Clinical Algorithm for Screening of HIV Among High-risk Children – Need of the Hour!

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Human immunodeficiency virus (HIV) transmission in children is most commonly due to vertical transmission [1]. Under the Prevention of parent to child transmission (PPTCT) program, all pregnant women are screened for HIV, which enables early diagnosis of HIV-exposed infants as they are at high risk for malnutrition, growth failure, developmental delay and repeated infections with common as well as uncommon organisms [1]. Without treatment, about one third of infants living with HIV will die in their first year and 50% by the second year of life [1]. The National HIV program provides access to early diagnosis for HIV testing of infants and children younger than 18 months who are HIV-exposed, and ensures that they receive the required essential package of care as part of the country's commitment on achieving 90-90-90 target by 2030, which aims at ending the acquired immunodeficiency syndrome (AIDS) epidemic [1]. In an attempt to achieve the target, screening of sick infants/children with unknown HIV exposure is important.

Perinatally-infected adolescents are more likely to suffer from chronic diseases, neurodevelopmental delay, growth and pubertal delays, unlike adolescents who acquire HIV behaviorally [1]. In 2014, it was estimated that 15% of all persons living with HIV in United States had undiagnosed HIV infection [2]. As per the US Preventive Services Task Force Recommendation (USPSTF) recommendation, persons aged 15-65 years should be screened for HIV at least once, and younger adolescents and older adults at increased risk should also be screened [3]. As per World Health Organization (WHO) 2010 guideline, it is strongly recommended to use the clinical algorithm and serologic test in the absence of virologic testing in sick infants for presumptive clinical diagnosis of HIV infection [4]. The adverse social and economic factors like poverty, broken families, parental sickness/drug abuse, and stigmatization by the society are the factors hindering access to medical care [1]. Integrated Management of Neonatal and Childhood Illness (IMNCI), which is adapted from the global

version of Integrated Management of Childhood Illness (IMCI), is a strategy to address high infant mortality and to meet sustainable developmental goals with target of reducing under five mortality to 25 per 1000 live birth [5,6]. There are mainly seven clinical features included in the IMCI/HIV algorithm for the clinical diagnosis of HIV infection in children – pneumonia, persistent diarrhea, ear discharge (acute or chronic), very low weight for age, oral thrush, parotid enlargement, and generalized persistent lymphadenopathy [7].

In an African study [8], the performance of the IMCI HIV algorithm in a cohort of 444 HIV-exposed Kenyan infants was studied. The overall sensitivity, specificity, positive predictive value (PPV) and negative predictive value (NPV) were 58%, 87%, 52% and 90%, respectively. It was noted that sensitivity was lowest at 1 month of age, when majority of HIV infections already had occurred and initiation of treatment is most important. The use of IMCI was estimated to delay diagnosis in HIV-infected infants by a median of 5.9 months. Oral thrush (67%), lymphadenopathy (55%) and pneumonia (55%) were the most commonly identified features in HIV-1 infected infants. However, IMCI still is useful in identifying older children with undiagnosed HIV-1 infection, acquiring infection through late breastmilk transmission [8]. Sensitivity and specificity estimates of HIV clinical algorithms over various studies have ranged from 9-89% and 42-99%, respectively [8].

Integrated Management of Adolescence and Adult Illness (IMAI) is a facility level health care service, which presents a syndromic case management protocol to diagnose and manage common adult illnesses [9]. The sensitivity and specificity of IMAI acute care algorithm in a HIV positive Ethiopian cohort was above 85% and above 92%, respectively [10].

In this issue of *Indian Pediatrics*, Sinha, *et al.* [11] present a cross-sectional study on the utility of Indian Council of Medical research (ICMR) modified integrated

algorithm as a screening tool in sick children for pediatric HIV case detection in health care facilities. The WHO generic IMCI-HIV screening algorithm for children up to 5 years of age, modification from Integrated Management of Adolescence and Adult Illness for children 5-14 years of age and 'other clues' for all children which includes risk factors and certain clinical conditions of WHO staging of HIV infection were used as screening tools. The HIV prevalence estimated in this study was 19.1% (5% in <5 years and 28% in 5-14 years), which is high, and is attributed to screening of sick children. The important predictors of HIV infection noted in this study were parents with HIV, unexplained fever (>1 month) and orphaned child. The strength of this study is the use of standard algorithms from IMCI HIV algorithm and IMAI which were modified as screening tools [11]. However, this screening algorithm could not be validated.

This study [11] is a multicentric study, but the population was limited to one state. Authors highlight the need for routine surveillance of HIV infection amongst children aged 5-14 years considering the high proportion of this population [11]. They concluded that one should have a high index of suspicion to consider the clinical diagnosis of HIV infection, when an infant/child with unknown HIV status but with risk factors like orphaned child, child having a single parent, child with high risk behavior, presents with symptoms and/or signs as per the clinical algorithm of WHO-IMCI. However, since this study included children predominantly above 1 year of age, the validity of algorithm in infants, especially HIV-exposed infants, needs to be evaluated.

The need of the hour is to have an estimation of HIV disease burden in symptomatic infants/children with risk factors at community level. It is important to understand that the modified integrated algorithm is a screening tool and not a diagnostic test. It is important to sensitize the healthcare workers regarding the use of this algorithm with appropriate training. The utility of this modified integrated algorithm needs to be further tested in different field studies in different states of India to confirm that these results are replicated.

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