

Original Research Article

Low prevalence and risk factors related to 6 weeks HIV-1 Mother to Child Transmission under Option B+ at military hospitals of Douala and Yaounde, and at the Bertoua regional hospital

UNDER PEER REVIEW

ABSTRACT

Aims: ~~To~~ evaluate the impact of Option B + program through the determination of Human Immunodeficiency Virus type 1 (HIV-1) Mother-To-Child Transmission (MTCT) prevalence and its predictors. ~~Because of~~ following the limited success achieved with the previous prevention ~~of~~ MTCT programs, the Cameroon's public health ministry adopted in 2014 the *Option B+* program that recommends a systematic lifelong treatment to ~~all any~~ HIV positive pregnant woman.

Study design: ~~A~~ case-control study was conducted within two groups: a reference group constituted of exposed infants from HIV positive mothers undergoing Option B+ program, and a control group ~~constituted~~ of infants from ~~no~~ Anti-Retroviral Treatment (ART) ~~naive~~ HIV positive mothers during pregnancy.

Place and Duration of Study: Douala and Yaounde ~~Military Hh~~ hospitals (*HMR2* and *HMR1* respectively) as well as the Bertoua Regional Hospital (HRB), From October 2017 to March 2018.

Methodology: ~~This this~~ research included infected mother - exposed child pairs ~~m~~. Infected mothers' sociodemographic and clinical characteristics were reported. Infants sampled ~~at for~~ six weeks ~~testing at~~ the HIV MTCT prevention units were tested for HIV-1 RNA early detection through rtPCR with Abbott m2000sp automated system at the Military Health Research Center. Multivariate logistic regression model was built to assess the predictors of MTCT and to compare groups.

Results: ~~Within the study period in 6 months~~, the overall HIV-1 prevalence in ~~the~~ 107 six weeks old reference group infants and 23 control group infants was nil and 4.35% (1/23) respectively. Logistic regression showed that predictors of HIV-1 MTCT were : home delivery $p=0.03$ and absence of ART during pregnancy $p = 0.04$.

Conclusion: ~~While u~~ untreated infected women during pregnancy were more likely to transmit the virus to their exposed ~~babies as compared to child than~~ their treated counterparts, implementation of the option B + seems a hope to eliminate HIV-1 MTCT in Cameroon. However, systematic enrollment of HIV pregnant women should be strengthened.

Keywords: Cameroon, case-control study, HIV-1, Mother-to-child Transmission, option B +, prevalence, risk factors

1. INTRODUCTION

Human Immunodeficiency Virus (HIV) is a positive sense single stranded Ribonucleic Acid (RNA) virus of the Lentiviruses family responsible for Acquired Immunodeficiency Syndrome (AIDS). HIV infection remains a serious public health problem worldwide, affecting adults, teenagers and children of both sexes [1]. In 2016, 2.75% (2.1 million) of the global population of individuals living with HIV (estimated at 76.1 million) were children and adolescents under 15 years of age. About 4000 of this group were infected through Mother To Child Transmission (MTCT) [2]. MTCT of HIV1 may occur during pregnancy, labor, delivery, or breastfeeding. In the absence of prophylactic therapy, the risk of HIV-1 MTCT, both in utero and during childbirth is estimated at 15-30%; this risk increases to 20-45% in breastfed children. This vertical transmission is ~~the~~ highest in resource-constrained countries [3,4].

In Africa, pediatric HIV-1 infection is a common cause of morbidity and mortality, and MTCT is the main route of transmission. Prevention of mother-to-child transmission (PMTCT) of HIV is therefore a priority intervention of HIV/AIDS programs, yet a serious challenge particularly in Sub-Saharan Africa where 90% of the world pediatric HIV-1 infection is recorded [3,5].

To address this public health concern in Cameroon, Option B was implemented in 2009 following Option A whose implementation started in 2004. Under Option A HIV-1 PMTCT program, when infected mothers' cluster of differentiation (CD4) ~~cell count number~~ was ~~comprised~~ between 350 and 500

Comment [A1]: Please check the statistics again for accuracy. Out of 2.1 million children and adolescents living with HIV, only 4000 contracted it through MTCT? That is only about 0.002%. So how did the rest contract the virus?

cells/mm³, they were ~~initiated on~~ ~~subjected to~~ antiretroviral (ART) monotherapy from the 14th week of pregnancy till one week after the delivery while newborns ~~receive~~ nevirapine (NVP) prophylaxis ~~and the duration of the medication~~ depended on the feeding mode. The mothers' monotherapy gave way to ~~triple therapy~~ ~~tritherapy~~ in Option B while systematic 4-6 weeks NVP prophylaxis of newborns was applied [6]. The achieved goals for both programs was very limited; 12.1% in 2011 for Option A [7] and 7.1% in 2014 for Option B [8], indicating a still very high transmission rate. With the support of USA President's Emergency Plan for AIDS Relief (PEPFAR) in Cameroon, the Ministry of Public Health adopted PMTCT program - Option B+ aiming to achieve the ambitious global targets of 95-95-95 set by the Joint United Nations Programme on HIV/AIDS[9]. The 95-95-95 targets is defined as 95% of all people living with HIV to know their HIV status, 95% of all people living with HIV to have access to antiretroviral therapy, and 95% of people on antiretroviral therapy to have an undetectable viral load [6]. These global targets for the infants aimed at eliminating the MTCT of HIV [2]. Option B + recommends a systematic and lifelong treatment of any HIV- positive pregnant or breastfeeding woman regardless of her viral load or CD4 count [10].

Option B+ ~~has been~~ ~~is~~ under implementation in Cameroon since 2014 and it is ~~therefore appropriate high time~~ to evaluate its added value in the PMTCT of HIV-1 in Cameroon with respect to the global targets for infants. Worldwide studies ~~have~~ identified potential risk factors for vertical HIV transmission [7]. These include: maternal factors (maternal immune status, ART regimen), obstetrical factors (traumatic birth, premature rupture of membranes, chorioamnionitis), fetal and infant factors (immune status, prophylaxis, feeding mode) [2,6,10,11]. As recommended by the World Health Organization (WHO), assessing the ~~implementation of~~ PMTCT of HIV-1 programs provides optimal programmatic choices for PMTCT [6]. Thus, this study aimed to determine the prevalence and investigate risk factors related to mother-to-child transmission of HIV-1 in infants at 6 weeks ~~of~~ age in hospitals where ~~the~~ option B+ prevention program is implemented, namely the Military Hospitals of Douala (HMR2) ~~and~~ the Military Hospital of Yaounde (HMR1) in comparison to ~~exposed neonates at the Bertoua Regional Hospital (HRB) where the option B+ strategy has not been adopted, newborns out of the program at the Bertoua Regional Hospital (HRB).~~

2. MATERIAL AND METHODS

2.1 Study design, period and population

~~A case control study with two groups: the reference group and the control group was conducted between October 2017 and March 2018. From October 2017 to March 2018 was conducted a case-control study with two groups: the reference group and the control group.~~ The reference group ~~was~~ constituted of six weeks ~~old~~ infants born ~~to~~ HIV-1 positive mothers undergoing ~~the~~ Option B+ program, and the control group (no ART group) ~~consisted was constituted~~ of six weeks ~~old~~ infants from HIV-1 positive mothers who discovered their HIV status during delivery. Participants of the reference group were sampled at the Douala and Yaounde military hospitals (*HMR2* and *HMR1* respectively), while participants

of the control group were sampled at the Bertoua Regional Hospital (BRH). BRH was chosen as study site for the control group because the Eastern region ~~is features~~ among the highly HIV affected regions in Cameroon and the ~~eastern populations~~-cultural practice ~~in that area~~ remains refractory to antenatal care (ANC) with only 49% of pregnant women going for ANC visit[12]. Recruitment was consecutive and non-probabilistic. References and controls were not matched because of the scarcity of ~~no-ART~~ ~~naiveniawe~~ HIV positive pregnant women.

2.2 Data collection and sample testing

Socio demographic and clinical information of consenting ~~retroviral positive~~ mothers including ~~mode of way of~~ delivery, ART regimen ~~and HIV-1 status~~ as well as ~~neonates newborns'~~ oral NVP prophylaxis regimen and feeding mode were collected using a questionnaire. Dried blood spots (DBS) samples on Whatmann 903 paper were collected following recommended sampling standards from all six weeks ~~old~~ participating infants [13]. Samples were then tested for early infants' diagnosis of HIV-1 RNA by real time reverse transcriptase polymerase chain reaction (RT-PCR) using Abbott m2000 Real Time HIV-1 Qualitative test as described ~~elsewhere~~, at the Military Health Research Center (CRESAR), Yaounde-Cameroon[14].

2.3 Quality assurance

Samples were tested according to the ISO 15189:2012 standard of medical analysis laboratories [13]. External quality control consisted of samples with known HIV-1 results from an accredited laboratory. The internal processing control (one negative and two positive manufactured controls with high and low levels) consisted of HIV nonspecific primers designed to evaluate the amplification. A test was validated under 3 conditions: internal control should have been amplified in all the samples, negative and positive controls should have been negative and positive respectively.

2.4 Administrative and ethical considerations

Prior to conducting this study, authorizations were obtained from the Cameroon's Ministry of Defense (No 006027/LE/MINDEF/01) and from the East Regional Delegation for Public health (No 896/L/MINSANTE/SG/DRSPE/BCASS/BFP). Consent for the HIV-exposed infants enrolled in the study was given by the mother or caregiver orally or by signing the consent form. Samples were labelled and treated following the principles of Helsinki declaration.

2.5 Data analysis

Statistical analysis was done using the Statistical Package for the Social Sciences (SPSS), software (IBM, Version 25.0. Armonk, NY: IBM corp). Descriptive statistics were done using frequency distribution. Multivariate logistic regression model for multimodal variables and binary logistic regression model for bimodal variables were built to assess the predictors of MTCT and to compare groups. Variables were considered statistically significant for a p-values ~~s~~ less than 0.05.

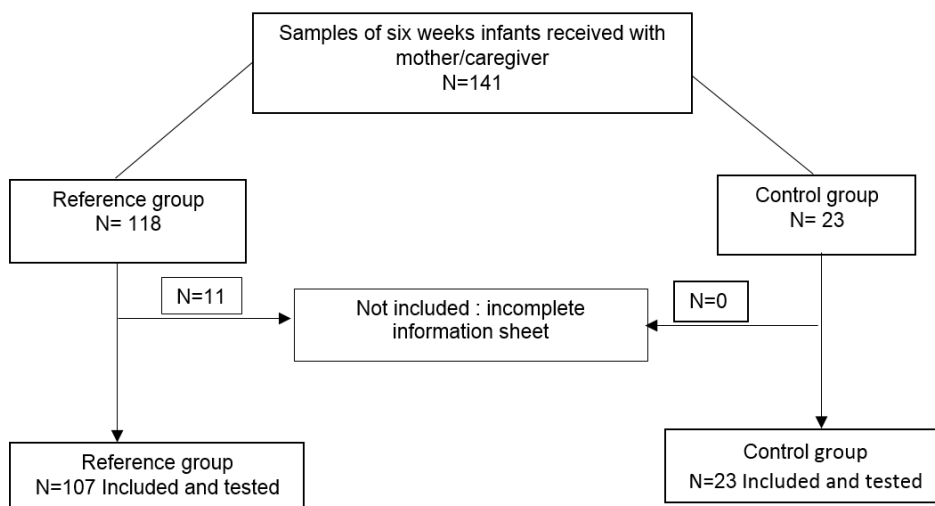
3. RESULTS AND DISCUSSION

3.1 Results

3.1.1 Sociodemographic characteristics of reference and control groups in mothers

During the six months, 141 samples of six weeks old infants seen at the PMTCT received to the MTCT Prevention units with their mothers/caregivers were collected and distributed as follows: 118 in the reference group and 23 in the control group. Out of these them, 11 were not included because of lack of suitable information about the infected mother from the caregiver in the reference group (Fig 1).

Comment [kcs2]: The top box in the diagram would read better if changed toSamples of six weeks old infants..



N : number

Figure 1: Flow diagram showing the study population

Mothers were aged 17 to 42 years with an average of 29 ± 5 years. The most represented age range was 25-34 years old with 70 (83.3) and 14 (16.7) in reference and control groups respectively. According to the results (see of Table 1), the later age was a risk for vertical transmission of the virus ($p=0.02$). As the samples were collected in the Center and Littoral regions for the reference group, and in the East region for the control group. Most infected mothers of the reference group resided in Douala 49 (98%) in the Littoral region, and 89,5% of the control group were from the East region. The lowest educational level (primary school) was the most observed in women of the control group (table 1).

Formatted: Font color: Red

Formatted: Font color: Red

Formatted: Font color: Red

Table 1. Sociodemographic characteristics of reference and control groups in mothers

Characteristics	Reference group mothers (N=107), n (%)	Control group mothers (N=23), n (%)	p-value
Mean±sd, years	29.60± 5.0	30.87± 6.1	
Age range (years)			
15 – 24	17 (85.0)	3 (15.0)	.06
25 - 34	70 (83.3)	14 (16.7)	
35 – 44	17 (73.9)	6 (26.1)	.02
East Region Origin	2(10.5)	17(89.5)	.9
Douala Residence	49 (98.0)	1 (2.0)	1.0
Primary school education level	6 (31.6)	13 (68.4)	.01
Married status	10 (47.6)	11 (52.4)	.8

Key: N: total number; n: number, % : percentage

3.1.2 Gender, age, place of birth and mode of delivery in infants

The National algorithm for the follow-up of HIV infected mothers and their children recommends that the first early infant diagnosis should be done [early after birth](#), about six weeks after the delivery. Accordingly, out of the 130 infants sampled, 69 (76.7%) and 21 (23.3%) were aged less than six weeks in the reference and control group respectively. [The](#) frequency of 84.6% of caesarian was observed in the reference group, while the control group registered 100% of infants born at home with a high risk of HIV MTCT transmission (p=0.03) ([See Table 2](#)).

Table 2. Demographic characteristics of reference and control groups in infants

Characteristics	Reference infants (N=107) n (%)	Control infants (N=23) n (%)	p-value
Age range (weeks)(reference=1-10)			
1 - 10	69 (76.7)	21 (23.3)	
11 - 20	7 (87.5)	1 (12.5)	.03
21 - 30	14 (93.3)	1 (6.7)	.02
31 - 40	13 (100.0)	0 (0.0)	1
41 - 50	4 (100.0)	0 (0.0)	1
Home birth place	0 (0.0)	2 (100.0)	.03
Caesarean birth	11(84.6)	2(15.4)	.07
Male	44 (75.9)	14 (24.1)	.005

Key: N: total number; n: number, % : percentage

3.1.3 HIV-1 Mother-to-child transmission prevalence

An Unundetected HIV-1 RNA was considered negative. And-anywhile detected HIV RNA was considered as a positive qualitative HIV-1 rtPCR test. Analysis of exposed newborns and infants taken from DBS gave a MTCT prevalence of 4.35% (1/23) in control group (Fig.2a) and nil prevalence (0/110) in the reference group under option B+ (Fig.2b)

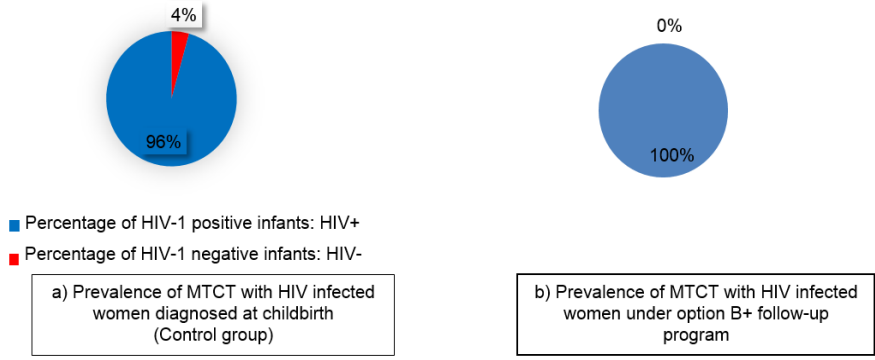


Figure 2: HIV-1 Mother-To-Child Transmission (MTCT) prevalence among infants of control and reference groups

3.1.3 Risk factors related to vertical transmission of HIV-1 infection

The table 3 summarizes the distribution of risk factors of MTCT related respectively to mother and infants according to their infectious status and socio-demographical characteristics, as well as the statistical analysis of maternal and infant's parameters with HIV MTCT (Tab. 3). During the study period, 1 each, representing 50% of infants were symptomatic at the sampling moment in each group. *As age range of infection known, 11-15 years, traditional medicine taken during pregnancy, smoking during pregnancy, no regular physical activities concern, there is no correlation with these factors of exposure and the vertical transmission of the virus.*

Formatted: Font: Italic

On At the contrary, the place of delivery was significantly related to MTCT, with home birth place being a major risk factor for vertical HIV transmission ($p = 0.03$), as well as the absence of mother's ART medication before or at birth ($p = 0.04$).

Comment [kcs3]: For the risk factors to be clear to readers, author may need to explain the parameters in this paragraph properly. The sentence structure may also need revision.

The two known delivery ways were represented, namely vaginal delivery the low normal way and caesarean section; the caesarean section was applied only in 11/107 in reference and 2/23 in control group (Tab. 2). However, vaginal delivery the delivery way was not associated related to MTCT in this study ($p > 0.05$), neither the a absence of prophylaxis to the newborn at delivery and the late infants' diagnosis were also not linked to MTCT ($p = 0.9$) (Tab. 3).

Formatted: Font: Italic

Formatted: Font: Italic

Table 3. Risk factors related to MTCT of HIV-1 in the study population under option B+ and [those who were not](#)

Exposure	Reference group	Control group	p-value
Symptomatic infant	1 (50.0)	1 (50.0)	.7
Artificial feeding	11 (78.6)	3 (21.4)	.0
Absence of prophylaxis to the newborn	0(0.0)	2 (100.0)	.9
Home birth place	0 (0.0)	2 (100.0)	.03
Late infant diagnosis	0 (0.0)	1 (100.0)	.9
known age range of infection, 11-15 years	3 (75.0)	1 (25.0)	.9
ART of women before or at birth (reference= HAART (TDF+3TC+EFV))			
AZT+3TC+EFV	10 (100.0)	0 (0.0)	1
AZT+3TC+NVP	4 (100.0)	0 (0.0)	1
HAART (TDF+3TC+EFV)	71 (82.6)	15 (17.4)	
TDF+3TC+NVP	5 (50.0)	5 (50.0)	1
Absence	5 (83.3)	1 (16.7)	.04
Traditional medicine taken during pregnancy	0 (0.0)	3 (100.0)	1
Smoking during pregnancy	0 (0.0)	1 (100.0)	1

Comment [A4]: Absence of what? ART at birth?

Alcohol drinking during pregnancy	8 (40.0)	12 (60.0)	.9
No regular physical activities	44 (71.0)	18 (29.0)	.9

ART: Antiretroviral Treatment; AZT: Azithromycin; 3TC: Lamivudine; EFV: Efavirenz; Nevirapine; TDF: Tenofovir

3.2 Discussion

3.2.1 Socio-demographic and clinical data

Mother-to-child transmission causes about 90% of pediatric HIV-1 infections worldwide. Data on MTCT from various countries and regions are as diverse as the social contexts are not the same. MTCT of HIV may be enhanced or better controlled depending on of each local social context, including cultural practices and levels of education, income, access to healthcare, implementation of preventive measure, the prevalence of infection among women in reproductive age in general and in pregnant women in particular [15].

In Cameroon, available data on HIV-1 MTCT are mainly those on the implementation of options A and B in public hospitals and civilian research centers [5]. However, no published data on Option B+ and MTCT of HIV1 are available from for military hospitals. The present work is a complement of what has been done so far. Option B + holds two advantages; it reduces infant mortality and morbidity and it increases the mother's life expectancy up to 3.75 years [16]. As one of the WHO world health organization recommendations was to determine the best program adapted in to any context for the elimination of the vertical transmission of HIV, it would be important to address the question of associated risk factors, and to identify gaps in the follow-up of HIV-positive pregnant women in Cameroon.

In this study, one infant out of 130 was infected via MTCT, a baby boy. Although this number is not significant enough to infer a gender influence in MTCT of HIV-1, reports from India, (14.2% boys vs. 8.2% girls) and Kenya (23% boys vs. 11% of girls) indicated that baby boys were more vulnerable to MTCT-HIV than baby girls [11,17].

Concerning the study site, it is noted that the present research was carried out for reference group in two military hospitals, HMR1 and HMR2 respectively in Yaounde and Douala. In Cameroon, the general public are not quite is not well aware that non-military patients are welcome in these hospitals. This is an important break in communication information break that has perhaps prevented the entire population of the towns of Douala and Yaounde from partaking in the to profit from the efficient implementation of Option B+ in these hospitals. This may also and as well this may justify the low sample size within the time period of this study[18].

The maximum frequency of caesarian sections (84.6%) was observed in the reference group. This is in accordance with a study where it was has been established that delivery through caesarean

Comment [kcs5]: Please indicate where and when study was done in the sentence to make it more comprehensive.

section dropped the risk of HIV transmission ~~by about 50%~~ ~~as compared to with~~ other types of delivery. That percentage increases if the seropositive female ~~adheres to her ARVs. follows antiretroviral therapy correctly.~~ The combination antiretroviral therapy plus caesarean section before or shortly after membrane ruptures drops the transmission to 87% [11,17]

One major socio-demographic parameter in this study was ~~the~~ home delivery. In fact, ~~two out of twenty three 2/23~~ mothers from the control group delivered at home. Home delivery was found to influence HIV-1 MTCT and represents an ~~adverse impairing~~ factor of PMTCT ($p = 0.03$). The non-application of preventive treatment during pregnancy and delivery, and the absence of newborn prophylaxis are ~~also~~ important risk factors accounting for more than 60% ~~towards contribution of~~ MTCT of HIV1 [2,10]. A study conducted in Zambia showed that it was possible to offer home-based HIV testing and NVP to traditional birth attendants for women who do not have easy access to health facilities in rural areas [19]. Cameroon through the ministry of public health should envisage extension of Option B+ in remote areas where access to healthcare facilities is not easy. This will lower significantly the number of undeclared cases, and the deficiency of follow-up. In France, Frange and Blanche founded that the rare residual cases of MTCT of HIV-1 were not related to the failure of the antiretroviral therapy, but rather associated with deficiencies in ~~factors like:~~ the follow-up of pregnant women, unrecognized screening tests, specialized medical follow-up initiated late or stopped early ~~and,~~ occurrence of an undiagnosed primary infection during pregnancy or breastfeeding ~~among others~~[20].

3.2.2 Impairing Prevalence of HIV-1 MTCT

HIV-1 MTCT prevalence of 0% was found in ~~the~~ reference group. This prevalence seems encouraging with regard to the UNAIDS global targets ~~of 95-95-95~~. However, further studies extended to the 10 regions of Cameroon would ~~be have been~~ necessary to confirm the closeness to the above ~~UNAIDS USAIDS~~ global target by 2025. This prevalence is close to that of developing and industrialized countries such as Cuba and France where the rate of HIV1 MTCT is lower than 2% [20,21], and is indicative of an improvement if compared to the 0.9% global prevalence of HIV1-MTCT [9]. Between 2008 and 2014 in Cameroon under the application of option B, the prevalence of MTCT was 7.1% in the rural part of Bamenda subdivision, in the North-West region, 8,7% (197/2254) in 58 health facilities in three regions of varying levels [8]. This observed significant decline in mother-to-child transmission of HIV has been achieved thanks to ~~the~~ consistent increase of awareness and a larger coverage ~~of~~ PMTCT measures [2,10]. Other contributing factors could be the availability of resources with the support of

PEPFAR towards the promotion of PMTCT programs implementation in military hospitals [22]. As the matter of fact, the present result in the reference group is far ~~lower than -from t-~~ that of Njom-Nlend (where the rate of transmission was at 4.20% at 6 weeks), but similar ~~to in-~~ the control group [23].

3.2.3 Risk factors related to HIV-1 MTCT

The observed significant risk factors associated with the prevalence of HIV MTCT included the lack of protection of the fetus by the mother through ~~not taking~~ the highly active antiretroviral treatment (HAART) during pregnancy ($p = 0.04$). Indeed, ~~vertical materno-fetal~~ transmission of HIV-1 occurs mainly from the end of the second trimester of pregnancy and ~~is~~ accounts for 15 to 25% of MTCT. It occurs during the passage of the virus from the mother to the fetus via the placenta, amniotic fluid or micro-transfusions [6,24]. The risk of infection in utero varies with maternal viral load, infections of the placenta by other pathogens, and the presence of sexually transmitted diseases. Nevertheless, it can be reduced by early maternal HAART ~~medication~~ continued throughout the pregnancy. The practice helps via two mechanisms: the reduction of maternal viral load in blood and genital secretions, and the pre-exposure prophylaxis through ART that cross the placenta and produce adequate systemic levels in infants especially during the passage into genital canal [1]. In the present study, 71/107 participants in the reference group and 15/23 participants in the control group had early HAART. Thus, (as also noted by Ngwej *et al*) the absence of maternal ART during pregnancy had a 10-fold increased risk $P < 0.0001$ [25].

Absence of Nevirapine (NVP) administration to the newborn was not statistically related to vertical transmission ($p=0.9$). Nevirapine is a non-nucleotidic inhibitor of HIV-1 reverse transcriptase. In case of exposure, it blocks viral replication by binding to the catalytic site of the reverse transcriptase, thus inhibits RNA-dependent DNA polymerase activities. In the absence of NVP prophylaxis, the process of viral replication remain active, so viral colonization of the newborn remains possible [1,4]. A different result was obtained in Burkina Faso by Ouedraogo *et al* (2015) where is was found that the risk of vertical HIV transmission was increased with the absence of antiretroviral administered to the newborn immediately after birth [26].

Comment [kcs6]: For clarity please state the early HAART the reference group and control group received because the subsequent sentence about the absence of maternal ARVs during pregnancy makes it a bit confusing... You stated in the design that the control group was not on ART.(See study design section 2.1 under materials and methods) and so how come 15/23 had early HAART?

In the control group, the prevalence of MTCT-HIV was 4.35%. Since the global distribution of the MTCT indicates 15-25% of transmission in utero in the absence of ART, the babies from these women would have been more infected than that because of ignorance and lack of awareness [2,6,10]. In Lubumbashi Democratic Republic of Congo in a similar situation, 25% of the unaware mothers transmitted the virus to their newborns [25]. A contrary result was obtained with the study of Njom-Nlend where infection discovery during the last pregnancy was not a risk factor to the vertical transmission of HIV [23].

3.2.4 Limitations and strength

This study ~~had has~~ some limitations. Reference and control group mothers were not paired because of the difficulty ~~in enrolling to meet~~ urban mothers who do not visit antenatal care units during pregnancy (control group). Also, the sample size was not consistent within the collection period because of the exclusion of some samples for which information sheets did not contain reliable data relevant to this research. This could be the reason why there is ~~a~~ probable ~~omission missing~~ of positive cases. Nevertheless, one of the strengths of this research is that it adds value to previous studies in terms of comparison between the two groups of infected mothers: ~~those~~ on ART and ~~those~~ not on ART during pregnancy, although the study population was not divided into subgroups.

4. CONCLUSION

Transmission of HIV-1 from mother to child was non-existent under option B+ in the military hospitals of Douala and Yaounde: 0% (1/107), while the rate of transmission was 4.35% in women who discovered their seropositivity at or after birth in the BRH (1/23) within the study period. The risk factors associated with MTCT were home delivery (p =0.03) ~~and~~ absence of highly active antiretroviral therapy (HAART) during pregnancy (p = 0.04). This prevalence is encouraging with respect of the ambitious 95-95-95 targets of UNAIDS, which aims to eliminate the MTCT. However significant awareness to raise adherence and compliance to prevention programs are needed to ensure that all pregnant HIV-positive women are enrolled in a management program so that the risk factors associated with the infection of the newborn are ~~brought~~ under control, including the Highly active antiretroviral treatment of HIV positive pregnant women, the respect of the ANC schedule and the delivery ~~at to~~ an appropriate health facility. We recommend a nationwide implementation of Option B+ to considerably reduce MTCT of HIV1 in Cameroon.

Comment [kcs7]: ...if the transmission rate is 0% then shouldn't it be (0/107)

CONSENT AND ETHICAL APPROVAL

The present study received Ethical approval N° 006027/LE/MINDEF/01 from the Cameroon's Institutional Research Board of the Ministry of Defense as well as the ethical authorization N° 896/L/MINSANTE/SG/DRSPE/BCASS/BFP from the Bertoua Regional Hospital prior to carrying out the research. Prior to the sample collection, mothers or caregivers were given an information's notice, afterwards consents for the HIV-exposed infants enrolled in the study were obtained by signing the consent form. All authors hereby declare that experiments have been examined and approved by the appropriate ethics committee and have therefore been performed in accordance with the ethical standards laid down in the 1964 Declaration of Helsinki.

REFERENCES

- [1] Lusic M, Siliciano RF. Nuclear landscape of HIV-1 infection and integration. *Nat Rev Microbiol* 2017;15:69–82. <https://doi.org/10.1038/nrmicro.2016.162>.
- [2] World Health Organization. Prevention of mother-to-child transmission of HIV: review of the evidence. World Health Organization; 2001. English.
- [3] Billong S-C, Fokam J, Billong E-J, Nguéfack-Tsague G, Essi M-J, Fodjo R, et al. Epidemiological distribution of HIV infection among pregnant women in the ten regions of Cameroon and strategic implications for prevention programs. *Pan Afr Med J* 2015;20:79. English. <https://doi.org/10.11604/pamj.2015.20.79.4216>.
- [4] Tejiokem MC. Early management of HIV infection in infants in a sub-Saharan African country with medium HIV prevalence. Paris Sud-Paris XI University, 2012. French.
- [5] Minhealth. National Health Development Plan PNDS 2016-2020 2016. French. https://www.minsante.cm/site/sites/default/files/PNDS_FRANCAIS-min.pdf (accessed June 29, 2023).
- [6] World Health Organization. Use of antiretroviral drugs to treat pregnant women and prevent HIV infection in infants. *WwwWhoInt* 2012. English. https://apps.who.int/iris/bitstream/handle/10665/70891/WHO_HIV_2012.6_eng.pdf?sequence=1 (accessed July 4, 2023).
- [7] Nguéfack HLN, Gwet H, Desmonde S, Oukem-Boyer OOM, Nkenfou C, Tejiokem M, et al. Estimating mother-to-child HIV transmission rates in Cameroon in 2011: a computer

- simulation approach. *BMC Infect Dis* 2016;16:11. <https://doi.org/10.1186/s12879-016-1336-2>.
- [8] Fondoh VN, Mom NA. Mother-to-child transmission of HIV and its predictors among HIV-exposed infants at Bamenda Regional Hospital, Cameroon. *Afr J Lab Med* 2017;6:589. <https://doi.org/10.4102/ajlm.v6i1.589>.
- [9] UNAIDS. UNAIDS announces 2 million more people living with HIV on treatment in 2015, for a total of 17 million in 2023. English. https://www.unaids.org/en/resources/presscentre/pressreleaseandstatementarchive/2016/may/20160531_Global-AIDS-Update-2016 (accessed June 29, 2023).
- [10] World Health Organization. Programmatic update: use of antiretroviral drugs for treating pregnant women and preventing HIV infection in infants: executive summary. World Health Organization; 2012.
- [11] The International Perinatal HIV Group. The Mode of Delivery and the Risk of Vertical Transmission of Human Immunodeficiency Virus Type 1 — A Meta-Analysis of 15 Prospective Cohort Studies. *N Engl J Med* 1999;340:977–87. <https://doi.org/10.1056/NEJM199904013401301>.
- [12] Anoubissi J de D, Gabriel EL, Kengne Nde C, Fokam J, Tseuko DG, Messeh A, et al. Factors associated with risk of HIV-infection among pregnant women in Cameroon: Evidence from the 2016 national sentinel surveillance survey of HIV and syphilis. *PloS One* 2019;14:e0208963. <https://doi.org/10.1371/journal.pone.0208963>.
- [13] Veenhof H, Koster RA, Brinkman R, Senturk E, Bakker SJL, Berger SP, et al. Performance of a web-based application measuring spot quality in dried blood spot sampling. *Clin Chem Lab Med* 2019;57:1846–53. <https://doi.org/10.1515/cclm-2019-0437>.
- [14] Pyne MT, Konnick EQ, Phansalkar A, Hillyard DR. Evaluation of the Abbott Investigational Use Only RealTime HIV-1 Assay and Comparison to the Roche Amplicor HIV-1 Monitor Test, Version 1.5. *J Mol Diagn JMD* 2009;11:347–54. <https://doi.org/10.2353/jmoldx.2009.080166>.
- [15] Ministry of Health, WHO, African Health Observatory. Analytical health profile 2016 Cameroon 2016. French. https://www.afro.who.int/sites/default/files/2017-07/Profil%20sanitaire_Cameroun_2016_%20FRENCH.pdf (accessed June 29, 2023).
- [16] Hassan AS, Sakwa EM, Nabwera HM, Taegtmeier MM, Kimutai RM, Sanders EJ, et al. Dynamics and constraints of early infant diagnosis of HIV infection in Rural Kenya. *AIDS Behav* 2012;16:5–12. <https://doi.org/10.1007/s10461-010-9877-7>.

- [17] Cambrea SC, Pinzaru AD, Cambrea SC, Pinzaru AD. Value of Caesarian Section in HIV-Positive Women. *Caesarean Sect., IntechOpen*; 2018. <https://doi.org/10.5772/intechopen.76883>.
- [18] Nlo'o AE, Kuate LM, Yayehd K, Nkoke C, Abissegue YG, Bell NW, et al. Management of Heart Failure in the Yaounde Military Hospital - Cameroon. *Health Sci Dis* 2016;17.
- [19] Brennan AT, Thea DM, Semrau K, Goggin C, Scott N, Pilingana P, et al. In-home HIV testing and nevirapine dosing by traditional birth attendants in rural Zambia: a feasibility study. *J Midwifery Womens Health* 2014;59:198–204. <https://doi.org/10.1111/jmwh.12038>.
- [20] Frange P, Blanche S. HIV and mother-to-child transmission. *Medical Press* 2014;43:691–7. <https://doi.org/10.1016/j.lpm.2014.02.015>. English.
- [21] Lenzer J. How Cuba eliminated mother-to-child transmission of HIV and syphilis. *BMJ* 2016;352:i1619. <https://doi.org/10.1136/bmj.i1619>.
- [22] Awungafac G, Njukeng PA, Ndasi JA, Mbuagbaw LT. Prevention of mother-to-child transmission of the Human Immunodeficiency Virus: investigating the uptake and utilization of maternal and child health services in Tiko health district, Cameroon. *Pan Afr Med J* 2015;20:20. <https://doi.org/10.11604/pamj.2015.20.20.5137>.
- [23] Njom Nlend AE, Nguedou Marcelle K, Koki Ndombo P, Brunelle Sandié A. [12-months efficacy of option B+ for prevention of mother-to-child transmission of HIV in Yaoundé, Cameroon]. *Rev Epidemiol Public Health* 2019;67:163–7. <https://doi.org/10.1016/j.respe.2019.03.119>.
- [24] Sama C-B, Fetei VF, Tindong M, Tanyi JT, Bihle NM, Iii FFA. Prevalence of maternal HIV infection and knowledge on mother-to-child transmission of HIV and its prevention among antenatal care attendees in a rural area in northwest Cameroon. *PLOS ONE* 2017;12:e0172102. <https://doi.org/10.1371/journal.pone.0172102>.
- [25] Ngwej DT, Mukuku O, Mudkereza R, Karaj E, Odimba EBF, Luboya ON, et al. Study of risk factors for HIV transmission from mother to child in the "option A" strategy in Lubumbashi, Democratic Republic of Congo. *Pan Afr Med J* 2015;22. English. <https://doi.org/10.11604/pamj.2015.22.18.7480>.
- [26] Ouédraogo Yugbaré SO, Zagré N, Koueta F, Dao L, Kam L, Ouattara DY, et al. Effectiveness of the prevention of mother-to-child transmission of the Human Immunodeficiency Virus by the 2010 protocol of the World Health Organization at the Saint Camille Medical Center in Ouagadougou (Burkina Faso). *Pan Afr Med J* 2015;22:303. English. <https://doi.org/10.11604/pamj.2015.22.303.7720>.