# **Original Research Article**

# Obstetric and new born outcome in HIV infected pregnant women: a prospective cohort study in Bangalore Medical College Hospitals, India

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# ABSTRACT

**Background:** The effect of HIV on obstetric complications is known to vary across regions of world. The variation may be due to HIV infection per se or it may be due to complex interaction of related medical and social conditions that affect pregnancy. Incidence of these obstetric complications in HIV infected pregnant women is not well reported in India. So, this prospective cohort study was carried to observe the demographics and incidence of obstetrics complications like abortions, still birth, premature rupture of membranes, preterm delivery, opportunistic infections in HIV infected pregnant women.

**Methods:** All pregnant women who were screened positive for HIV test, irrespective of their gestational age were included in the study. Apart from routine obstetric care, CD4 cell count was carried out. The patients were followed up till term, delivery and up to 6-week postpartum period. Obstetric outcomes like incidence of abortion, intrauterine death, preterm delivery and premature rupture of membrane were noted. Baby's birth weight and the incidence of maternal opportunistic infection were noted and correlated with maternal CD4 cell count.

**Results:** Maternal HIV infection is associated pre-term labour in 34.5%, PROM in 30%, low birth weight in 52.3%. Increased incidences of these adverse outcomes have an inverse relationship with CD4 count.

**Conclusions:** Maternal HIV infection is associated with increased incidence of adverse obstetric outcome and opportunistic infection, and their incidences increase with lower CD4 count.

Keywords: CD4 T cell, Low birth weight, Maternal HIV infection, Obstetric complications, Preterm labour, PROM

### **INTRODUCTION**

In 2009 it was estimated that 2.4 million people were living with HIV in India, which equates to a prevalence of 0.3%.<sup>1</sup> India had the third highest number of people living with HIV. Women are among the fastest growing populations getting infected with HIV and AIDS and most of them are in childbearing age.<sup>1</sup> With the early detection and increasing use of antiretroviral therapy, more number of HIV positive women are entering family cycle and are delivering. HIV infection has been reported to have little effect on pregnancy outcome or complications in the developed world.<sup>2,3</sup> Adverse pregnancy outcomes have, however, been reported more commonly in a number of African studies.<sup>4-7</sup> It is often difficult to determine the relative contribution of HIV infection, drug abuse and inadequate antenatal care to adverse outcomes in these women.<sup>8,9</sup> Thus the adverse pregnancy outcome in HIV may directly by HIV infection or it may be a marker of a complex interaction of related medical and social conditions in that community. But the effects of HIV infection on maternal morbidity, obstetric complications are not well reported in India. So, this prospective cohort study was carried out to observe the demographics and obstetrics complications in HIV infected pregnant women.

### **METHODS**

This hospital based prospective cohort study was carried out after taking approval from the institutional ethics committee. The study was carried out in the antenatal clinics of the two tertiary referral hospitals of Bangalore medical college and research institute; namely Vani Vilas hospital and Bowring and Lady Curzon Hospital, Bangalore. All the pregnant women attending the antenatal clinic at the above-mentioned hospitals from November 2009 to November 2010 were screened for HIV infection. Diagnosis of HIV infection was made as per the NACO guidelines, using rapid ELISA tests.<sup>10</sup> Those patients, who were found to be seropositive for HIV, underwent post-test counselling.

#### Inclusion criteria

All pregnant women who were screened positive for HIV test, irrespective of their gestational age and irrespective of whether they received any antiretroviral treatment (ART), were included in this study.

#### Exclusion criteria

Those HIV seropositive pregnant women who had other pre-existing chronic medical conditions like cardiac disease, diabetes mellitus, chronic renal failure and chronic liver disease were excluded, since these medical conditions are known to independently affect the pregnancy outcome.

Informed consent was obtained from all the HIV positive pregnant women to be part of this study. Pregnant women were given routine antenatal care including history, clinical examination. Routine antenatal investigations were done which included: complete haemogram, blood grouping and Rh typing, urine routine and microscopy, random blood sugar, serum for Hepatitis B antigen and VDRL.

The patients underwent obstetric ultrasound scan as needed. Once they were counselled about the HIV status, their sexual history, including history of multiple sexual partners by the woman herself and her partner was taken. Other high-risk factors for HIV transmission like blood transfusion or intravenous drug abuse were also noted.

Histories of other sexually transmitted diseases were noted and if present, they were treated. The parturient were explained the possibility of mother- to-child HIV transmission, the available methods to reduce the transmission rate and the facilities available for the follow up of the infants. If the pregnancy was within 20 weeks of gestation, then the option of termination of pregnancy was offered. Those desirous of continuing the pregnancy were followed up.

Based on the CD4 count and WHO stage of the disease, pregnant women were considered for ART at the earliest after detection. They were given ART as per NACO guideline 2007.<sup>11</sup> The neonate received a single dose of 2 mg per kg body weight of syrup Nevirepine within 72 hours of delivery and it was continued daily for 6 weeks. The parturient, who requested for elective lower segment caesarean section (LSCS) to reduce the risk of mother-tochild HIV transmission, were offered LSCS. Other parturient had either normal vaginal delivery or LSCS as per the obstetric indication. All women received 5-day course of antibiotics with first generation cephalosporin, irrespective of whether it was vaginal delivery or LSCS.

Demographic data like age of the parturient, their religion, educational status, occupation and occupation of their partner were collected. Obstetric data like number of years of married life, obstetric score, gestational age at the time of delivery, status of membranes at onset of labour, mode of delivery, incidence of postpartum complications like retained placenta, puerperal pyrexia, urinary infection, puerperal sepsis and burst abdomen were noted.

New born birth weight was also noted. Authors wanted to assess whether the severity of HIV infection, as indicated by CD 4 count, affects the incidence of obstetric complications. Hence the patients were categorised based on their CD4 cell count in to those with CD4 cell count more than 350 and those with count less than 350.

### Statistical analysis

SPSS version 16 was used for analysing the data. Before analysis, the data was checked for consistency in range. Categorical data like obstetric complications, foetal complications, opportunistic infections were analysed using Chi square test and data is expressed as frequency and percentages. Continuous data like birth weight was analysed by independent sample t-test and expressed as mean and standard deviation.

# RESULTS

A total 120 HIV seropositive pregnant women were followed up prospectively. The pregnant women were aged between 17 to 45 years, with mean age of 24.1 years (Table 1). The majority were detected to be seropositive within 4 years of married life.

(Table 2) Forty per cent of couple agreed to have polygamous relation and 17% of the women had previous history of blood transfusion. In remaining obvious risk factor for HIV were not available in history. Authors noted 92 (77%) of the husbands were HIV positive and 27 (23%) of the husbands were seronegative. In 1 patient, husband's HIV status could not be ascertained. Seventeen per cent of women had some other sexually transmitted diseases and were treated for the same.

#### Table 1: Demographic characters.

	Number of	Percentage			
	patients, n=120				
Age (years)					
15-19	5	4			
20-25	79	66			
26-30	29	24			
>30	7	6			
Religion					
Hindu	98	81.6			
Muslim	21	17.5			
Christian	1	0.8			
Patient education					
Illiterate	62	51.6			
Primary	39	32.5			
Secondary	17	14.1			
College	2	1.6			
Husband's education					
Illiterate	42	35			
Primary	39	32.5			
Secondary	23	19.1			
College	16	13.3			
<b>Occupation of patient</b>					
House wife	96	80			
Coolie and farmer	11 and 4	9.1 and 3.3			
Factory and shop worker	2 and 3	1.6 and 2.4			
Commercial sex worker	2	1.6			
Occupation of husbane	d				
Driver	33	27.5			
Coolie	28	23.3			
Factory	9	7.5			
Farmer	19	15.8			
Other services (painter, tailor, cook, mechanic)	19	15.8			

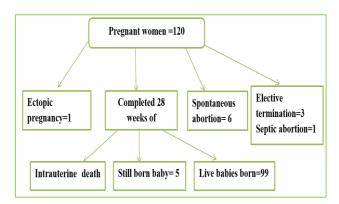
Pregnant women received ART based on their CD4 count and time of them first reporting to health care system. Twenty-six (21.6%) of the women received antiretroviral therapy during the pregnancy and 94 (78.4%) did not receive antiretroviral therapy. Thirteen of the 120 patients developed opportunistic infection. Opportunistic infections included; Tuberculosis in 3 patients, recurrent urinary infection in 3, lower respiratory infection with pneumonia in 5, MRSA pneumonia in 1 patient and opportunistic genital ulcer in 1 patient.

Of the 120 HIV positive women in the study, 6 patients had spontaneous abortion, 3 patients opted for termination of pregnancy, 1 patient had septic abortion and 1 patient had an ectopic pregnancy (Figure 1). Remaining 109 patients crossed 28 weeks of gestation and were followed till the delivery for maternal and new born outcome. Eighty-nine (82%) of women had vaginal deliveries and 20 (18%) had LSCS. The vaginal deliveries included 3 forceps assisted deliveries. All of the LSCS were for obstetric reasons and none of women asked for LSCS to reduce the HIV transmission per se. Puerperal pyrexia was seen in 11 (9%) women, wound infection was noted in 4, puerperal sepsis was seen in 2 women. Burst abdomen and retained placenta were seen in 1 woman each. There were no maternal deaths. The remaining women did not have any complication during puerperium.

#### Table 2: Obstetric parameter.

	Number of patients, n=120	Percentage			
Married life					
<2 years	41	34.1			
2-4 years	37	30.8			
>4 years	42	35			
Gravida	· · · · · ·				
1	47	39.1			
2	46	38.3			
3	22	1.1			
4 or more	5	4.1			
Multiple sexual partner					
By woman	5	4.1			
By husband	41	34.1			
Both	5	4.1			
None	69	57.5			
Membrane status at labour					
Intact	80	73.3			
PROM	29	26.6			
Term of delivery					
Preterm	37	34.5			
Term	64	58			
Post-term	8	7.5			

There were 5 cases (4.5%) of still birth and 5 cases (4.5%) with intrauterine death out of 109 births. Hence 99 babies were delivered. Baby weight ranged from 1.0 to 3.75 kg with the mean birth weight being 2.33 kg. Fifty seven of 99 babies (57.5%) had birth weight less than 2.5 kg and were classified as low birth weight babies.



# Figure 1: Flow diagram showing number of patients studied.

Of the 120-total parturient analysed, 39 patients had CD4 count <350 and 81 had their CD4 count >350 (Table 3). In order to assess whether the obstetric and foetal complications were related to the severity of HIV as depicted by CD4 count, these complications were analysed with low and high CD4 count as two categories. The complications like spontaneous abortions, ectopic pregnancy, preterm labour, PROM, puerperal pyrexia, postoperative wound infection, IUD and still birth were clubbed under obstetric complications. The chances of

obstetric complications were significantly higher in patients with low CD4 count than in patients with high CD4 count group (75% versus 54%). The incidence of opportunistic infection was also higher in low CD4 count group than in high CD4 count group. Intrauterine death, still birth and low birth weight, were classified as foetal complications. The incidence of foetal complications were high in those with the low CD4 counts than in high CD4 count group (61% versus 48%) But it was statistically not significant.

# Table 3: Obstetric complication for all pregnant women.

Complications	CD4 count ≤349 (n=39)	CD4 count ≥350 (n= 81)	Total	
Obstetric complications			N=120	
No	10 (25%)	37 (46%)	47	-0.020
Yes	30 (75%)	43 (54%)	73	p=0.030
Fetal complications			N=109	
No	14(39%)	38 (52%)	52	n = 0.129
Yes	22 (61%)	35 (48%)	57	p=0.138
Opportunistic infections			N=120	
Yes	8 (20%)	5 (6%)	13	n = 0.022
No	32 (80%)	75 (94%)	107	p=0.022

Table 4: Complications in women who crossed 28 weeks of pregnancy.

Complications	CD4 count ≤349 (n=35)	CD4 count ≥350 (n= 74)	Total N=109	
Term of delivery				
Preterm	15 (43%)	22 (30%)	37	- 0.292
Term	19 (54%)	45 (60%)	64	p=0.383
Post term	1 (3%)	7 (9%)	8	
Membrane status				
Intact	25 (72%)	55 (74.3%)	80	- 0.626
PROM	10 (28%)	19 (25.7%)	29	p=0.626
Birth weight				
Low (<2.5 kg)	23 (65.7%)	35 (47.3)	58	
Normal (>2.5 kg)	12 (34.3)	39 (52.7)	51	p=0.072

Of the 109 women who crossed 28 weeks of gestation, 35 pregnant women had low CD4 count and 74 women had high CD 4 count (Table 4). Even though the incidences of preterm labour and PROM were higher than in low CD4 count group than high CD4 group, it did not reach statistical significance.

# DISCUSSION

In the present study, majority of women were in the age group of 20 to 25 years, illiterates and had a married life of less than 4years. The study population had 81% Hindus, 18% Muslims and less than 1% Christians. These parameters fairly represent the demographic parameters of Indian city, like Bangalore. Only seven per cent women had history of blood transfusion and none had history of IV drug abuse. Hence sexual route is the most common route of HIV transmission in India. Pattern of seropositivity in husbands confirms the heterosexual mode of transmission as the most common mode. High risk behaviour (polygamous relationship) was observed in 4% of the patients in contrast to 34% in husbands and in 4% in both the partners. In recent years married monogamous women have been identified as a population at increasing risk for HIV in India. In India, drivers and commercial sex workers are already recognised as high risk for HIV infection. Authors noted high incidence in coolies working on daily wages, most of whom are immigrants to city from other villages as other high-risk group. Seventeen per cent of the women in this study group had either previous history or an ongoing presence of STDs. In another study conducted in India, 20.7% HIV positive women had additional STDs compared to only 4.6% HIV negative women.<sup>12</sup>

Of the 120 HIV positive women in the study, 6 patients had spontaneous abortion and 1 patient had an ectopic pregnancy. HIV infection in Africa has been linked to a higher rate of spontaneous abortion.<sup>13</sup> In a prospective follow-up American study showed a three-fold increase in early spontaneous abortion.<sup>14,15</sup> Authors noted incidence of preterm delivery around 34.5% in the present study, which is slightly higher than reported in other studies in HIV positive women. Temmerman et al reported that prematurity was observed in 21.1% of neonates born to HIV positive women.<sup>6</sup> In other studies in HIV infected women the incidence of premature delivery varied from 14-23%.<sup>16,17</sup>

The incidence of PROM in the present study was 26.6%. Other studies have observed incidence of PROM varying from 15-50%.<sup>7,12</sup> Normal vaginal delivery was the most common route of delivery in the present study and LSCS was carried out only in 18.35%.

Authors noted very high incidence of low birth weight in present patients at 52%. Most of the western studies have reported incidence like 18.9%, 19.6%, where incidence of low birth weight in general population is very low.<sup>6,16</sup> Even other studies from India and Africa have given incidence from 9%- 22%.<sup>12,18</sup> This is still far from 52% noted in present study. Incidence of low birth weight is known to vary with environmental and many other parameters.

There were 5 still births and 5 IUD out of 109 births. This is higher than incidence seen in normal population. Results are consistent with another large study in Nairobi where it was found that there was increased association between HIV and IUD and still birth rate, after controlling for the presence of other STDs.<sup>6</sup>

Authors noted 17% incidence of puerperal complications, which was even higher in women who underwent LSCS. These results are consistent with other studies, who have noted higher infectious complications during the postpartum period in HIV-positive women.<sup>4</sup>

Authors noted lower CD4 count is associated with higher incidence of obstetric complications and opportunistic infections. This was statistically as well as clinically significant. The inverse relationship between CD4 count and incidence of opportunistic infections was noted, which is well established fact. Bacterial pneumonia, urinary tract infections and other infections are more common during pregnancy in HIV seropositive women.<sup>20</sup> The study also showed that lower CD4 count was associated with increased incidence of PROM, Low birth weight, preterm delivery. Though not statistically significant (due to small sample size), it showed that there is trend towards more adverse outcomes with lower CD4 count.

## CONCLUSION

To conclude, maternal HIV infection adversely affects the pregnancy outcome and is significantly associated with higher incidences of obstetric complications and opportunistic infections, more so when the CD4 count is less. HIV infection is also associated with increased incidence of Preterm labour, PROM, low birth weight and postpartum complications and show the increasing trend as the CD4 count decreases.

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