

Sero-Epidemiology of Human Parvo B19 Virus among Pregnant Women Attending Ante-Natal Federal Medical Center Keffi, Nasarawa State, Nigeria

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Abstract Trans placental transmission of parvovirus B19 during pregnancy has been reported as one of the leading causes of non-immune fetal hydrops, spontaneous abortion or intra-uterine fetal death. We therefore set out to determine the presence of parvovirus B19 (IgG and IgM) antibodies and their risk factors among pregnant women attending ante-natal clinic, at Federal Medical Center, Keffi, Nasarawa State, Nigeria. A total of 273 pregnant women (Age range 15-40 years) participated after informed consent and clearance from the research ethics committee. Each filled a self-administered questionnaire and donated blood samples between January and August 2012. A recombinant parvovirus B19 IgM and IgG ELISA kit from DeMediTec (USA) was used for the assays. Of the 273 participants, 36 (13.2 %) had IgM antibodies, while 75 (27.5%) had IgG antibodies with those aged 15-20 and 36-40 years having the highest incidence and prevalence rates. Being a house wife, having more children and histories of blood transfusion, miscarriage and still birth were associated with infection ($P < 0.05$). The study reveals high incidence and prevalence of parvovirus B19 infection among the pregnant women. The presence of parvovirus B19 IgM antibody particularly in the first and second trimesters of pregnancy poses the danger of likely transplacental transmission leading to spontaneous abortion and intra-uterine fetal death.

Keywords Parvo B19, Pregnant, Risk, Nasarawa State, Trans-placental

1. Introduction

Human parvovirus B19 is the only known parvovirus pathogenic to human[1,2]. The infection during pregnancy may account for thousands of incidences of fetal loss per annum in Europe, North America and beyond[3].

Transplacental transmission of parvovirus B19 during pregnancy is one of the leading causes of non-immune fetal hydrops, spontaneous abortion or intra uterine fetal death [4,5]. The disease which was first discovered in 1975[6] was first associated with disease in 1981, when it was linked to aplastic crisis, fifth disease, spontaneous abortion and some form of acute arthritis[7]. It also has a 33% vertical transmission rate with a 2-3 incidences in all spontaneous abortion[8].

B19 may also stimulate a cellular process initiating apoptosis[9]. This may account for the minimal inflammatory response noted in tissues infected with B19[10].The virus infection of megakaryocytes, placental

cells, fetal liver cells and myocardial cells results in anemia, which can lead to high output congestive heart failure, causing fetal circulatory compromise. The risk of fetal demise is highest in the first trimester and is thought to be as high as 10% in women who are infected prior to 20 weeks of gestation[11]. The P blood group antigen, which serves as a receptor for B19, has been detected on cells of the villous trophoblast of placental tissues in varying amount during the course of pregnancy. And in the first trimester of pregnancy, the level of the P-antigen is very high. This high level of globoside receptor on placental cells in early pregnancy may act as a pathway for B19 to be transmitted from the mother to the fetus where by the virus can then infect erythroid progenitor cells for replication[12]. There are limited data on the incidence and prevalence of parvovirus B19 in Nigeria in spite of the importance of such data in health policy formulation. We therefore, undertook a survey to establish the incidence, prevalence and risk factors of this agent among pregnant women attending Plateau specialist hospital in Jos.

2. Materials and Methods

2.1. Study Area

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The study was carried out at Federal Medical Center, Keffi, a city in the middle belt region of Nigeria. It is located at 9°56'N 8°53'E 9.933°N 8.883°E, with an altitude of 4,062 feet (1,217m) above sea level. It enjoys a more temperate climate than the rest of Nigeria and has an average monthly temperature range from 70° to 77°F or 21° to 25°C[13].

2.2. Study Population

This consisted of pregnant women attending ante-natal clinic in Federal Medical Center, Keffi, between January and August, 2012. The hospital has an average of 70 pregnant women attending ante-natal clinic weekly and population of two hundred and seventy three (273) subjects participated in this study.

2.3. Ethical Considerations

Clearance from the health research ethical committee of Federal Medical Center, Keffi was obtained in accordance with the code of ethics for biomedical research involving human subjects. Signed informed consent was also obtained from each of the subjects.

2.4. Exclusion and Inclusion Criteria

The study inclusion criteria were being pregnant, ante-natal clinic attendance records and signed informed consent; while the exclusion criteria included absence of pregnancy, age below 15 and above 40 years and failure to sign a consent form.

2.5. Specimen Collection

After collection of the signed informed consent form, and the self-administered questionnaires with the relevant personal information, 5mls of blood sample from each subject was aseptically collected by venipuncture of the cubital vein using disposable needles and syringes. The samples were collected between January and August, 2012, and then taken to Innovative Biotech Virology Laboratory where the sera were separated, put into sample vials and stored at -20°C until assayed.

2.6. Assaying Technique

The **DeMeditec** parvovirus B19 IgM and IgG (Recombinant) Enzyme immunoassay kit [Lot; 123G/K038] USA was used for the in-vitro qualitative and semi-quantitative determination of immunoglobulin class antibodies assay for parvovirus B19 in serum, according to the manufacturer's instructions.

2.7. Interpretation of Results

Incidence was defined as the presence of IgM antibodies to parvovirus B19 in the tested subjects, while prevalence was defined as the presence of parvovirus B19 IgG antibodies. The presences of IgM or IgG antibodies were determined by comparing the absorbance value of each sample to that of the cut-off value. Specimens with absorbance values less than the cut-off values were recorded

as negative while those above are recorded as positive according to the manufacturer's instructions.

2.8. Statistical Analysis

Pearson chi-squares were calculated and compared with values on a chi-square table at 95% confidence interval. P values ≤ 0.05 were considered statistically significant.

3. Results

Blood samples were collected from 273 pregnant women. The serum samples were analyzed for parvovirus B19 (IgM and IgG) antibodies. Of the 273 samples assayed, 36(13.2%) were positive for Parvovirus B19 IgM and 75(27.5%) for B19 IgG.

Table 1 shows the distribution of parvovirus B19 antibodies according to age. The highest IgM antibodies (16.7%) and IgG antibodies (37.5%) antibodies were observed in the age groups 15-20 and 36-40 years respectively, while the lowest Parvovirus B19 IgM antibodies and IgG antibodies were observed in the age group 36-40 and 15-20 years respectively ($P > 0.05$).

Table 1. Age Distribution of Parvovirus B19 Antibodies Among pregnant Women Attending Ante-natal Federal Medical Center, Keffi Nigeria

Age (Year)	No Tested	No Positive	IgM antibodies	IgG antibodies
15-20	54	18(16.7%)	9(16.7%)	9(16.7%)
21-25	78	33(42.3%)	12(15.4%)	21(26.9%)
26-30	69	30(43.5%)	9(13.05)	21(30.4%)
31-35	48	21(43.8%)	6(12.5%)	15(31.3%)
36-40	24	9(37.5%)	0 (0.0%)	9(37.5)
Total	273	111(40.7%)	36(13.2%)	75(27.5%)
		P > 0.05		

Table 2 shows the distribution of parvovirus B19 antibodies according to gestational age. The highest IgM antibodies (13.3%) and IgG antibodies (31.1%) were observed among pregnant women in their second trimester, followed by those in their first trimester. Those in their third trimester gave the least prevalence (For IgM, $P > 0.05$ while for Ig G, $P = 0.05$).

Table 3 shows the distribution of parvovirus B19 antibodies according to history of pregnancy complications (miscarriage and stillbirth). The highest IgM antibodies (20.0%) and IgG antibodies (40.0%) were observed among those who have had history of both miscarriage and still birth. The lowest IgM antibodies (10.5%) and IgG antibodies (22.8%) were observed among pregnant women with no history of pregnancy complications ($P > 0.05$).

Table 3 shows the distribution of parvovirus B19 antibodies according to the number of children in the household. The highest IgM antibodies (19.5%) and IgG antibodies (42.9%) were observed among those with 5 or more children. The lowest IgM antibodies (10.6%) and IgG antibodies (19.2%) were observed among those with 0-2 children (For IgM, $P > 0.05$, while for Ig G, $P < 0.05$).

Table 2. Distribution of Parvovirus B19 IgM antibodies by Risk Factors Among Pregnant Women Attending Ante-natal Federal Medical Center, Keffi Nigeria

Risk factor Value	No Tested	No Positive (%)	X ²	P
Gestational Age				
First Trimester	114	15 (13.2%)		
Second Trimester	135	18 (13.3%)	0.03	>0.05
Third Trimester	24	3 (12.5%)		
History of Complication				
Miscarriage	54	9 (16.7%)		
Still Birth	33	6 (18.2%)	5.93	>0.05
Both	15	3 (20.0%)		
None	171	18 (10.5%)		
No of Children in Household				
0-2	141	15 (10.6%)		
3-4	69	9 (13.0%)	5.02	>0.05
5 or more	63	12 (19.0%)		
History of Blood Transfusion				
Yes	39	9 (23.1%)	5.57	<0.05
No	234	27 (11.5%)		
Occupation				
Students	12	3 (25.0%)		
Business	135	15 (11.1%)		
Civil Servants	81	6 (7.4%)	19.99	<0.05
House wife	45	12 (26.7%)		

Table 3. Distribution of Parvovirus B19 IgG antibodies by Risk Factors Among Pregnant Women Attending Ante-natal Federal Medical Center, Keffi Nigeria

Risk factor Value	No Tested	No Positive (%)	X ²	P
Gestational Age				
First Trimester	114	30 (26.3%)		
Second Trimester	135	42 (31.1%)	5.99	=0.05
Third Trimester	24	3 (12.5%)		
History of Complication				
Miscarriage	54	21 (38.9%)		
Still Birth	33	9 (27.3%)	4.89	>0.05
Both	15	6 (40.0%)		
None	171	39 (22.8%)		
No of Children in Household				
0-2	141	27 (19.1%)		
3-4	69	21 (30.4%)	10.65	<0.05
5 or more	63	27 (42.9%)		
History of Blood Transfusion				
Yes	39	12 (30.8%)	0.2	>0.05
No	234	63 (26.9%)		
Occupation				
Students	12	3 (25.0%)		
Business	135	33 (24.4%)		
Civil Servants	81	21 (25.9%)	3.86	>0.05
House wife	45	18 (40.0%)		

Table 3 shows the distribution of B19 antibodies according to history of blood transfusion. The highest IgM antibodies (23.1%) and IgG antibodies (30.8%) were

observed among those who has been transfused with blood at one occasion or another, while the lowest IgM antibodies (26.9%) and Ig G antibodies (11.5%) respectively were found among those who have not received blood transfusion (For IgM, P<0.05, while for IgG, P>0.05).

Table 3 shows the distribution of parvovirus B19 antibodies according to occupation of the subjects. The highest IgM antibodies (26.7%) and Ig G antibodies (40.0%) were observed among pregnant women who are housewives. The lowest IgM antibodies (7.4%) and IgG antibodies (24.4%) were observed among those who are civil servants and business women respectively (For IgM, P<0.05, while for IgG, P>0.05).

4. Discussion

This study identifies an incidence (13.2%) and prevalence (27.5%) of parvovirus B19 at the Federal Medical Center, Keffi, Nasarawa State which has an average of 70 pregnant women attending ante-natal clinic weekly. The incidence of 13.2% and prevalence of 27.5% agrees closely with the reports of Keikha et al in Iran who reported 10.3% incidence and 21.8% prevalence in pregnancy [14]. It also accords with the 30-50% range reported in India by Kaur and Basu, and also with 30-60% prevalence range reported in United States by the American Academy of Pediatrics Committee on infectious Diseases, [1,15]. Our finding is however, lower than the 63-71% prevalence range reported in Dublin, Ireland, by Eis-Hubinger, and the 70% reported in United Kingdom and Finland among adults by Massong et al [16,17]. It is however very high compare to the 2.0% incidence among blood donors in North Africa (Tunisian) and Western Europe (Belgian) reported by Letalef et al [18]. There is no clear explanation for this difference.

The incidence of parvovirus according to age groups was observed to decrease simultaneously with increase in ages while the prevalence was found to increase with increase in age groupings. This trend could be due to chances of exposure to the infection with age. A report of increase seroprevalence with age and greater than 70% in adult's populations was reported by Kerr et al, [19].

The outcome of the highest incidence (13.3%) and prevalence (31.1%) in the second trimester in the gestational age of pregnancy could probably be explained as susceptibility due to hormonal changes with progress in pregnancy. Since the presence of IgM antibodies signifies recent infection, it therefore poses the danger of transplacental transmission with the risk of fetal anaemia and demise highest in the first and second trimester especially for women infected prior to 20 weeks of gestation. A similar report was made by Markenson, and Enders et al [11,24].

The observation from the history of pregnancy complications, in which those with both miscarriage and still birth experience a higher parvovirus B19 incidence and prevalence when compared with those with no such history, further emphasizes the possible outcome of the effect of

parvovirus B19 infection during pregnancy.

The outcome of the incidence and prevalence of parvovirus B19 antibodies among those with more number of children as against those with few or no children, suggests the involvement of contact with more number of children as a possible source of spread of the infection. However, previous reports identified children are the main sources of transmission, and outbreaks can persist for months in schools and day care centers, due to the relatively large number of seronegative children and close contact of children within this environment,[20,21]. A similar result was also reported by Xu *et al.*, and Laura and Caraciolo[2,10]. The findings from our study was statistically significant at $P < 0.05$.

The incidence and prevalence obtained among those who have had blood transfusion as against those who have not also emphasizes the possible implication of blood transfusion in the transmission of the infection, especially if there is contamination. This study identifies the incidence to be statistically significant at $P < 0.05$. A report by Corcoran and Doyle,[3], observed that despite the fact that parvovirus B19 infection can be transmitted via contaminated blood products; there is presently no strict regulatory prerequisite governing B19 contamination of pooled plasma or blood products prior to products release.

The prevalence of B19 antibodies among housewives when considering the occupation of the pregnant women studied, may implicate household contact in the transmission of the infection. Previous studies reported that, the infection is commonly passed through respiratory tract secretions from close personal contact[2]. Statistically, there was no significant difference between B19 incidence and occupation during pregnancy but there was significant difference between B19 prevalence and occupation. A report of no significant differences found in maternal infection from occupational exposures was reported by Harger *et al*[22], and Valeur *et al*[23].

The high incidence and prevalence of infection with B19 virus among the pregnant women and the known dangers posed in pregnancy raise transfusion policy questions in Nigeria. Like other blood-transmissible viral infections, routine screening for B19 virus infection before transfusion should, as a matter of policy, be made compulsory in Nigeria.

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