without treatment and eventually develops life threatening morbidities. Overall, our results highlight the need to involve the entire community in schistosomiasis control programs along the Lake Victoria basin. To reduce and control the transmission of schistosomiasis and its associated hepatosplenic morbidities, it is strongly recommended that annual or biannual community-based MDA be offered to the exposed population for an extended period of time to reduce the transmission potential of the parasite and control the hepatosplenic morbidities. Treatment of hepatosplenic disease in adult patients (on a case-by-case basis) in healthcare facilities should be part of this effort. Activities of the current project include improving the

detection and treatment of severe hepatosplenic diseases in primary healthcare facilities and transferring patients to the next level of healthcare when necessary to reduce severe morbidity and mortality. In the district hospital, the project is building the capacity of healthcare workers to deal with these serious cases through endoscopic procedures established within this framework. Currently, the country is working to achieve the 2030 vision of controlling and eliminating schistosomiasis as a public health challenge. In the elimination phase, the remaining work is mainly to treat the chronic hepatosplenic disease caused by *S. mansoni*, which may not resolve or may take time to resolve. Therefore, capacity building of

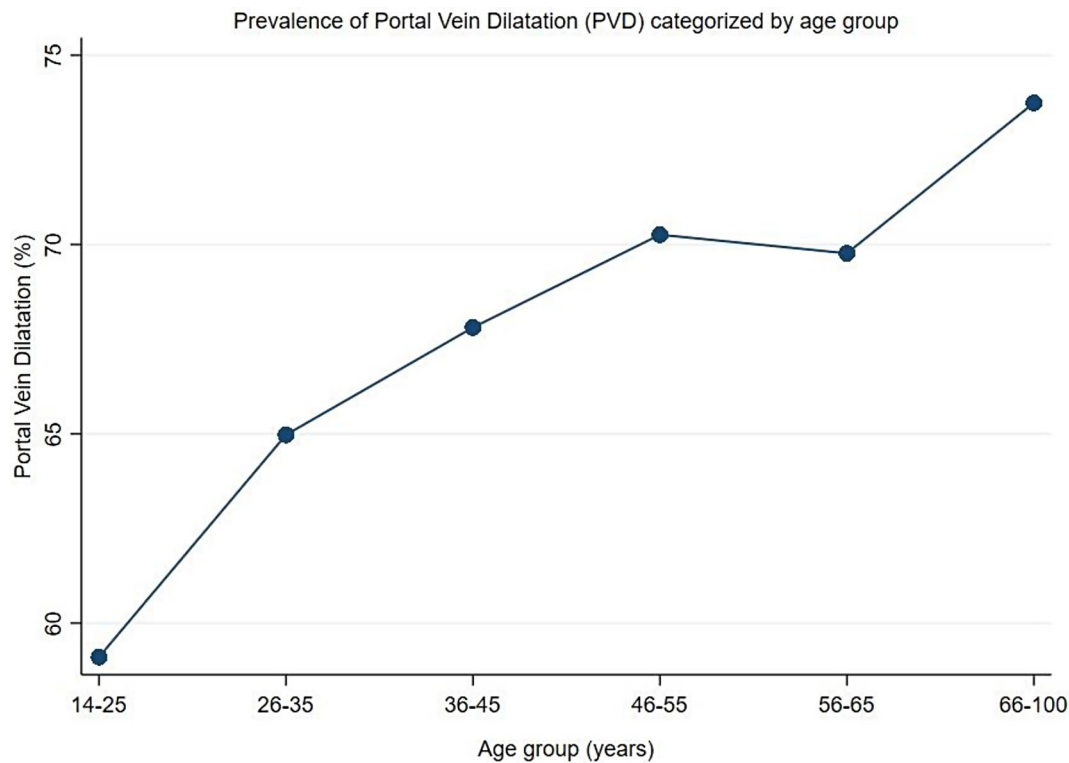


Fig. 5 Prevalence of Portal Vein Dilatation categorized by age groups, the prevalence of PVD increased with increase in age

the primary healthcare system is strongly recommended, especially in endemic areas.

In *S. mansoni* endemic areas, various demographic, clinical and epidemiological factors have been described to explain the continuity of transmission of *S. mansoni* infections and the development of associated morbidities. In the current study setting, *S. mansoni* infection was observed to be associated with the youngest age groups, male gender, blood in stool, no use of praziquantel and place of residence. These demographic factors have been described previously to be associated with *S. mansoni* infection along the Lake Victoria areas [21, 42]. The current study participants reported enterocolitis symptoms and signs such as abdominal pain, diarrhoea, hematemesis, and bloody stools. This was similar to a previous study in the same setting [9]. We found that reported bloody stools continue to be independently associated with *S. mansoni* infection. From a pathogenic perspective, *S. mansoni* eggs that pass through the intestinal wall and are accompanied by inflammatory reactions can cause blood to seep into the intestinal lumen and be detectable in excreted stool. Due to rupture of oesophageal varices, digested blood maybe found in stool [43]. In tropical environmental, blood in stool can also be caused by other endemic intestinal infections such as intestinal amoebiasis [44] and other bacterial and viral infections that can cause dysentery. Due to their geographical location on the Lake Victoria shorelines, the

population living in these villages frequently contacts lake water during fishing, recreational activities, farming, carrying household chores and farming along the shoreline. Almost, all study villages remain at risk of *S. mansoni* infection. Epidemiological studies have shown that, *S. mansoni* infection is associated with village of residence, age, occupation, specifically fishing, being male, and hepatosplenic morbidities [21, 42].

For hepatosplenic morbidities (left liver lobe hepatomegaly, PPF and PVD), sex, older age groups, non-use of praziquantel drugs, reported ever having vomited blood, abdominal distension, liver image pattern (C-F) and village of residences were common explanatory factors that remained independently associated with these morbidities. The association of these morbidities with older age groups clearly shows that the current study population suffering from *S. mansoni* infection for a long time without treatment. The development of *S. mansoni* related morbidities is a function of intensity of infection, duration of infection exposure, and a genetical component of the host [11, 42]. These morbidities develop slowly, and the severe manifestations and life-threatening symptoms and signs occur in older age groups [43]. Some of these factors, such as the intensity of infection and duration of exposure can be modified through various interventions. For example, the delivery of PZQ drug through community MDA can be expanded to reach the entire population. This will help to reduce the transmission

Table 6 Factors associated with portal vein dilatation among adult participants from 20 villages of Ukerewe district, north-western Tanzania

| Variable | cPR | 95%CI | P-value | aPR | 95%CI | P-value |
|--|-----|---------|---------|-----|---------|---------|
| Sex | | | | | | |
| Female | 1 | | | 1 | | |
| Male | 1.1 | 1.0–1.1 | 0.001 | 1.0 | 0.9–1.0 | 0.2 |
| Age groups (in years) | | | | | | |
| 14–25 | 1 | | | 1 | | |
| 26–35 | 1.1 | 1.0–1.1 | 0.04 | 1.0 | 0.9–1.1 | 0.3 |
| 36–45 | 1.1 | 1.0–1.2 | 0.002 | 1.0 | 0.9–1.1 | 0.1 |
| 46–55 | 1.1 | 1.1–1.2 | 0.001 | 1.1 | 1.0–1.1 | 0.03 |
| 56–65 | 1.1 | 1.0–1.2 | 0.001 | 1.1 | 1.0–1.1 | 0.02 |
| ≥ 66 | 1.2 | 1.1–1.2 | 0.001 | 1.1 | 1.0–1.2 | 0.001 |
| History of blood vomiting | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.8–1.1 | 0.3 | --- | --- | --- |
| History of blood in stool | | | | | | |
| No | 1 | | | 1 | | |
| Yes | 1.2 | 1.0–1.3 | 0.01 | 1.1 | 0.9–1.2 | 0.2 |
| History abdominal pain | | | | | | |
| No | 1 | | | | | |
| Yes | 1.0 | 1.0–1.1 | 0.4 | --- | --- | --- |
| History of abdominal distension | | | | | | |
| No | 1 | | | 1 | | |
| Yes | 1.2 | 1.0–1.2 | 0.002 | 1.0 | 0.9–1.1 | 0.4 |
| History of praziquantel treatment | | | | | | |
| No | 1.0 | 1.0–1.1 | 0.01 | 1.1 | 1.0–1.1 | 0.002 |
| Yes | 1 | | | | | |
| HIV serostatus | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.8–1.1 | 0.8 | --- | --- | --- |
| Hepatitis C | | | | | | |
| No | 1 | | | | | |
| Yes | 1.1 | 0.8–1.3 | 0.6 | --- | --- | --- |
| Hepatitis B | | | | | | |
| No | 1 | | | | | |
| Yes | 0.9 | 0.9–1.1 | 0.5 | --- | --- | --- |
| S. mansoni infection status | | | | | | |
| No | 1 | | | | | |
| Yes | 1.0 | 0.9–1.0 | 0.4 | --- | --- | --- |
| Intensity of S. mansoni infection | | | | | | |
| Low | 1 | | | | | |
| Moderate | 1.0 | 0.9–1.1 | 0.4 | --- | --- | --- |
| Heavy | 0.9 | 0.9–1.1 | 0.9 | --- | --- | --- |
| Village of residence | | | | | | |
| Bugula | 1 | | | 1 | | |
| Bugorola | 1.3 | 1.2–1.5 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Muriti | 1.1 | 0.9–1.2 | 0.1 | 1.1 | 1.0–1.2 | 0.01 |
| Bukiko | 1.3 | 1.3–1.5 | 0.001 | 1.2 | 1.2–1.3 | 0.001 |
| Bukindo | 1.3 | 1.2–1.4 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Bukondo | 1.1 | 0.9–1.2 | 0.2 | 1.1 | 0.9–1.2 | 0.1 |
| Bukungu | 1.2 | 1.1–1.3 | 0.001 | 1.1 | 0.9–1.2 | 0.2 |
| Busumba | 1.4 | 1.2–1.5 | 0.001 | 1.1 | 1.1–1.3 | 0.001 |
| Bwisya | 1.4 | 1.2–1.5 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Chibasi | 1.2 | 1.1–1.4 | 0.001 | 1.1 | 1.0–1.2 | 0.01 |

Table 6 (continued)

| Variable | cPR | 95%CI | P-value | aPR | 95%CI | P-value |
|----------------------------|-----|---------|---------|-----|---------|---------|
| Chifule | 1.4 | 1.2–1.5 | 0.001 | 1.2 | 1.1–1.4 | 0.001 |
| Gallu | 1.0 | 0.9–1.1 | 0.9 | 1.0 | 0.9–1.1 | 0.9 |
| Kamea | 1.2 | 1.1–1.4 | 0.001 | 1.1 | 1.0–1.2 | 0.003 |
| Kome | 1.4 | 1.3–1.5 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Kweru | 1.1 | 1.0–1.2 | 0.003 | 1.1 | 1.0–1.2 | 0.002 |
| Murutilima | 1.4 | 1.2–1.5 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Musozi | 1.4 | 1.3–1.5 | 0.001 | 1.3 | 1.2–1.4 | 0.001 |
| Nebuye | 1.2 | 1.1–1.3 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Nyamanga | 1.3 | 1.2–1.4 | 0.001 | 1.1 | 1.0–1.2 | 0.02 |
| Nyang'ombe | 1.3 | 1.2–1.4 | 0.001 | 1.2 | 1.1–1.3 | 0.001 |
| Liver image pattern | | | | | | |
| A and B | 1 | | | 1 | | |
| C | 1.5 | 1.4–1.6 | 0.001 | 1.5 | 1.4–1.5 | 0.001 |
| D | 1.4 | 1.4–1.5 | 0.001 | 1.4 | 1.4–1.5 | 0.001 |
| E | 1.7 | 1.6–1.8 | 0.001 | 1.6 | 1.5–1.7 | 0.001 |
| F | 1.7 | 1.6–1.9 | 0.001 | 1.6 | 1.5–1.8 | 0.001 |

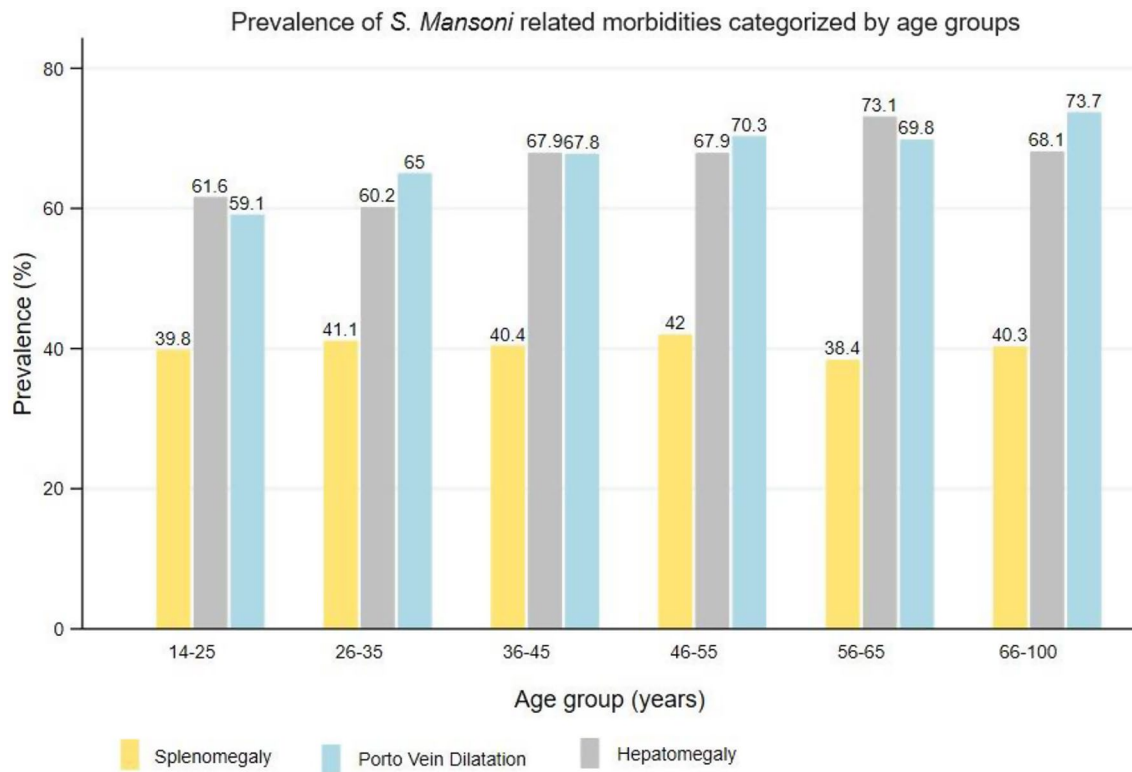


Fig. 6 Prevalence of *S. mansoni* related ultrasound detectable morbidities (splenomegaly, hepatomegaly, and portal vein dilatation) categorised by age groups

of *S. mansoni* infections and lead to regression of these hepatosplenic morbidities [28, 45]. In addition, providing water, sanitation and hygiene facilities to endemic communities can reduce human faecal pollution and human-water contact, thereby helping to facilitates effort to control the disease and push the country to the elimination phase. Public health education that focuses on understanding disease transmission routes and

compliance with prevention and control measures can increase target audience awareness, which in turn can lead to improved compliance with control measures such as MDA. These efforts should be accompanied by the search for alternative sources of income for these populations whose livelihood are highly dependent on fishing.

We acknowledged that the current study was not conducted without any limitations. The use of duplicate KK

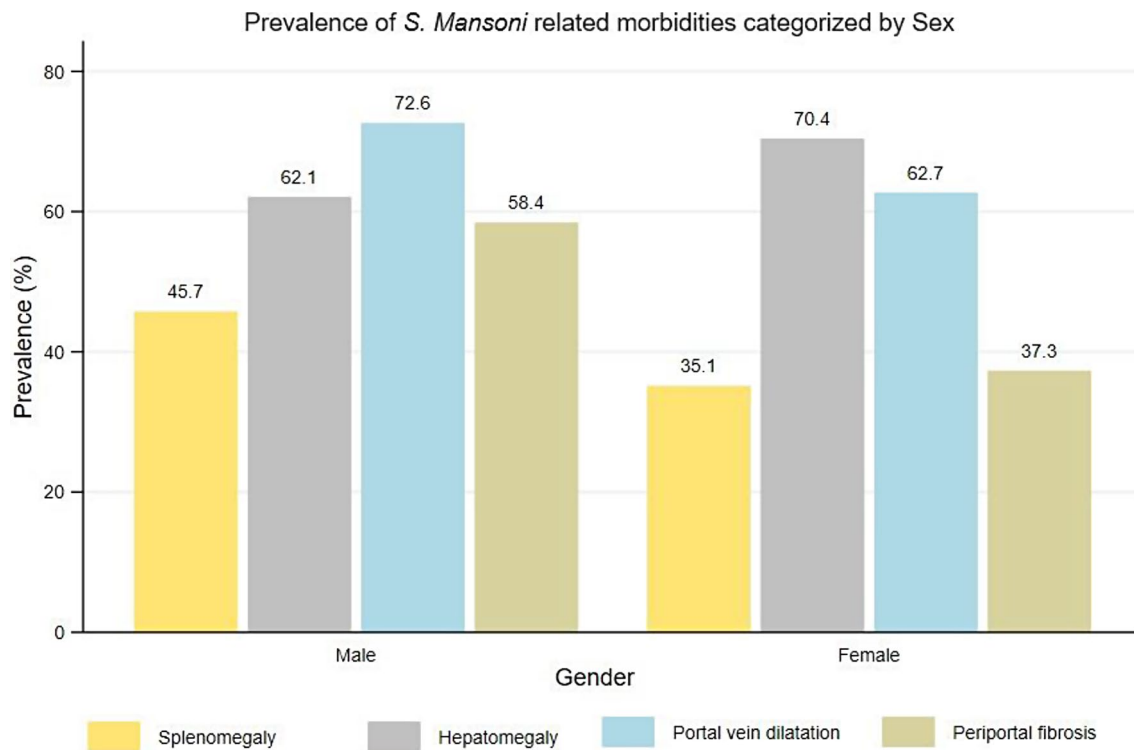


Fig. 7 Prevalence of *S. mansoni* infections and its related ultrasound detectable morbidities (splenomegaly, hepatomegaly, periportal fibrosis and portal vein dilatation) categorised by sex

thick smears for a single stool sample to screen for *S. mansoni* infection may have resulted in an underestimate of the true prevalence of the disease due to daily fluctuation in egg output of infected individuals [46]. To reduce the impact of this limitation, a point-of-care CCA test was used, which has higher sensitivity than the KK technique, but this test also has its own limitations. The cross-sectional nature of the study design, with limited ability to demonstrate the association of various explanatory factors with the outcome, may have contribute to the lack of association between some of the explanatory variables and the outcome of interest. Nevertheless, the current epidemiological study was one of the largest ultrasound surveys in an endemic foci of *S. mansoni* in northwestern Tanzania and has clearly demonstrated that the disease is still endemic and associated with life-threatening morbidities and therefore requires integrative measures. The current project activities close this gap.

Conclusion

The current study setting is endemic for *S. mansoni* infection and the population has a high prevalence of hepatosplenic morbidities associated with the disease, characterized by hepatomegaly of the liver lobe, splenomegaly, ascites, thickness of the gall bladder wall thickness, periportal fibrosis and portal vein dilatation. Several

demographic, clinical and epidemiological factors remain independently associated with *S. mansoni* infection and associated morbidities. These findings call for integrative intervention measures, starting with whole- community MDA, that covers all out-of-school going community members. The current project conducts annual MDA for high-risk communities living in 20 villages. However, all the villages on Ukerewe Island require community MDA interventions, which should be combined with other intervention measures such as WASH, public health education and capacity building of the health facilities to detect and treat advanced complications of hepatosplenic schistosomiasis.

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Author contributions

HDM, SK, CK, AM and AF fund acquisition and study design CEC, TB, ESM, GMK, MM and SM fieldwork coordination, data collection and data management HDM and TL data analysis HDM and CEC wrote the first draft of the manuscript SK, CK, AF and AM reviewed the manuscript HDM, CEC, SK and CK were responsible for the overall project administration.

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Data availability

The datasets used for analysing the presented data are available from the corresponding authors upon reasonable request. This is because the ethical approval process made no specific mention of making the data publicly available.

Declarations

Ethics approval and consent to participate

The ethical approval to implement the project activities including the current study was obtained from the National Ethical Review Committee board (NIMR/HQ/R.8a/Vol.IX/3590 and NIMR/HQ/R.8 C/Vol.I/1973) and further implementation permission were received from the Prime Minister office for local governments, Mwanza region administrative office and the Ukerewe district council, where the study is being implemented. Written Informed Consent to participate in the current study was obtained from all study participants. The informed consent form was written in Kiswahili language and each participant received a copy of written informed consent. For illiterate participants, the consent form was signed using thumbprint after they had received a clear oral description of the study objectives and the treatment options. The consent form described the objective, durations, risks, and benefits of participating in the study. Participation in the study was voluntary and participants had the right to continue or to withdraw their consent from the study at any time.

Consent for publication

Not applicable.

Competing interests

The authors declare no competing interests.

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