Prevalence of parasitic infections in Iranian stable hemodialysis patients *Running title: parasitic infections in hemodialysis patients*

Shiva SEYRAFIAN¹ M.D; Associate professor of Nephrology.

Nader PESTECHIAN² PhD; Assistant professor of Parasitology, Department of Parasitology, Faculty of Medicine.

Nasrin NAMDARI¹ M.D, Mahshid KAVIANI¹ M.D, Maryam KERDEGARI¹ M.D Farzad PARVIZIAN¹ M.D, Leila KASSAII¹ M.D, Afrouz ESHAGHIAN¹ M.D Hamid NASRI¹'* M.D; Professor of Nephrology

Abstract:

Background: Hemodialysis (HD) patients are prone to infections as a result of impaired immune system. Early detection of disease helps to prevent complications. The aim of this study was to evaluate the prevalence of intestinal parasite infections in HD patients and compare it with control groups. Methods: In a cross sectional study, the stool sample of 155HD patients, and 294 controls were examined for parasitic contaminations. Control groups included: 130 patients' family, 16 staffs of three HD wards and 148normal populations . 3 stool samples were taken from each participant. After perceiving the appearance of the samples and preceding the stages of concentration with Formaline-ether method, direct smear were prepared and inspected by trichrome staining. Then, groups were compared with SPSS version12 by chi-square and T-test methods. Results: 43.9% of 155HD patients were infected by intestinal parasites. There was 40% parasite infection in non diabetic and 45% in diabetic case groups with no significant difference between the 2 groups (P>0.05). There was no relation between parasite infection with sex, HD duration and use of immunosuppressive drugs. 43.1% of control group was also infected. No significant difference was seen between the 2 groups (P>0.05). The most common parasite was Blastocystis hominis in the 2 groups. Conclusion: This study could not show increased parasite infection in HD patients compared to control groups. The high prevalence of

¹Kidney research center, Isfahan University of Medical Sciences, Isfahan, Iran

² Isfahan University of Medical Sciences, Isfahan, Iran

^{*}Author to whom correspondence should be addressed; Tel.: +98 311 2208082 (hospital); Fax: +98 311 2235043. Email; hamidnasri@yahoo.com

intestinal parasites in HD patients and control groups, may indicate that population hygiene status is not well controlled, and emphasizes more health care providers' attention.

Keywords: End-stage renal disease, Intestinal Parasite infections; Kidney Failure, Chronic

Introduction:

Uremia impairs antigen presentation, T-cell activation, and cause impaired antibody production[1]. Hemodialysis (HD) patients are susceptible to opportunistic infections, as a result of leukocyte dysfunction[2,3], and impaired immunologic response (like phagocytosis, migration, bactericidal action of neutrophils)[4-8]. Infection is the second cause of mortality in dialysis survivors[9,10]. One of the hygienic indices is parasite infections in population. Cryptosporidiosis, Isospora belli, Chilomastix mesnili, Blastocystis hominis, Endolimax nana, Entamoeba Coli, Entamoeba Hartmani and Dientamoeba fragilis are some of the opportunistic parasites which cause serious and recurrent infections in immunocompromised patients, but are self limited in normal healthy populations[11]. Studies regarding parasitic infections in HD patients revealed various results and may be related to socioeconomic state and region of living[12,1] . In this study we aimed to evaluate the prevalence of parasite infections in a group of stable hemodialysis patients, and compare it with normal population.

Methods:

This cross-sectional study was conducted on 155 hemodialysis patients from three university hospitals and three control groups (16 HD ward staffs, 130 persons from patients' family and 148 normal random population). Family groups were similar in eating and drinking habits, environment and home conditions, and normal random healthy population who had been referred to laboratory for other causes. The propose for selecting several control groups was to obviate the effect of hygiene and living environments on the infection rate. None of the control groups suffered from diabetes mellitus, malignancy, leukemia, renal failure, autoimmune diseases and chronic cardiopulmonary disease. All of the patients participated with informed consent. The study was approved by Isfahan university of medical sciences' ethics committee, Isfahan, Iran. Demographic variables consisting age, sex, the length of the time they were on hemodialysis, cause of renal failure, history of any immunosuppressive drug treatment ,history of renal transplantation, and diabetes mellitus were recorded.

Patients with history of taking antibiotics, mineral oil, Barium, Bisthmooth, anti malarial drugs, non-absorbable anti-diarrheal dugs (e.g., hydrated aluminomagnesium silicates and kaolinpectin) during 2 weeks before getting the specimen were excluded from the study [14]. Three containers were given to the patients, and three consecutive specimens were taken during tree alternate days. The watery or loose specimens immediately sent to laboratory for examination; if the specimen was formed and not watery, preserved in refrigerator in 10% aqueous formalin until examination. The stool samples were studied by a single technician. The specimens were evaluated by direct inspection with physiologic serum, and Lugol's solution, then concentration procedure was done for study sedimentation with formalin-ethyl acetate technique. After preparing thin smear from specimens, they stained with trichrom staining. The specimens were studied for 15 types of parasites included Giardia Lamblia, Cryptosporidiosis, Isospora belli, Iodamoeba Büchel, Hymenolepsis nana, Teniae, Chilomastix mesnili, Ascaris lumbricoides, Blastocystis hominis, Strongyloides stercoralis, Endolimax nana, Entamoeba Coli, Entamoeba histolytica, Entamoeba Hartman and Dientamoeba fragilis[15]. The examiner was blinded to the clinical informations. Data were analyzed by SPSS statistical package version 12 (SPSS Inc, Chicago, USA), with chi-square, T-student and Fisher Exact tests. P-value < 0.05 was considered significant.

Results:

In this study we entered 449interviewee including 155hemodialysis patients , 130patients 'family, 148normal individuals and 16 HD ward staffs. They were 224women (49.2 %) and 228 men (50.8%). Mean (\pm SD) age of the patients was 54(\pm 17) year. Mean length of hemodialysis was 36(\pm 33) months. Forty patients (25.8%) had history of cardiovascular disease; Thirty-five patients (22.5%) had diabetes mellitus; Twenty-one patients (13.5%) had history of renal transplantation; Twenty patients (12.9%) had history of using immunosuppressive agents. Sixty-eight (43.9%) of dialysis patients, 61 (46.6%) of patients' family, 48 (32.6%) of normal population and 8 (50%) of dialysis staffs infected by intestinal parasites. All groups had higher protozoan infection than helminthes infection. There was not significant difference in parasite infections between 4 groups (P =0.129).

The rate of parasitic infections is described in table 1.

Table 1- The rate of parasite infections in all groups of the study. (p=0.45)

Groups	Infection with one	Infection with at
	parasite	least two parasites
HD patients	51.5%(35)	48.5%(33)
patients' family	55.7%(34)	44.3% (27)
normal population	54.1%(26	45.9%(22)
staffs of dialysis wards	37.5%(3)	62.5%(5)
staffs of dialysis wards	37.5%(3)	62.5%(5)

There was not relationship between age and rate of infection in HD patients. There was a significant relation between age and parasite infection in patients' family (P = 0.03) (figure 1).

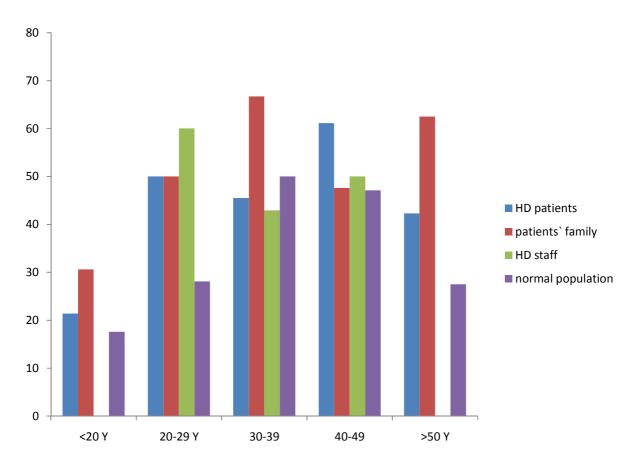


Figure 1-Age of 4 groups with parasite infection. There was a significant relation between age and parasite infection only in patients' family group. (p=0.038)

There was not relationship between parasite infection and sex in all study groups (P > 0.05).

Parasite infections in diabetic and non diabetic case groups was 40% (14 patients) and 45% (54 patients) respectively. There was not significant difference between diabetic and non diabetic case groups (P=0.372). There was no significant difference in length of dialysis time (P=0.791), history of renal transplant (P=0.27), history of immunosuppressive agents (P=0.094), cause of renal failure (P=0.82), and history of diabetes mellitus (P=0.372) with parasite infection.None of the specimens were infected by Ascaris lumbricoides, Strongyloides stercoralis, Isospora Belli and Taniae. The most common cause of parasite infection was Blastocystis Hominis(8%), Entamoba Coli(5.6%) and Endolimax Nana(4.2%) in whole studied population. The highest prevalence rate in HD patients was 48.7% for Blastocystis hominis, followed by Cryptosporidium (17%) and Entamoeba colli (12.1%). There is not any significant relationship in intestinal parasite infection types between all groups (P=0376).

Discussion:

Protozoa and helminthes can cause serious infections in immunocompromised hosts, especially patients with impaired cellular immunity[16] Uremic status causes acquired immunodeficiency syndrome[2-8]. In our study, increases in the prevalence of parasite infection in HD patients in comparison to control groups was not found, also increased risk of infection in diabetic patients was not seen too. This study has shown 43.9% of dialysis patients, 46.9% of patients' families, 32.6% of normal populations and 50% of dialysis staffs infected by intestinal parasites. This prevalence rate was higher than previous report by Nasiri in normal population of Karaj of Iran, in which prevalence of parasite infection was 4.7%[15]. Kulik et al. in the study on 86 hemodialysis and 146healthy controls found, 45.1% of dialysis patients and 25.7% of controls were infected with intestinal parasites. There was a significant difference in prevalence of parasite infections in HD group in comparison to control participants [16]. While we have shown higher rate of Cryptosporidiosis in HD patients in our previous study too[17], in an another study in Iran, no higher rate of this fungi in hemodialysis was seen [18]. In present study, there was not significant association between duration of dialysis and age of the patients with parasite infection. This result was found in our previous study too [17], however in contrast to this finding Hazarati et al. showed significant increase in parasite infection with duration of dialysis[18]. In the present study, no increase in parasitic infection, in HD patients who have a history of renal transplant immunosuppressive agents taking also causes of renal failure was not contributed to parasitic infection in HD patients, similar results was in agreement with our finding [13,19-21]. In this study, the most common cause of parasitic was Blastocystis hominis (8%), Entamoeba Coli (5.6%) and Endolimax nana infections (4.2%).In the study of Kulik et al. the most frequent parasitic infections was Blastocystis hominis(% 18-20.1), Endolimax nana (%14-16.3), Cryptosporidiosis (%4-4.7) and Entamoeba an another study in 1997 in Tehran, Iran, the majority of parasitic Coli (%4-4.7)[16].In infections in cancer patients was Blastocystis hominis (18.2%) and Giardia Lamblia (11.9%) [19].It seems that Blastocystis hominis is a common cause of parasitic infections in these immunocompromised patients. While it was assumed that increased time of uremic status might weaken immune system progressively and resultant increased risk of infectious disease, however other factors like water and food hygiene, population general health may influence the findings.

<u>Conclusion</u>: This study could not show increased parasite infection in HD patients compared to control groups. The high prevalence of intestinal parasites in HD patients and control groups, may indicate that population hygiene status is not well controlled, and emphasizes more health care providers' attention.

Recommendations: As a result of immunosuppressive status in HD patients, we recommend stool exam for parasite detection in HD patients suffer from diarrhea.

Authors' Contributions

Shiva Seyrafian carried defined the aim of research and the design of experiment and carried out the experiments. Nader PESTECHIAN, Nasrin NAMDARI, Mahshid KAVIANI, Maryam KERDEGARI, Farzad PARVIZIAN, Leila KASSAII and Afrouz ESHAGHIAN participated in the design of the study and performed the laboratory exams and also statistical analysis. Hamid NASRI coordinates and helped to draft the manuscript. All authors read and approved the final manuscript.

Acknowledgments

We thanks to Mrs. Mohebrasool, Malekahmadi and Rokooei head nurses of dialysis centers for all helps in this study.

References

- 1-Ocak S. H. and Eskiocak A. F.; The evaluation of immune response to hepatitis B vaccination in diabetic and nondiabetic haemodialysis patients and the use of tetanus toxoid, Nephrology 2008; 13:487.
- 2-Massry Y. G, Alexiewicz J.M., Gacioug Z. Secondary hyperparathyroidism and immune system in chronic renal failure. Semin-Nephrol, 1996; 2: 186 201
- 3-Descomps L.B., Chatenoud L. T cells and B cells in chronic renal Failure. Semin Nephrol, 1996; 16; 182 191
- 4-Dohrety CC., Labelle P., Collins J.F., Massry S.G. Effect of parathyroid hormone on random migration of human PMN. Amj Nephrol 1988, 8, 212-19.
- 5-Alexiewicz J. M, Smoger Zewski: My., Massry S.G. Impaired phagocytosis in dialysis patients: Studies on mechanisms. Anj Nephrol, 1991; 11; 102 111
- 6-Haag W.M. Horl W.H. Uremia and infection: mechanisms of imparied cellular host defense editorial. Nephrol, 1993; 63; 125-131
- 7-Vanholer R., Ringoir S, Dhondt A. Hakim R. Phagocytosis in uremic and hemodialysis. Kindy int, 1991; 39; 320 27.

- 8-<u>Chonchol M.</u> Neutrophil dysfunction and infection risk in end-stage renal disease. <u>Semin Dial.</u> 2006 Jul-Aug;19(4):2946.
- 9-Brenner B M. Chronic renal failure ,Brenner and Rector 's the kidney. 8th edition volume 2; 2008. 1193-1194.
- 10-<u>Vanholder R, Ringoir S</u>. Infectious morbidity and defects of phagocytic function in end-stage renal disease: a review. J Am Soc Nephrol. 1993 Mar;3(9):1541-54.
- 11-Garcia L S, Diagnostic Medical Parasitology, 5th Edition, 2007, p22, 23, 26, 27, 29, 48, 50, 61, 81.
- 12-ttp://www.dpd.cdc.gov/dpdx/HTML/Frames/DiagnosticProcedures/body_dp_stoolcollect.htm.
- 13-World Health Organization. Basic laboratory methods in medical parasitology. World Health Organization, Geneva, 1991. (http://www.dpd.cdc.gov/dpdx/HTML/DiagnosticProcedures.htm).
- 14-Ferreira MS, Borges AS. Some aspects of protozoan infections in immunocompromised patients A review. Mem Inst Oswaldo Cruz Rio de Janeiro 2002; 97:443-57.
- 15-Nasiri V, Esmailnia K, Karim G, Nasir M, Akhavan O. Intestinal parasitic infections among inhabitants of Karaj City, Tehran province, Iran in 2006-2008. Korean J Parasitol. 2009 Sep;47(3):265-8.
- 16-Kulik R. A., Morais Falaovigna D. L.; Nishi L., Araujo S. M. Blastocystis sp. And other intestinal parasites in hemodialysis patients. The Brazilian Journal of Infectious Disease 2008 12 (4): 338-341
- 17- Seyrafian Sh, Pestechian N, Kerdegari M, Yousefi HA, Bastani B, Prevalence rate of cyptosporidium infection in hemodialysis patients in Iran. Hemodialysis International 2006; 10: 375-379
- 18-Hazarati Tappeh KH, Gharavi MJ, Makhdoumi K, Rabat M, Tagizadeh A. Prevalence of cyptosporidium spp. Infection in Renal Transplant and hemodialysis patients. Iranian J publ Health, 2006, vol. 35, No. 3: 54-57.
- 19-Athari E, Marfi H, Relationsheep Between Immunosupressive Drugs and Intestinal Parasitic infections in Iran . Abstract of Papers in 3rd universal Parasitolyg Congress in Mazandaran Medical University 1379; 78.
- 20-Azami M, Sharifi M, Hejazi SH, Tazhibi M. Intestinal parasitic infections in renal transplant recipients. Braz J Infect Dis. 2010 Feb;14(1):15-8.