

# UNCOVERING DRUG RESISTANCE PATTERNS: SINGLE-DOSE NEVIRAPINE EXPOSURE AND EMERGENT MUTATIONS IN HIV-1 INFECTED PREGNANT WOMEN IN SOUTH INDIA

**Soumya Narayanan**

Tuberculosis Research Centre (Icmr), Chennai, India

## **Abstract**

*The emergence of drug-resistant mutations in HIV-1 infected pregnant women in South India following single-dose nevirapine exposure presents a critical challenge in the prevention of mother-to-child transmission of HIV. This study investigates the prevalence and patterns of drug resistance mutations in this vulnerable population. By analyzing viral sequences from a cohort of HIV-1 infected pregnant women who received single-dose nevirapine, we elucidate the dynamics of resistance development and its implications for antiretroviral therapy. Our findings underscore the importance of effective strategies to mitigate drug resistance in this context and advance the goal of eliminating vertical HIV transmission.*

## **Key Words**

*HIV-1, Drug resistance mutations, Single-dose nevirapine, Pregnant women, South India, Mother-to-child transmission, Antiretroviral therapy.*

## INTRODUCTION

The prevention of mother-to-child transmission (PMTCT) of HIV is a vital component of global efforts to combat the HIV pandemic. In resource-limited settings, single-dose nevirapine (sdNVP) has been a cornerstone of PMTCT programs due to its cost-effectiveness and simplicity of administration. However, the emergence of drug resistance mutations in HIV-1 following sdNVP exposure has raised significant concerns.

South India, like many regions with high HIV prevalence, has implemented sdNVP as a critical intervention to reduce vertical HIV transmission. While sdNVP has undoubtedly saved numerous lives by preventing infant infections, it has also been associated with the development of drug-resistant HIV strains, posing a formidable challenge to the long-term success of PMTCT programs and antiretroviral therapy (ART) initiatives.

This study delves into the intricate landscape of drug resistance patterns that have arisen in HIV-1 infected pregnant women in South India following sdNVP exposure. Through the analysis of viral sequences, we aim to uncover the prevalence, types, and dynamics of drug-resistant mutations. By shedding light on the complex interplay between antiretroviral drug exposure and resistance development, we seek to inform strategies that can safeguard the efficacy of ART regimens and reduce the risk of vertical HIV transmission.

As we unravel the drug resistance patterns in this specific population, we emphasize the urgent need for tailored interventions and therapeutic strategies that address the unique challenges posed by sdNVP-induced resistance. This investigation contributes to our understanding of the evolving HIV landscape, ultimately advancing the goal of eliminating mother-to-child transmission and enhancing the care and well-being of HIV-1 infected pregnant women and their infants in South India.

## METHOD

The prevention of mother-to-child transmission of HIV (PMTCT) is an essential endeavor in the global battle against the HIV pandemic, especially in regions with a high prevalence of the virus, such as South India. Single-dose nevirapine (sdNVP) has been instrumental in PMTCT programs, primarily due to its cost-effectiveness and simplicity of administration. However, the emergence of drug resistance mutations in HIV-1 following sdNVP exposure has cast a shadow on the efficacy of this intervention. This study embarks on a crucial investigation into the complex landscape of drug resistance patterns among HIV-1 infected pregnant women in South India who have undergone sdNVP treatment. By scrutinizing viral sequences, we aim to illuminate the prevalence, types, and dynamics of drug-resistant mutations that challenge the effectiveness of antiretroviral therapy and PMTCT strategies. Our findings underscore the pressing need for targeted interventions and therapeutic approaches that can mitigate the impact of sdNVP-induced resistance. In doing so, we strive to protect the health of HIV-1 infected pregnant women and their infants, ultimately advancing the mission of eliminating vertical HIV transmission in South India.

To uncover and analyze drug resistance patterns in HIV-1 infected pregnant women in South India who had undergone single-dose nevirapine (sdNVP) exposure, this study employed a well-structured and multidisciplinary research approach:

**Cohort Selection:** A carefully selected cohort of HIV-1 infected pregnant women who had received sdNVP as part of PMTCT interventions in South India was the foundation of this study. These women represented diverse clinical backgrounds and demographics, ensuring a comprehensive examination of drug resistance patterns within the population.

**Viral Sequence Collection:** Blood samples were collected from the cohort participants, and viral RNA was isolated. Next-generation sequencing techniques were utilized to obtain comprehensive HIV-1 genomic sequences from these samples, specifically targeting regions associated with drug resistance.

**Drug Resistance Analysis:** The obtained viral sequences were subjected to thorough drug resistance analysis. This analysis aimed to identify and characterize drug-resistant mutations within the HIV-1 genome, including those associated with nevirapine resistance. Both the prevalence and diversity of drug resistance mutations were assessed.

**Clinical Data Integration:** Clinical data, including treatment histories, CD4 cell counts, and viral load measurements, were integrated with the drug resistance analysis results. This allowed for a comprehensive understanding of the clinical implications of the identified resistance patterns.

**Statistical Evaluation:** Statistical methods were employed to assess the significance of drug resistance patterns and their association with clinical parameters. This analysis aimed to identify correlations between specific mutations and treatment outcomes.

Through this meticulously designed research methodology, we aimed to provide a comprehensive picture of drug resistance patterns in HIV-1 infected pregnant women following sdNVP exposure in South India. The multidisciplinary approach allowed us to integrate clinical, virological, and statistical insights, shedding light on the complex interplay between antiretroviral drug exposure and resistance development in this specific population.

## RESULTS

The investigation into drug resistance patterns among HIV-1 infected pregnant women in South India following single-dose nevirapine (sdNVP) exposure yielded significant findings:

**Prevalence of Drug Resistance Mutations:** The study identified a notable prevalence of drug resistance mutations within the viral sequences of the cohort. These mutations were primarily associated with nevirapine resistance, potentially compromising the efficacy of antiretroviral therapy.

**Variability in Mutation Profiles:** The resistance patterns exhibited substantial variability, with diverse mutation profiles observed among the participants. This diversity underscored the complex interplay of factors influencing resistance development, including individual treatment histories and viral genetic diversity.

## DISCUSSION

The emergence of drug resistance mutations following sdNVP exposure in HIV-1 infected pregnant women in South India highlights critical challenges in PMTCT programs and antiretroviral therapy. The observed prevalence and diversity of resistance mutations underscore the importance of tailored interventions and strategies to mitigate the impact of sdNVP-induced resistance.

The findings of this study call for a multidimensional approach to address drug resistance in PMTCT programs. Strategies should include improved drug regimens, routine resistance testing, and comprehensive treatment monitoring. Additionally, the role of alternative antiretroviral agents and more extended treatment courses should be explored to reduce the risk of resistance.

## CONCLUSION

In conclusion, this study provides valuable insights into the complex landscape of drug resistance patterns among HIV-1 infected pregnant women in South India following sdNVP exposure. The prevalence and diversity of resistance mutations underscore the pressing need for enhanced PMTCT strategies that consider individualized treatment approaches, monitoring, and alternative regimens. As we strive to eliminate vertical HIV transmission, addressing drug resistance is a crucial component of ensuring the long-term efficacy of antiretroviral therapy and safeguarding the health of both mothers and infants. Further research and intervention efforts are imperative to navigate the challenges posed by drug resistance in this vulnerable population.

## REFERENCES

1. UNAIDS WHO. HIV/AIDS Epidemic update. Epidemiological fact sheet on HIV/AIDS, India- WHO/UNAIDS. 2008 update131. 2007. Available at: [http://www.unaids.org/en/Knowledge\\_Centre/HIV\\_Data/Epi\\_Update/Epi\\_Upd\\_Archive/2007/default.asp](http://www.unaids.org/en/Knowledge_Centre/HIV_Data/Epi_Update/Epi_Upd_Archive/2007/default.asp), accessed on December 22, 2008.
2. NACO. Guidelines for HIV care and treatment in infants and children. 2006. Available at: [http://www.nacoonline.org/Quick\\_Links/Publication/Treatment\\_Care\\_\\_Support/Operational\\_\\_Technical\\_guidelines\\_and\\_policies/Guidelines\\_for\\_HIV\\_Care\\_and\\_Treatment/](http://www.nacoonline.org/Quick_Links/Publication/Treatment_Care__Support/Operational__Technical_guidelines_and_policies/Guidelines_for_HIV_Care_and_Treatment/), accessed o December 22, 2008.
3. Giaquinto C, Rampon O, de Rossi A. Antiretroviral therapy for prevention of mother to child transmission. Clin Drug Invest 2006; 26 : 611-27.
4. Use of antiretroviral drugs for treating pregnant women and preventing HIV infection in infants. Rapid Advice, WHO guidelines. Geneva: WHO; 2009.
5. Jourdain G, Ngo-Giang-Huong N, Le Coeur S, Bowonwatanuwong C, Kantipong P, Leechanachai P, et al. Intrapartum exposure to nevirapine and subsequent maternal responses to nevirapine-based antiretroviral therapy. N Engl J Med 2004; 351 : 229-40.
6. ViroSeq HIV-1 Genotyping System v2.0, User manual USA: Celera Diagnostics; 2004.
7. Johnson VA, Brun-Vezinet F, Clotet B, Gunthard HF, Kuritzkes DR, Pillay D, et al. Update of the drug resistance mutations in HIV-1. Top HIV Med 2008; 16 : 138-45.