

### PUBLICATION REPORT

Line of the states

1775

9/94

SCHISTOSOMA MANSONI INFECTION 3 MONTHS AFTER PRAZIQUANTEL THERAPY AMONG FARMERS IN QALYUB, EGYPT

B¥

27.R. Aba-Elyazeed, N.S. Mansour, M. Habib and J.K. Podgore

U.S. NAVAL MEDICAL RESEARCH UNIT NO. 3 (CAIRO, ARAB REPUBLIC OF EGYPT) PSC 452, BOX 5000 FPO AE 09835-0007

DTIC QUALITY HALF SORED &

REPORT D	Form Approved OMB No. 0704-0188		
Particle on a story consider to software devices of a galaxies y order continuing the data meeted a scalastic of software trees including supported to a software of software 1204. Arbungton, VA 222	nlight the constituted to average 1 hour oils completing and reviewing the collection is for reducing this burden, to Washington 22 4102, and to the office of Management	per response, including the time fo o of information - Send comments re Headquarters Services, Directorale and Budget, Paperwork Reduction	r reviewing instructions, searching existing data sources, erarding this burden estimate or any other aspect of this for information Operations and Reports, 1215 Jefferson Project (0704-0188), Washington, DC 2050
1. AGENTY USE ONLY (Leave bla	nk) 2. REPORT DATE 1993	J. REPORT TYPE	AND DATES COVERED
4. HILL AND SUBTILE Schistosoma mansoni Infectio among Farmers in Qalyub, E	5. FUNDING NUMBERS PE- 64758A WU- 3M464758D849.EB		
6. AUTHOR(5) Abu-Elyazeed, R.R., Manson and Podgore, J.K.	ur, N.S., Habib, M.,		
7. PERFORMING ORGANIZATION NAME(S) AND ADDRESS(ES)			8. PERFORMING ORGANIZATION REPORT NUMBER
U.S. Naval Medical Research PSC 452, Box 5000 FPO AE 09835-0007	9/94		
9. SPONSORING/MONIFORING AGENCY NAME(S) AND ADDRESS(ES) Naval Medical Research and Development Command, National Naval Medical Center Building 1, Tower 12 Batharda, MD 20890 5044			10. SPONSORING/MONITORING AGENCY REPORT NUMBER
11. SUPPLEMENTARY NOTES			
Published in: J. Trop. Med.	, <u>2</u> (4):3-7, 1993; Acc. No. 1	175.	
12a. DISTRIBUTION / AVAILABILITY	STATEMENT		12b. DISTRIBUTION CODE
Approved for public release; Distribution is unlimited.			
13. ABSTRACT (Maximum 200 wor The prevalence of Schistosom communities in the Qalyub re body weight in a single oral d volunteers) submitted three da winter when transmission was studies to evaluate optimal do transmission are indicated.	os) a mansoni infection was 52% gion of the Nile Delta. Trea lose was taken by 668 (87%) ily faecal samples and 186 (2 low this 31% post treatment sage of praziquantel and sequ	6 among 1494 male farm tment with praziquantel of the infected farmers. 31%) were infected. Sint infection is most likely uential treatment schedul	hers aged 15-42 years old in 3 rural at the recommended dose 40 mg/kg After 3 months 607 (91% of the nee this study was conducted during the due to treatment failure. Prospective es to achieve effective control of
14. SUBJECT TERMS Schistosoma mansoni; Treatm Nile Delta, Egypt.	ent; Praziquantel; Farmers;		15. NUMBER OF PAGES
17. SECURITY CLASSIFICATION OF REPORT	18. SECURITY CLASSIFICATION OF THIS PAGE	19. SECURITY CLASS	FICATION 20. LIMITATION OF ABSTRACT
UNCLASSIFIED	UNCLASSIFIED	UNCLASS	SIFIED

anuan	υr	0101	23	o (ne	Ψ.	4-0:
rescribed	by	ANSI	Std	239-18	1	

## Journal of Tropical Medicine

and the second secon

# JTM

Trealcal

The Society of Fellows of the Egyptian branch of the

Royal Society of Tropical Medicine & Hygiene

Tager Building, 1, Ozoris Street, Garden City, Cairo, Egypt. Tel.: 3541857 Legalized No. 3605

Editorial Board Address : Professor Kabil S M, 4 Green Street off 6 Souk El-Tawfiqia St., Supreme Court Sq., Central Cairo, Egypt.Tel/Fax (202):767688. 

#### JOURNAL OF TROPICAL MEDICINE JTM May, (1993). Vol. 2, No.4, Abu-Elyazeed et al., 3 - 7

#### SCHISTOSOMA MANSONI INFECTION 3 MONTHS AFTER PRAZIQUANTEL THERAPY AMONG FARMERS IN QALYUB, EGYPT

#### Abu-Elyazeed R R, \*Mansour N S, Habib M and Podgore J K Naval Medical Research Unit No. 3, Cairo, Egypt and \*Center for Field and Applied Research, Egyptian Ministry of Health Cairo, Egypt.

#### Abstract

The prevalence of Schistosoma mansoni infection was 52% among 1494 male farmers aged 15-42 years old in 3 rural communities in the Qalyub region of the Nile Delta. Treatment with praziquantel at the recommended dose 40 mg/kg body weight in a single oral dose was taken by 668 (87%) of the infected farmers. After 3 months 607 (91% of the volunteers) submitted three daily faecal samples and 186 (31%) were infected. Since this study was conducted during the winter when transmission was low this 31% post treatment infection is most likely due to treatment failure.

Prospective studies to evaluate optimal dosage of praziquantel and sequential treatment schedules to achieve effective control of transmission are indicated.

#### Introduction

Schistosomiasis is one of the most important public health problems in Egypt. It is the major occupational disease in rural areas where agricultural workers

Address Correspondence and Request for Reprints to:

Research Publications Branch, U.S. NAMRU-3, On the Extent of Ramsis Street, Postal Code No. 11517 Cairo, Egypt.

acquire the infection through repeated daily contact with schistosomal cercarial infested water (Kloos et al., 1990). Waterresource development projects for irrigation purposes have increased the areas of disease transmission. Treatment of schistosomiasis was simplified with the advent of praziguantel, a safe and effective anti-schistosomal drug. The cure rate with a single dose of 40 mg/kg body weight ranges from 72% to 100% for S. haematobium and 63% to 97% for S. mansoni (King and Mahmoud, 1989), Although praziguantel treatment achieves high cure rates, it does not prevent reinfection. How frequent it is necessary to re-treat patients to achieve lasting impact on re-infection rates is not known.

This study was designed to evaluate effectiveness of praziquantel therapy in a cohort of Egyptian farmers with *Schistosoma mansoni* infection three months after a single dose treatment during the low transmission season.

#### Materials and Methods

#### Study area and study population

The study was conducted in three rural sites in the Qalyub district in the Nile Delta about 30 km north-west of Cairo, Egypt. Three villages (Halaba, Sanafeir and Aghour) were chosen because of a known high prevalence of schistosomiasis

in farmers. Male farmers 18-42 years of age, with no anti-schistosomal treatment during the previous three months, were invited to participate in this study which was the initial step of a study to evaluate a schistosomal topical antipenetrant.

#### Schistosomiasis Survey

Between September and November 1990, all participants were given a labeled 50 ml. plastic centrifuge tube for urine and a wide mouthed plastic screw-cap container for faeces and were asked to provide specimens the next morning. Specimen examinations were done within 2-4 hours of collection. Urine samples were examined by a sedimentation concentration technique (Mansour et al., 1981) and faecal samples were examined using the Kato-Katz thick smear method (Katz et al., 1972).

#### **Treatment Procedure**

Praziquantel in a single oral dose of 40 mg/kg was given under the supervision of Egyptian Ministry of Health physician to each farmer immediately after urine and faecal samples indicated schistosome infection.

#### Post-treatment stool survey

Twelve weeks after treatment, a faecal sample was collected on each of three consecutive days from each subject. Faecal samples were examined by the modified Ritchie technique, a more sensitive procedure to determine complete cure following therapy (Knight et al., 1976). If the first or second faecal sample was positive for *S. mansoni* the subject was determined to be infected and subsequent samples were not examined.

#### Results

Of the 1494 farmers participating in the study, 374 were from Halaba, 615 from Sanafeir and 505 from Aghour. The prevalence of Schistosoma haematobium infection was 0.9% and that of Schistosoma mansoni was 51.5%. Of the 14 individuals with S. haematobium, 11

Table- 1: Prevalence of Schistosoma infection in the study sites.

Type of infection	Halaba	Sanafeir	Aghour	Total
	n=374	n=615	n=505	n=1494
S. mansoni	169(45.2%)	<b>309(50.2%)</b>	291(57.6%)	769(51.5%)
S. haematobium	4(1.1%)	<b>4(0.7%)</b>	6(1.2%)	14(0.9%)

Table- 2: Schistosoma mansoni infection 3 months after praziquantel therapy in the study sites.

Type of infection	Halaba	Sanafeir	Aghour	Total
	n=140	n=257	n=210	n=607
S. mansoni	47(33.6%)	80(31.1%)	59(28.1%)	186(30.6%)

and the second second and a

also had S. mansoni. The prevalences of S. mansoni and S. haematobium positive individuals among the three villages (table-1) were not significantly different. All infected individuals were offered treatment with a single dose of praziquantel, but because the prevalence of S. haematobium was very low (0.9%), only S. mansoni infected individuals were included in this study. Of the 668 (86.9%) individuals treated, 607 (90.0%) provided the required consecutive daily faecal specimens during post treatment followup. Three months after treatment 168 of the 607 individuals (30.6%) were still infected with S. mansoni. There was no significant difference in prevalence of post treatment infection among the study villages (table- 2). The value of three consecutive daily faecal examinations post treatment is shown in figure- 1. The first sample identified 50% or less of those infected.





#### Discussion

The prevalence of S. mansoni infection (51.5%) at the time of this study was higher than that reported previously in this area (El-Alamy and Cline, 1977 and Cline et al., 1977). The increase in prevalence of S. mansoni infection is probably due to the selective nature of this study focusing only on male farmers. The prevalence of S. haematobium (<1%) was much lower than that reported by other investigators in this Qalyub area, but the decrease in S. haematobium prevalence is consistent with its dramatic decline in the Nile Delta (Abdel Wahab et al., 1979). Persistent S. mansoni infection after treatment may be due to infection just prior to treatment, re-infection after treatment, or treatment failure. Praziquantel affects only the mature worm stages of schistosomes, so immature stages at the time of treatment may survive, reach maturity and excrete eggs 4-6 weeks later. Exposure after treatment may result in re-infection with egg excretion after 6-8 weeks. The posttreatment S. mansoni infection of 31% in this study is most likely due to drug failure since the study was done during the low transmission season from December through February. This winter period is considered a low season of schistosomiasis transmission because snail breeding is minimal, most snails either die or hibernate in mid winter and human contact with infested water is greatly reduced. The 69% cure rate is consistent with studies in Sudan (Kardaman et al., 1985); Burundi (Gryseels et al., 1987); Botswana (Friis and Byskow, 1989) and Ethiopia (Simonsen et al., 1990).

Single treatment of S. mansoni infected individuals with praziquantel has a limited effect on schistosomiasis control. A single mass treatment may achieve 70% cure rate but will not prevent re-infection. Repeated treatment of infected individuals, or repeated mass treatment if prevalence rates are high, could achieve long-term control. A field trial to evaluate the efficacy of praziquantel in Sudan, (Kardaman et al., 1985) suggested repeating chemotherapy every 6 months for school children living in high prevalence area.

The present study suggests that examination and treatment of a population should be done at the end of the high transmission season with re-examination three months after therapy and re-treatment of infected individuals prior to the beginning of the next high transmission season. If re-examination and treatment are not conducted the high number of infected patients remaining in the population will result in a high level of transmission in the next season with persistence of endemic patterns of the disease. Improved cure rates up to 85% such as those reported by El-Masry et al., (1988), utilizing a 60 mg/kg body weight in divided doses alone or in conjunction with a 3 months follow-up treatment, would be another possible approach to reducing the high level of residual infection due to single 40 mg/kg body weight praziquantel treatment failure.

#### Acknowledgements

This research was supported by the U.S. Naval Medical Research and Development Command, Bethesda, MD, work unit No. 3M464758D849EB.

The opinions and assertions contained herein are the private ones of the authors and are not to be construed as official or as reflecting the views of the Navy Department, Department of Defense, the U.S. Government or the Egyptian Ministry of Health.

#### References

Abdel-Wahab MF, Strickland GT,

6

El-Sahly A, El Kady N, Zakaria S and Ahmed L. (1979): Changing pattern of schistosomiasis in Egypt 1935-1979. Lancet. 2, 242:244.

Cline BL, Richards FO, El-Alamy M, El-Hak S, Ruiz-Tiben E, Hughes JM and Mc Neely DF. (1989): 1983 Nile Delta schistosomiasis survey: 48 years after Scott. Am. J. Trop. Med. Hyg., 41, 56:62.

El-Alamy MA and Cline BL. (1977): Prevalence and intensity of Schistosoma haematobium and S. mansoni infection in Qalyub, Egypt. Am. J. of Trop. Med. Hyg., 26, 470:472. El-Masry NA, Bassily S and Farid Z. (1988): A comparison of the efficacy and side effects of various regimens of praziquantel for the treatment of schistosomiasis. Trans. Roy. Soc. Trop. Med. Hyg., 82, 719:720.

Friis H and Byskow J. (1989): The effect of praziquantel against Schistosoma mansoni infection in Botswana. Tropical Geographical Medicine. 41, 49:51.

Gryseels B, Nkulikyinka L and Coosemans MH. (1987): Field trials of praziquantel and oxaminiquine for the treatment of schistosomiasis mansoni in Burundi. Trans. Roy. Soc. of Trop. Med. Hyg., 81, 641:644.

Kardaman MW, Fenwick A, El-Igail AB, El-Tayeb M, Affalia AA and Dixon HG. (1985): Treatment with praziquantel of school children with current S. mansoni and S. haematobium infections in Gezira. Sudan journal of Tropical Medicine and Hygiene. 88, 105:109.

Katz N, Chaves A and Pallegrino J. (1972): A simple device for quantitative stool thick smear technique in schistosomiasis mansoni. Revista Do Instituto De Medicina Tropical De Sao, 14, 397:400.

King CH and Mahmoud AF. (1989): Drugs Five Years Later: Praziquantel. Annals of Internal Medicine. 110, 290:296.

Kloos H, Higashi GI, Schinski VD, Mansour NS, Murell KD and Miller FD. (1990): Water contact and Schistosoma haematobium infection in a Rural village in upper Egypt. Am. J. Trop. Med. Hyg., 19, 3, 749:758.

Knight WB, Huitt RA, Line BL and Ritchie LS. (1976): A modification of the formol-ether concentration technique for increased sensitivity in detecting *Schistosoma* mansoni eggs. Am. J. Trop. Med. Hyg., 25, 818:823.

Mansour NS, Higashi GI, Schinki VD and Murell KD. (1981): A longitudinal study of Schistosoma haematobium infection in Quena Governorate, upper Egypt. Am. J. Trop. Med. Hyg., 30, 795.

Simonsen PE, Nega A and Furu P. (1990): Intestinal schistosomiasis among children in a labor village of Wonji Sugar Estate, Ethiopia. East African Medical Journal. 67, 532: 538.

