Assessment of Prothrombin Time Activated Partial Thromboplastin Time, and Platelets Count among Children with Schistosomiasis at Alhajalej School, Assalay Locality, White Nile State, Sudan

Elham Elamin<sup>1</sup>, Abdelhakam G. Tamomh<sup>2</sup>, Ahmed M. E. Elkhalifa, <sup>1,3</sup> Emmanuel Ifeanyi Obeagu<sup>4</sup>\* Almanna A. HassbAllah<sup>1</sup>, Ibrahim E. Mustafa <sup>1</sup>, and Yunus B. Y. Ahmed<sup>1</sup>

<sup>3</sup>College of Health Sciences, Public Health Department, Saudi Electronic University, Kingdom of Saudi Arabia, Riyadh

# \*Corresponding Author: Emmanuel Ifeanyi Obeagu

Department of Medical Laboratory Science, Kampala International University, Ishaka, Uganda

**Tel**: +234-8037369912

Email: emmanuelobeagu@yahoo.com https://orcid.ORG/0000-0002-4538-0161

#### Abstract

Schistosomiasis plays critical role to increase the risk for several diseases worldwide particular in developing country, also are related to hematologic changes by disturbing blood flow and endothelial function, which leads to hypercoagulability. The study was done to estimate a Prothrombin time [PT], and international normalization ratio [INR], Activated Partial Thromboplastin Time [APTT], and Platelets count among school Children infected and non-infected Schistosomiasis. Cross – section study included 149 school children [101 cases, and 48 controls] were enrolled in this study, all were matched age and sex. PT [16.80 ± 4.18 Sec], INR Citation: Elamin E, Tamomh AG, Elkhalifa AME, Obeagu EI, HassbAllah AA, Mustafa IE, Ahmed YBY. Assessment of Prothrombin Time Activated Partial Thromboplastin Time, and Platelets Count among Children with Schistosomiasis at Alhajalej School, Assalay Locality, White Nile State, Sudan. *Elite Journal of Haematology*, 2024; 2(4): 10-22

<sup>&</sup>lt;sup>1</sup> Department of Hematology& Immunohematology, Faculty of Medical Laboratory Sciences, University of El Imam El Mahdi, Kosti, White Nile State, Sudan.

<sup>&</sup>lt;sup>2</sup> Departments of Parasitology and Medical Entomology, Faculty of Medical Laboratory Sciences, University of El Imam El Mahdi, Kosti, White Nile State, Sudan.

<sup>&</sup>lt;sup>4\*</sup>Department of Medical Laboratory Science, Department of Medical Laboratory Science, Kampala international university, Uganda

[1.58  $\pm$  0.27], APTT [35.45 $\pm$  3.85 Sec], were significantly higher, while the platelet count [261000 $\pm$  6500 count/ cum] was significantly lower in schistosomiasis infected than non - infected students, PT [11.50  $\pm$  1.32 Sec], INR [0.90  $\pm$  0.19], APTT [29.67  $\pm$ 1.4 Sec], were significantly higher, while the platelet count [280000  $\pm$  12000 count/ cumm\_] with *P. value* < 0.001. Prothrombin time [PT], international normalization ratio [INR], and activated partial thromboplastin time [APTT], were significantly higher, while the platelet count was significantly lower in schistosomiasis infected than non - infected students.

**Keywords:** Schistosomiasis, PT, INR, APTT, platelet count, Alhajalej School Children.

# Introduction

Blood coagulation is host defense system closed to maintain the integrity of high circulation system after blood vessel injury [1]. Hemostasis is functional process to maintain blood in the fluid state and minimize blood loss via the arrest of bleeding at sites of vascular injury [2]. The coagulation system is involved in the conversion of soluble fibrinogen to fibrin clot and consists of various protein factors produced by the liver [3]. Platelets play a critical role in the process of stopping bleeding at the site of interrupted endothelium through platelet adhesion, aggregation, and activation of the coagulation system [4, 5]. Activated platelets release adenosine Diphosphate (ADP), which induce vasoconstriction, stimulate secondary coagulation, and promote further platelet activation and aggregation [6, 7]. The sequential activation of certain plasma, proenzyme that proceeds of blood through the intrinsic or extrinsic pathway [8].

Schistosomiasis is a significant health parasitic infection problem increasing risk for several diseases worldwide particular in developing country [9].

Globally, around 207 million people are infected with schistosome and 120 million of these suffer chronic symptoms [10]. Schistosomiasis is grades second only to malaria in standing among parasitic diseases [11], which can cause abdominal pain, portal hypertension, hepatic and intestinal fibrosis in chronically infected patients, as well as nutritional Iron deficiency anemia [12].

Patients with hepatosplenic Schistosomiasis are disposed to develop complex potential risk of bleeding in those patients which changes in procoagulant- anticoagulant balance, associated with low level of vitamin K-dependent and contact factor proteins were prominent in hepatosplenic Schistosomiasis, main while state of low-grade association chronic disseminated intravascular coagulation [DIC], likewise reduced to presence of immunological and / or inflammatory stimulus [13, 14]. Schistosomiasis usages various mechanisms to prevent primary hemostasis. The schistosome tegument contains several enzymatic activities that lead to the degradation of ADP, resulting in inhibition of ADP-mediated platelet activation and aggregation [15, 16].

#### **Materials and Methods**

This is cross-sectional study was conducted to assess the Prothrombin time [PT], and international normalization ratio [INR], Activated Partial Thromboplastin Time [APTT], and Platelets count among Children with Schistosomiasis infected and non-infected student

Total of 149 students [101 infected schistosomiasis students were classified into two groups [65 infected with *Schistosoma mansoni [S. mansoni]*, and 36 students infected with urinary Schistosomiasis [*S.haematobium*], all are male aged 10-15 years; compare with 48 students non-infected schistosomiasis used as controls. from Alhajalej, Assalay Locality, White Nile State, Sudan, during the period of September 2022 through to March 2023.

#### **Inclusion Criteria**

All students attending Alhajalej children schools in Assalay locality; and who agreement to participate during the study period.

### **Exclusion Criteria**

Multiple parasite infections, students who were on anticoagulant therapy, students having history hypertension, cardiac disease, and diabetes mellitus, chronic renal disease, bleeding disorders, liver disease, Student who received treatment of parasitic infections during the last two weeks before the study were excluded, and/ or who refused to give their consent.

#### **Ethical consideration**

The study agreement received ethical permission from the Ministry of Health and Ministry of Education, then administration of the schools. Objective of study were well explained to school student and their parents, then asked for verbally.

# **Data collection**

Data was collected by using a questionnaire which includes personal, clinical information and laboratory investigation.

#### **Data Analysis:**

Data was exported into the statistical package for social sciences (SPSS) software, version 26 [Chicago, IL, USA] from Microsoft Excel 7. Descriptive statistics were used to describe the variables of the study samples, the P. value of < 0.05 was considered statistically significant

### Sample collection

Four milliliter (ml) of venous blood was collected by venipuncture, and then divided into two tubes (one with 2.25 ml of blood in 0.25ml 3.2% tri-sodium citrate for coagulation tests PT, INR, APTT and other tube with Ethylene Demine Tetra-Acetic Acid (EDTA) (1.75 ml of blood for Hb, PCV, platelet count.

#### **Methods**

**Prothrombin Time (PT)** 

Preparation of Platelet Poor Plasma (PPP)

Platelet Poor Plasma (PPP) is prepared by centrifugation of citrate blood at 2000g for 15 minute at 4 °C, and the test was performed immediately after samples were prepared.

#### Method

Deliver 0.1ml of PPP into small test tube (65x10mm) placed in water bath at **37°C**. Added 0.1 ml of Thromboplastin, Wait for 1-2 min to allow the mixture to worm (Thromboplastin without calcium); then add 0.1 ml of warmed cacl<sub>2</sub> and mixed well, Start stop watch until the cacl<sub>2</sub> was added. Expressed the PT in second as the mean of duplicated for control and test plasma [**28**].

# **Activated Partial Thromboplastin Time (APTT):**

APTT test measures the clotting time of plasma after the activation of contact Factors, calcification by adding of  $CaCl_2$  to phospholipids.

# Method

Pre-warmed CaCl<sub>2</sub> reagent in the water bath at 37°C for at least 10 min.

In clean test tube, 0.1 ml of plasma from test or control samples was added in water bath at 37°C, mixed with equal volumes of APTT reagents (phospholipids reagent and the kaolin), wait for 1-3 min, after that, 0.1 ml of pre-warmed CaCl<sub>2</sub> was added and start a stopwatch immediately. The time for clot formation was observed and recorded; the results of APTT were expressed in second as the mean of duplicated for control and test plasma [28].

#### **Results**

Table 1: General characteristic of student infected, and non-infected schistosomiasis in Alhajalej School

Variables	infected Schistosomiasis	Non- infected [control]
Male	101 [68%]	48[32%
Age /years	13.11 ±1.56	13±2.00
Hb g/dl	$11.46 \pm 1.85$	$14.96 \pm 1.5$
PCV %	$35.51 \pm 5.80$	$45.00 \pm 1.90$
APTT/ Sec	$35.45 \pm 3.85$	29.67± 1.4
PT/ Sec	$16.80 \pm 4.18$	11.50 ±1.32
INR	$1.58 \pm 0.27$	$0.90 \pm 0.19$
Platelet count/cumm	$261000 \pm 6500$	280000± 12000

**Hb**: Hemoglobin, **g/dl**: gram per disliter, **PCV**: Packed Cell Volume, **APTT**: Activated Partial Thromboplastin Time, PT: Prothrombin Time, **Sec**: Second, and **cumm**: Cell per cubic millimeter.

Table 2: Compression of Age, Hb, PCV, APTT, PT, INR and Platelet count among infected Schistosomiasis student with non-infected in Alhajalej school children

Variables	infected- Schistosomiasis	Non- infected [control]	P-value
Male	101 [68%]	48[32%]	0.003
Age /years	13.11 ±1.56	13±2.00	0.71
Hb g/dl	$11.46 \pm 1.85$	$14.96 \pm 1.5$	0.001
PCV %	$35.51 \pm 5.80$	$45.00 \pm 1.90$	0.001
APTT/ Sec	$35.45 \pm 3.85$	29.67± 1.4	0.001
PT/ Sec	$16.80 \pm 4.18$	$11.50 \pm 1.32$	0.001
INR	$1.55 \pm 0.27$	$0.90 \pm 0.19$	0.001
Platelet	$261000 \pm 6500$	$280000 \pm 12000$	0.001
count/cumm			

**Hb**: Hemoglobin, **g/dl**: gram per disliter, **PCV**: Packed Cell Volume, **APTT**: Activated Partial Thromboplastin Time, PT: Prothrombin Time, **Sec**: Second, and **cumm**: Cell per cubic millimeter.

Table 3: Compression of Hb, PCV, APTT, PT, INR and Platelet count among infected Schistosomiasis students according to Schistosoma type in Alhajalej school children

Variables	S. mansoni	S.haematobium	P. value
Infected students	65 [64%]	36[36%]	0.01
Hb g/dl	$11.7 \pm 2.3$	13.03 ±1.06	0.001
PCV %	$35 \pm 7.00$	$36.01 \pm 4.68$	0.001
APTT/ Sec	$37.82 \pm 3.78$	$33.08 \pm 3.92$	0.001
PT/ Sec	17.03± 4.99	13.07± 3.36	0.004
INR	$1.57 \pm 0.26$	$1.32 \pm 0.27$	0.001
Platelet count/cumm	$257000 \pm 7100$	$266000 \pm 5900$	0.001

**S.** *mansoni*: Schistosoma Mansoni, **S.** *haematobium*: Schistosoma *Haematobium*, **Hb**: Hemoglobin, **g/dl**: gram per disliter, **PCV**: Packed Cell Volume, **APTT**: Activated Partial Thromboplastin Time, PT: Prothrombin Time, **Sec**: Second, and **cumm**: Cell per cubic millimeter.

Table 4: Correlation of Hb, PCV, APTT, PT, INR and Platelet count among infected with non-infected Schistosomiasis students according to Schistosoma type in Alhajalej school children

Variables	infected Schistosomiasis		non infected
	S. mansoni	S.haematobium	[control]
Male	65 [43%]	36[24%]	48[32%]
Age /years	13.18 ±1.71	13.03 ±1.06	13±2.00
Hb g/dl	$11.7 \pm 2.3$	$11.21 \pm 1.41$	14.96 ± 1.5 *P
PCV %	$35 \pm 7.00$	$36.01 \pm 4.6.8$	29.00 ± 1.90 *P
APTT/ Sec	$33.82 \pm 3.78$	$33.08 \pm 3.92$	28.67± 1.4 *P
PT/ Sec	17.03± 4.99	13.07± 3.36	10.50±.50 *P
INR	$1.57 \pm 0.26$	$1.32 \pm 0.27$	$0.90 \pm 0.10$ *P
Platelet count/cumm	$257000 \pm 7100$	$266000 \pm 5900$	280.0 0± 20000 *P

**S.** *mansoni*: Schistosoma Mansoni, **S.** *haematobium*: Schistosoma *Haematobium*, **Hb**: Hemoglobin, **g/dl**: gram per disliter, **PCV**: Packed Cell Volume, **APTT**: Activated Partial Thromboplastin Time, PT: Prothrombin Time, **Sec**: Second, and **cumm**: Cell per cubic millimeter.

### **Discussion**

<sup>\*</sup>P. value consider statistically significant < 0.05

Globally schistosomiasis increases mortality and morbidity rate of death among school children, the prevalence of intestinal and urinary schistosomiasis is major public health problem [44, 45]. Schistosoma mansoni parasites can live for years within human blood vessels and appear to be refractory to intravascular thrombus formation deaths per year [24].

Urinary schistosomiasis causes chronic infection with negatively affect all aspects of health, nutrition, learning, considerable growth obstruction and anemia, hematuria, as well as coagulation changes in countries where the disease is endemic [44].

This study was aimed to evaluate basic coagulation profile [activated partial Thrombo-plastin time [APTT], prothrombin time [PT] as well as international normalization ratio [INR], likewise, platelets count among schistosomiasis infected, and non-infected students attending Alhajalej school children at Assalay locality, White Nile State, Sudan during the period of September 2021 to March 2022. Our results included total of 149 students 101[68%] infected schistosomiasis, 48 [32%] students non-infected], all were matched age and sex table: 1 Our revealed that from 101students 65[43%] infected S. mansoni, and 36 [24%] infected S. haematobium table 3. Our results revalued APTT, PT, INR, and platelet count [35.45 $\pm$  3.85 Sec, 16.80  $\pm$  4.18 Sec, 1.58  $\pm$  0.27, and 261000  $\pm$  6500 count/ cumm ], in effected schistosomiasis students in relation to non- infected [29.67  $\pm$  1.4 Sec, 11.50  $\pm 1.32$  Sec,  $0.90 \pm 0.19$ , and platelet count [  $280000 \pm 12000$  count/ cumm ] respectively, table 4.2, 4, 3 were statistically different p. value = 0.001. This results comparable by Eyayu et al 2020 both, intrinsic [APTT] and extrinsic [PT, INR), among an infected patients; could possibly indicate is either inhibition of both intrinsic and extrinsic coagulation pathways, or an inhibition of the common pathway [17], and apparently, S. mansoni was found by Da'dara et al 2016 to accelerate the formation of blood clots in murine blood in vivo, however possibly compensated by a rapid, abnormal, fibrinolytic clot breakdown due to reduced blood platelet count and fibrinolysis [46].

This finding is harmonized with the previous discussion on the study done by Lagler *et al.* [47], Da'dara *et al.* hypothesized that this may be a result of reduced levels of coagulation control

proteins which promotes rapid blood clotting, and as counter balance, rapid fibrinolytic clot break down may occur due to reduced blood platelet count and fibrinolysis [46].

Adult S. *mansoni* and eggs induced alteration in endothelial function and schistosomes have several electronegative charges on their tegument that could potentially activate platelets and coagulation cascade, leading to hypercoagulation granulomas, increased consumption of coagulation factors and decreased hepatic synthesis of these factors due to liver abnormality were the possible reason for the occurrence of prolonged PT, INR and APTT in S. mansoni-infected children [16, 35].

When compared hematological parameters Hb g/dl, PCV%, platelet counts in Schistosoma infected Hb g/dl [ $11.46 \pm 1.85$ ] while PCV% [ $35.51 \pm 5.80$ ], and with non-infected students Hb g/dl [ $14.96 \pm 1.5$ ] while PCV% [ $45.00 \pm 1.90$ ] we fount statistically different decreases p- value = 0.001 [table 4.3]. This study revealed that the platelet count had significantly decreased in S. mansoni-infected students more serious than Schistosoma haematobium—compared to non-infected controls p value < 0.001. This is in agreement with studies done in Brazil, in China, and Egypt [37, 39- 40]. These results explain by splenic retention due to poor portal blood drainage or platelet draped sinusoidal space of liver fibrotic [48].

In this study Schistosoma was evaluated only by basic coagulation profile [APTT, PT, INR], and platelets count as well as CBC], further studies contacted iron profile, D-dimer, factors assay particularly contact factors.

#### **Conclusion**

Prothrombin time [PT], international normalization ratio [INR], and activated partial thromboplastin time [APTT], were significantly higher, while the platelet count was significantly lower in schistosomiasis infected than non - infected students.

PT, INR, APTT coagulation tests, and platelet count should be used to monitor and manage schistosomiasis-related complications

#### References

- **1.** Antoniak S. The coagulation system in host defense. Research and Practice in Thrombosis and Haemostasis. 2018; 2(3):549-557.
- **2.** Hickman DA, Pawlowski CL, Sekhon UD, Marks J, Gupta AS. Biomaterials and advanced technologies for hemostatic management of bleeding. Advanced Materials. 2018; 30(4):1700859.
- **3.** Vilar R, Fish RJ, Casini A, Neerman-Arbez M. Fibrin (ogen) in human disease: both friend and foe. *Haematologica*. 2020; 105(2):284.
- **4.** Palta S, Saroa R, Palta A. Overview of the coagulation system. *Indian J Anaesth*.2014; 58(5):515–523.
- **5.** Hou Y, Carrim N, Wang Y, Gallant RC, Marshall A, Ni H. Platelets in hemostasis and thrombosis: novel mechanisms of fibrinogen-independent platelet aggregation and fibronectin-mediated protein wave of hemostasis. *J Biomed Res.* 2015; 29(6):437–444.
- **6.** Broos K, Feys HB, De Meyer SF, Vanhoorelbeke K, Deckmyn H. Platelets at work in primary hemostasis. *Blood reviews*. 2011; 25(4):155-67.
- **7.** Matus MF, Vilos C, Cisterna BA, Fuentes E, Palomo I. Nanotechnology and primary hemostasis: Differential effects of nanoparticles on platelet responses. *Vascular Pharmacology*. 2018; 101:1-8.
- **8.** Shamanaev A, Emsley J, Gailani D. Proteolytic activity of contact factor zymogens. *Journal of Thrombosis and Haemostasis*. 2021;19(2):330-41.
- **9.** Verjee MA. Schistosomiasis: still a cause of significant morbidity and mortality. *Research and reports in tropical medicine*. 2019; 10:153.
- **10.** Weatherhead JE, Hotez PJ, Mejia R. The global state of helminth control and elimination in children. *Pediatric clinics*. 2017; 64 (4):867-77.
- **11.** Kinung'hi SM, Mazigo HD, Dunne DW, Kepha S, Kaatano G, Kishamawe C, Ndokeji S, Angelo T, Nuwaha F. Coinfection of intestinal schistosomiasis and malaria and association with haemoglobin levels and nutritional status in school children in Mara region, Northwestern Tanzania: a cross-sectional exploratory study. *BMC research notes*. 2017;10(1):1-1.

- **12.**Thijs L, Messiaen P, van der Hilst J, Madoe V, Melis C, Van Eyken P, Vanmoerkerke I, Janssens F. Hepatic schistosomiasis with massive splenomegaly: a case report and literature. *Acta Gastro-Enterologica Belgica*. 2018;81.
- **13.** Yang ZL, Guo T, Zhu DL, Zheng S, Han DD, Chen Y. Risk factors of portal vein thrombosis after splenectomy in patients with liver cirrhosis. *Hepatoma Research*. 2020; 6:37.
- **14.** Abou-Alfa GK, Ang C. Dx/Rx: Liver Cancer: Liver Cancer. Jones & Bartlett Publishers; 2012.
- **15.** Lovászi M, Haas CB, Antonioli L, Pacher P, Haskó G. The role of P2Y receptors in regulating immunity and metabolism. *Biochemical pharmacology*. 2021; 187:114419.
- **16.** Mebius MM, van Genderen PJ, Urbanus RT, Tielens AG, de Groot PG, van Hellemond JJ. Interference with the host haemostatic system by schistosomes. *PLoS pathogens*. 2013;9(12): e1003781.
- **17.** Eyayu T, Zeleke AJ, Seyoum M, Worku L. Basic coagulation profiles and platelet count among Schistosoma mansoni-infected adults attending Sanja Primary Hospital, Northwest Ethiopia. *Research and Reports in Tropical Medicine*. 2020; 11:27.
- **18.** Li J. The potential of Schistosoma-derived substances for use as basis for novel anti-haemostatic therapeutics: A systematic review.
- **19.** Bica I, Hamer DH, Stadecker MJ. Hepatic schistosomiasis. Infectious disease *clinics of North America*. 2000; 14(3):583-604.
- **20.** Kaatano GM, Min DY, Siza JE, Yong TS, Chai JY, Ko Y, Chang SY, Changalucha JM, Eom KS, Rim HJ. Schistosoma mansoni-related hepatosplenic morbidity in adult population on Kome Island, Sengerema district, Tanzania. *The Korean journal of parasitology*. 201553(5):545.
- **21.** Olopade Bo. Intestinal Parasites, Nutritional Status and Cognitive Function among Pupils in Ile-Ife, Osun State. Faculty of Pathology. 2014.
- 22. Gryseels B. Schistosomiasis. Infect Dis Clin North Am. 2012;26 (2):383-97.
- **23.** Moné H, Boissier J. Sexual biology of schistosomes. *Adv Parasitol*. 2004; 57:89-189.

- **24.**Centers for Disease Control and Prevention Laboratory Identification of Parasites of Public Health Concern. Schistosomiasis Life Cycle [Image]. 2010.
- **25.** Mari L, Ciddio M, Casagrandi R, Perez-Saez J, Bertuzzo E, Rinaldo A, *et al.* Heterogeneity in schistosomiasis transmission dynamics. *J Theor Biol*. 2017; 432:87-99.
- **26.**Centers for Disease Control and Prevention and Control (CDC). Life cycle of schistosomiasis .2012.
- **27.**Tsiklidis E, Sims C, Sinno T, Diamond SL. Multiscale systems biology of trauma induced coagulopathy. Wiley Interdisciplinary Reviews: *Systems Biology and Medicine*. 2018;10(4): e1418.
- **28.** Mussbacher M, Kral-Pointner JB, Salzmann M, Schrottmaier WC, Assinger A. Mechanisms of hemostasis: Contributions of platelets, coagulation factors, and the vessel wall. In Fundamentals of Vascular Biology 2019: 145-169. *Springer, Cham.*
- **29.** Freato N, van Alphen FP, Boon Spijker M, van den Biggelaar M, Meijer AB, Mertens K, Ebberink EH. Probing activation driven changes in coagulation factor IX by mass spectrometry. *Journal of Thrombosis and Haemostasis*. 2021 Jun;19(6):1447-59.
- **30.** Bronić A, Coen Herak D, Margetić S, Milić M. Croatian Society of Medical Biochemistry and Laboratory Medicine: National recommendations for blood collection, processing, performance and reporting of results for coagulation screening assays prothrombin time, activated partial thromboplastin time, thrombin time, fibrinogen and D-dimer. *Biochemia medica*. 2019;29(2):262-283.
- **31.** Boroumand M, Goodarzynejad H. Monitoring of anticoagulant therapy in heart disease: considerations for the current assays. *The Journal of Tehran Heart Center*. 2010; 5(2):57.
- **32.** Normal ME. Proceedings of Réanimation 2020, the French Intensive Care Society International Congress. *Ann. Intensive Care*. 2020; 10(1):16.
- **33.** Da'dara AA, Skelly PJ. Schistosomiasis versus platelets. *Thromb Res*. 2014;134(6):1176–1181.
- **34.** Bhardwaj R, Skelly PJ. Purinergic signaling and immune modulation at the schistosome surface? *Trends Parasitol*. 2009; 25(6):256–260.

- **35.** Tanabe M. Haemostatic abnormalities in hepatosplenic schistosomiasis mansoni. *Parasitol Int.* 2003;52(4):351–359.
- **36.** Roberts DJ. Hematologic changes associated with specific infections in the tropics. *Hematol Oncol Clin*. 2016; 30(2):395–415.
- **37.** Leite LAC, Pimenta Filho AA, da Fonseca CSM, et al. Hemostatic dysfunction is increased in patients with hepatosplenic schistosomiasis mansoni and advanced periportal fibrosis. *PLoS Negl Trop Dis.* 2013; 7(7): e2314.
- **38.** Correia MC, **Domingues** AL, Lacerda HR, al. Platelet function et and factor antigen hepatosplenic Willebrand in the form of schistosomiasis mansoni. Trans R Soc Trop Med Hyg. 2009; 103 (10): 1053–1058.
- **39.** Shun L. Meng Q, Shao-Qian T. **Analysis** of coagulation related schistosomiasis parameters between patients with advanced cirrhosis 2016:29 and hepatitis В cirrhosis. Chin JSchistosomiasis Control. (1):68-71.
- **40.**Leite LAC, Domingues ALC, Lopes EP, et al. Relationship between splenomegaly and hematologic findings in patients with hepatosplenic schistosomiasis. *Rev Bras Hematol Hemoter*. 2013;35 (5):332–336.
- **41.**Osman MA. Evaluation of Some Haemostatic Parameter among Schistosomiasis Parameters in Algleea Village—Shendi. Sudan University of Science and Technology; 2012.
- **42.** Nagy ZP, Varghese AC, Agarwal A, editors. Practical manual of in vitro fertilization: advanced methods and novel devices. Springer Science & Business Media; 2012.
- **43.** Chernecky, Cynthia C. and Barbara Berger. Lab Tests and Diagnostic Processes. 3rd ed. Philadelphia: W. B. Saunders Company, 2001.
- **44.** Amin MA, Kardaman MAM, Mounkaila N, Abubaker H, Algali M, Homeida M. The transmission patterns of schistosomiasis in Khartoum State, Sudan. *Ann Clin Pathol* . 2016; 4(6): 1088.

- **45.**Da'dara AA, de Laforcade AM, Skelly PJ. The impact of schistosomes and schistosomiasis on murine blood coagulation and fibrinolysis as determined by thromboelastography (TEG). *J Thromb Thrombolysis*. 2016;41(4):671-7.
- **46.** Lagler H, Ay C, Waneck F, Gattringer R, Graninger W, Ramharter M. Characterisation of inflammatory response, coagulation, and radiological findings in Katayama fever: a report of three cases at the Medical University of Vienna, Austria. *BMC Infect Dis*. 2014; 14:357.
- **47.** de Araújo Souza MR, De Toledo CF, Borges DR. Thrombocytemia as a predictor of portal hypertension in schistosomiasis. *Dig Dis Sci.* 2000; 45 (10):1964–1970.