Infection Rate of Human Parvovirus B19 among Hem dialysis Patients in Bequeath City

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Abstract:Background: Dialyzed patients have an increasing risk of exposure to viral infection such as Human Parvovirus B19, It is a DNA virus that is responsible for causing several diseases in humans.

Objectives: To determine the infection rate of human parvovirus B19 among dialyzed patients in Baqubah City. **Patients and methods:** Cross-sectional research study which includes 51 dialyzed patients who have been selected randomly from Ibn-Sina Hemodialysis Center in Baqubah Teaching Hospital during the period from September 2016 till February 2017. Full information had been taken directly from the patients and the information had been arranged in an informative formula sheet which includes: age, gender, residence, marital status, education levels, occupation, history of blood transfusion, kidney transplant and associated with other disease. All study subjects were screened for anti-human parvovirus B19 IgM and IgG antibodies using enzyme-linked immunosorbent assay.

Result: Anti-human parvovirus B19 IgM antibody was found in 4(7.84%) dialyzed patients and anti-human parvovirus B19 IgG antibody was detected in 46 (90.19%) ones. There was significant difference between both of them. Multivariate analysis of demographic and risk factors showed that male gender, age, marital status, occupation, length of time on hemodialysis and family history were associated with IgM positivity while other showed non-significant differences.

Conclusion: The seroprevalence of human parvovirus B19 was relatively high in dialyzed patients in Baqubah-Iraq, and should be investigated after exclusion of other common causes.

Key words: Hemodialysis, human parvovirus B19, enzyme-linked immunosorbent assay, kidney disease.

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I. Introduction

Chronic kidney disease (CKD), also known as chronic renal disease, is progressive loss in kidney function over a period of months or years [1]. The most common recognized cause of CKD is diabetes mellitus. High blood pressure is also a very common cause of chronic kidney disease. Other causes of CKD include idiopathic (i.e. unknown cause, often associated with small kidneys on renal ultrasound) and glomerulonephritis [2]. Together, these cause about 75% of all adult cases. Historically, kidney disease has been classified according to the part of the kidney anatomy involved[3].

Hemodialysis (HD) is a process of purifying the blood of a person whose kidneys are not working normally. This type of dialysis achieves the extracorporeal removal of waste products such as creatinine and urea and free water from the blood when the kidneys are in a state of kidney failure. Hemodialysis is one of three renal replacement therapies (the other two being kidney transplant and peritoneal dialysis) [4].

Hemodialysis patients are particularly predisposed to infections. It seems that the HD procedure per se as well as disturbances in both innate and adaptive immunity significantly contribute to this susceptibility. Infections are the major cause of morbidity and the second cause of death following cardiovascular events in HD patients. Episodes of bacteremia and pneumonia account for the majority of severe infections in this population as well as blood transmitted viral infections [5].

Human parvovirus B19 has been linked to renal disease in three settings: As a cause of acute glomerulopathy and as a cause of anemia in ESRD and kidney transplantation [6]. Human parvovirus B19 is a non-enveloped icosahedral a single-stranded DNA virus that is a member of the parvoviridae family [7]. Immunosuppressed patients can fail to mount an effective immune response to B19, resulting in prolonged or persistent viremia [8]. The mode of transmission for human parvovirus B19 in normal hosts is through the respiratory tract. Other possible routes of transmission suggested include donor graft [9]. The transmission rate is about 50 percent for those living with infected persons [10]

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Most persons with human parvovirus B19 infection are asymptomatic or exhibit mild, non-specific, cold-like symptoms that are never linked to the virus[11]. There are two types of diagnostic tests to confirm parvovirus B19 infection: B19-specific antibody testing and viral DNA testing. Giant pronormoblasts on a peripheral blood smear or in a bone marrow aspirate are suggestive of parvovirus B19 infection but are not diagnostic [12].

The present study is designed to determine the infection rate of human parvovirus B19 among dialyzed patients and to investigate any related aspects with different parameters such as age, gender, residence, marital status, occupation, duration of diyalsis, family history, history of blood transfusion and having knowledge about the disease and others.

Patients and methods

Cross-sectional research study includes 51 patients with hemodiyalasis, who have been selected randomly from Ibn-Sina hemodialysis center in Baqubah Teaching Hospital during the period from September 2016 till February 2017.

Full information had been taken directly from the patients and the information has been arranged in an informative formula sheet which includes: age, gender, residence, marital status, education levels, occupation, history of blood transfusion and having knowledge about the disease and others.

Two milliliters of blood samples were collected via vein puncture and used to evaluate anti-human parvovirus B19 IgM (Ridascreen-K6031-Germany) and anti-human parvovirus B19 IgG (Ridascreen -K6021- Germany) using enzyme-linked immunosorbent assay.

Statistical analysis

All date were analyzed with the statistical package for the social sciences (SPSS), version 13, the researchers performed the Chi-square analysis and p-value \Box 0.05 and it was considered statistically significant.

II. Results

Table (1) summarizes the demographic characteristic of 51 hemodiyalasis patients. There was a higher proportion of males 32 (62.74%) than females 19(37.25%) but statistically not significant. The age range was 15 to 70 years with mean age 48.5 years. Most of the patients were in age group 19 (73.25%); 22(43.13%) were living in urban area and 29(56.56%) were living in rural area. Forty-nine patients (96.07%) were married while only 2(3.92%) patients were single. Regarding educational background, 10 patients (19.60%) were illiterate, 15 patients (29.41%) had primary school, 19 patients (37.25%) had secondary school and 7 patients (13.72%) had an academic degree. Significant differences were noticed with age, marital status, occupation and duration of diyalsis though there were no -significant difference with other factors.

Parameters	No. (%)	P value
Gender type		0.069
Male	32 (62.74%)	
Female	19(37.25%)	
Age groups		0.000*
5-20 years	1(1.96%)	
21-36 years	10(19.60%)	
37-52 years	19(73.25%)	
53-70 years	21(41.10%)	
Residence		0.372
Urban	22(43.13%)	
Rural	29(56.56%)	
Marital status		0.000*
Single	2(3.92%)	
Married	49(96.07%)	
Education		0.084
Illiterate	10(19.60%)	
primary school	15(29.41%)	
Secondary school	19(37.25%)	
High education	7(13.72%)	
Duration of diyalsis		0.036*
Less than 1 year	33(64.70%)	
More than 1 year	18(35.29%)	

Table (1): Socio-demographic characteristics among studied group.

*Significant at p < 0.05

The risk factor amongst the studied patients, 44 cases (86.27%) presented a family history, while 15/51(29.41%) gave a blood transfusion, only 3 cases (5.88%) had a kidney transplant, related with other disease were 15(29.41%) and 12 cases (88.23%) gave a smoking habit as shown in table 2.

Risk Factor	Yes	No	P value
	No. (%)	No. (%)	
Family History	44(86.27%)	7(13.72%)	0.000*
Blood transfusion	15(29.41%)	36(70.58%)	0.003*
Kidney transplant	3(5.88%)	48(94.11%)	0.000*
Associated with other disease	15(29.41%)	36(70.58%)	0.003*
Smoking habit	45(88.23%)	6(11.76%)	0.000*
* Significant at p < 0.05			

Table (2): Distribution of haemodialysis patients according to risk factor.

The results of enzyme linked immunosorbent assay revealed that 4 serum patients (7.84%) were positive to anti-human parvovirus B19 IgM and 47 serum patients (92.15%) were positive to anti-human parvovirus B19 IgG. Statistical analysis shows significant difference as shown in table 3.

Parameters	ELISA result No. (%)	No No. (%)	p-value
	Positive%	4(7.84%)	Chi-square- 69.2031
IgM	Negative%	47(92.15 %)	Significant at p < 0.05
	Total	51(99.99%)	
	Positive%	46 (90.19%)	
IgG	Negative%	5(9.80%)	
	Total	51(99.99%)	

 Table (3): Comparison between human parvovirus B19 IgM and IgG in studied group.

The positive anti-parvovirus B19 IgM among the studied patients was 4 cases. The relation between positive anti-parvovirus B19 IgM and different socidemographic characters which demonstrated significant differences with marital status, occupation and duration of diyalsis while non significant occur with others as shown in table 4.

Table (4): Comparison between human parvovirus B19 IgM positive and negative result in studied group.

Parameters	Positive No. (%)	Negative No. (%)	p-value	
Gender type				
Male	2(6.25%)	30(93.75%)	0.582	
Female	2(10.52%)	17(89.74%)	Non-significant- p < .05	
Age groups				
5-20 years	0	1(100%)	0.281	
21-36 years	2(20%)	8(80%)	Non-significant - p < .05	
37-52 years	0	19(100%)	1	
53-70 years	2(9.52%)	19(90.47%	1	
Residence	1	1		
Urban	1(4.45%)	21(95.45%)	P-0.445 Non-significant - p < .05	
Rural	3(10.34%)	26(89.65%)	Ivon-significant - p < .05	
Marital status	1			
Single	1(50%)	1(50%)	0.582 Similaritatin < 0.05	
Married	3(6.12%)	46(93.87%)	Significant at p < 0.05	
Education	Education			
Illiterate	2(20%)	8(80%)	0.416	
primary school	1(6.66%)	14(93.33%)	Non-significant - p < .05	
Secondary school	1(5.26%)	18(94.37%)	1	
High education	0	7(13.72%)	-	
Duration of diyalsis		0.653		
Less than 1 year	3(9.09%)	30(90.90%)	Significant at p < 0.05	
More than 1 year	1(5.55%)	17(94.44%)	1	

Regarding risk factor of all human parvovirus B19 IgM positive and negative results were distributed in table (5). Two (28.57%) participants were positive and had family history whereas all positive cases had not a history of blood transfusion and kidney transplant, 1(6.66%) was tested with other disease while smoking habit occurring in all positive cases.

Risk Factor	Yes	No	p-value
	Number(%)	Number(%)	-
Family History			
Positive	2(4.54%)	2(28.57%)	0.028
Negative	42(95.45%)	5(71.42%)	
Total	44(86.27%)	7(13.72%)	
Blood transfusion			
Positive	0	4(100%)	
Negative	15	32	0.179
Total	15(29.41%)	36(70.58%)	
Kidney transplant			
Positive	0	4(8.33%)	0.602
Negative	3(5.88%)	44(91.66%)	
Total	3(5.88%)	48(94.11%	
Associated with other disease			
Positive	1(6.66%)	3(8.33%)	0.840
Negative	14(93.33%)	33(91.66%)	
Total	15(29.41%)	36(70.58%)	
Smoking habit	· · · · ·		
Positive	4(8.88%)	0	0.447
Negative	41(91.11%)	6	
Total	45(88.23%)	6(11.76%)	

Table (5): Distribution of patients with anti-human	parvovirus B19 infection according to result of IgM.
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Some important patient characteristics such as age, gender, residence, marital status, educational level, occupation and duration of diyalsis were compared among the seropositive groups. As demonstrated in the table 6, there was no significant difference with all parameters.

 Table (6): Comparison between anti-human parvovirus B19 IgG positive and negative result in studied group.

Parameters	Positive No. (%)	Negative No. (%)	p-value	
Gender type				
Male	29(90.62%)	3(9.37%)	0.89	
Female	17(89.47%)	2(10.52%)	Non-significant - p < .05	
Age groups				
5-20 years	1(100%)	0		
21-36 years	9(90%)	1(10%)	0.794	
37-52 years	18(94.73%)	1(5.26%)	Non-significant - p < .05	
53-70 years	18(85.71%)	3(23.80%)		
Residence				
Urban	19(86.36%)	3(13.63%)	0.42	
Rural	27(93.10%)	2(6.89%)	Non-significant - p < .05	
Marital status	Marital status			
Single	2(100%)	0	0.634	
Married	44(89.79%)	5(10.20%)	Non-significant - p < .05	
Education				
Illiterate	7(70%)	3(30%)		
primary school	15(100%)	0	0.011	
Secondary school	19(100%)	0	Non-significant - p < .05	
High education	5(71.42%)	2(28.57%)		
Duration of diyalsis		0.45		
Less than 1 year	29(78.78%)	4(12.12%)	Non-significant - p < .05	
More than 1 year	17(94.44%)	1(5.55%)		

According to distribution of anti-human parvovirus B19 IgG, positive results were with risk factor. Majority of the participants, 43(97.72%) had family history and 44(97.77%) smoking habit. Only 10(66.66%) with blood transfusion and associated with other disease while low frequency of positive cases were registered among patients with kidney transplant. Statistical analysis showed significant differences as summarized in table (7).

Risk Factor	Yes	No	p-value
	Number(%)	Number(%)	•
Family History			
Positive	43(97.72%)	3(42.85%)	0.000
Negative	1(2.27%)	4(57.14%)	
Total	44	7	
Blood transfusion			
Positive	10(66.66%)	36(100%)	0.000
Negative	5(33.33%)	0	
Total	15	36	
Kidney transplant			
Positive	2(66.66%)	44(91.66%)	0.158
Negative	1(33.33%)	4(8.33%)	
Total	3	48	
Associated with other			
disease	10(66.66%)	36(100%)	0.000
Positive	5(33.33%)	0	
Negative	15	36	
Total			
Smoking habit			
Positive	44(97.77%)	2(33.33%)	0.000
Negative	1(2.22%)	4(66.33%)	
Total	45	6	

Table (7): Distribution of patients with human parvovirus B19 infection according to result of IgG.

III. Discussion

The current study is done due to few studies which had already been carried out on the B19 frequency in dialysis patients. So the result of this study showed that infection rates of human parvovirus B19 in Baqubah city of Iraq, 4(7.84%) were positive to human parvovirus B19 IgM and 46 (90.19%) were positive to human parvovirus B19 IgG. This result conforms with study carried out by Sharif *et al.*(2016) who found the same result among Iranian hemodiaylasis and peritoneal dialysis patients [13].

The prevalence of immunoglobulin G (IgG) antibodies directed against human parvovirus B19 ranges from 15 to 60% in children 6 to 19 years old: and from 30 to 60% in adults and is more than 85% in the geriatric population [14][15][16]. Furthermore, the result of present study conforms with many investigations which have been carried out in different parts of the world [17][18][19].

However, patients with renal failure on dialysis have disruptions in their immune system due to the immunosuppressive effects of uremia, deficient erythropoietin production, and significantly decreased erythrocyte survival [9][10]. Therefore, dialysis patients reveal an increasing susceptibility to human parvovirus B19 infection.

Human parvovirus B19 infection was higher in age group of 53-70 years than others age groups as registered, this result is relatively comparable with that reported by other studies [20][21]. This may be relevant to the rising incidence with age and may be explained by the thymus function which is known to decline with age. The thymus reaches its maximal size at puberty and then atrophies, with a significant decrease in both cortical and medullary cells and an increase in the total fat content of the organ. Whereas the average weight of the thymus is 30 grams in human infants, its age-dependent involution leaves an organ with an average weight of only 3 grams in the elderly [22]. Moreover, it may be related with decline in the number of NK, which plays an important role in early natural surveillance against cancer and infectious disease, a progressive age-related shift in the circulating lymphocyte population from conventional T cells to NK cells [23].

Approximately half of the adult population has been exposed to this agent and acquired immunity between the ages of five and 19 years, with seroprevalence increasing with age [24][25].

Chronic kidney disease infection occurred in all age groups and genders with no significant difference among these groups. This indicates that final stage of renal failure affects the economically productive age group which is unlike the situation in many developed countries in which the mean age of ESRF patients is generally over 60 years. This result is in agreement with Amin *et al* (2015). Who showed that the mean age of ESRF patients is 49 ± 15.8 years and 47.9 % of ESRF patients [26].

The result of current study revealed that most infection is nearly three times more common in males than in females but without significant difference.

The result of current study demonstrated that no significant difference in gender, educational level, history of HD, this disagrees with result of study done by Khameneh et al. (2014) who found that there was no significant difference in age, sex, educational status, history of blood transfusion, history of HD and immunosuppressive therapy among kidney transplant recipients with positive human parvovirus B19 [27].

In conclusion, the seroprevalence of human parvovirus B19 was relatively high in dialyzed patients in Bagubah-Iraq, and should be investigated after exclusion of other common causes. Further studies are needed to assess the role of other viruses in heamodiyalasis patients.

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References

- National Kidney Foundation. K/DOQI clinical practice guidelines for chronic kidney disease. Retrieved 2008-06-29. [1]. [2].
 - "United States Renal Data System (USRDS)".
- [3]. Rahman, Mahboob; Smith, Michael C. Chronic renal insufficiency: A diagnostic and therapeutic approach. Archives of Internal medicine.1998; 158: 1743-52.
- National Kidney and urologic disease information clearinghouse guidance kidney failure. Choosing a treatment that's right for you. [4]. [5]. Eleftheriadis T, Liakopoulos V, Leivaditis K, Antoniadi G, Stefanidis I. Infections in hemodialysis: a concise review. Part II: blood
- transmitted viral infections. Hippokratia. 2011, 15, 2: 120-126
- Waldman M, Kopp JB. Parvovirus B19 and the kidney. Clin J Am Soc Nephrol. 2007; 2 Suppl 1: S47-56. [6].
- Yango A, Morrissey P, Gohh R, Wahbeh A. Donor-transmitted parvovirus infection in a kidney transplant recipient presenting as [7]. pancytopenia and allograft dysfunction. Transpl Infect Dis 2002;4:163-6.
- [8]. Waldman M, Kopp JB. Parvovirus-B19-associated complications in renal transplant recipients. Nat Clin Pract Nephrol. 2007; 3(10):540-50.
- [9]. Murer L, Zacchello G, Bianchi D, Dall'Amico R, Montini G, Andreetta B, Perini M, Dossi EC, Zanon G, Zacchello F. Thrombotic microangiopathy associated with parvovirus B 19 infection after renal transplantation. J Am Soc Nephrol 2000; 11:1132-7.
- [10]. Parvovirus B19 (erythema infectiosum, fifth disease). In: Red Book 2006: Report of the Committee on Infectious Diseases. 27th ed. Washington, D.C.: American Academy of Pediatrics, 2006:484-7.
- Heegaard ED, Brown KE. Human parvovirus B19. Clin Microbiol Rev. 2002;15:485-505. [11].
- Cohen BJ, Buckley MM. The prevalence of antibody to human parvovirus B19 in England and Wales. J Med Microbiol. [12]. 1988;25:151-3.
- [13]. Sharif A, Arezoo A, Ali AV, Mohammad B, Mohammad RS, Effat R, Davood K, Monireh K, Anahita B, Amitis R. Frequency and Genotype of Human Parvovirus B19 among Iranian Hemodialysis and Peritoneal Dialysis Patients. Intervirology 2016;59:179-185.
- [14]. parvovirus B19 Anderson LL Role of in human disease. Pediatr. Infect. Dis. J. 1987; 6:711-718.
- [15]. Cohen BJ, Buckley MM. The prevalence of antibody to human parvovirus B19 in England and Wales. J. Med. Microbiol.1988; 25:151-153
- Tsujimura M, Matsushita K, Shiraki H, Sato H, Okochi K, Maeda Y. Human parvovirus B19 infection in blood donors. Vox [16]. Sang.1995; 69:206-212.
- Heegaard ED, Jensen IP, Christensen J. Novel PCR assay for differential detection and screening of erythrovirus B19 and [17]. erythrovirus V9. J Med Virol. 2000; 65: 362-367.
- [18]. Jordan J, Tiangco B, Kiss J, Koch W. Human parvovirus B19: prevalence of viral DNA in volunteer blood donors and clinical outcomes of transfusion recipients. Vox Sang. 1998; 75: 97-102.
- McOmish F, Yap PL, Jordan A, Hart H, Cohen BJ, Simmonds P. Detection of parvovirus B19 in donated blood: a model system for [19]. screening by polymerase chain reaction. J Clin Microbiol. 1993; 31:323-328.
- [20]. Cohen BJ, Buckley MM. The prevalence of antibody to human parvovirus B19 in England and Wales. J Med Microbial. 1988; 21-25.
- [21]. Kerr S, O'keeffe G, Kilty C. Undenatured parvovirus B19 antigens are essential for the accurate detection of parvovirus B19 IgG. J Med Virol. 1999;57:179-185.
- Kindt TJ, Goldsby RA, Osborne BA, Kuby J. Kuby immunology 6th Ed. WH. Freeman and company. New York. 2007.p41. [22].
- [23]. Ravaglia G, Forti P, Maioli F, Bastagli L, Facchini A, Mariani E, Savarino L, Sassi S, Cucinotta D, Lenaz G. Effect of micronutrient status on natural killer cell immune function in healthy free-living subject aged. Am Soci Clin Nutri 2000; 71(2):590-598.
- [24]. Centers for Disease Control. Risks associated with human parvovirus B19 infection. MMWR 1989;38:81-8.
- Torok TJ. Parvovirus B19 and human disease. Adv Intern Med 1992;37:431-55. [25].
- Banaga ASI, Elaf BM, Rania MS, Diana ES, Sara BE, Mohamed OK, Rasha A Babiker, Khalifa E, Mamoun MH. Causes of end [26]. stage renal failure among haemodialysis patients in Khartoum State/Sudan. BMC Research Notes. 2015; 8:502.
- Khameneh ZR, Nariman S, Vahid S, Nazafarin G. The seroprevalence of Parvovirus B19 among kidney transplant recipients: [27]. Asingle center study. Saudi J Kidney Dis Transpl. 2014; 25(1):16-21.

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