

Evaluating the Safety of Kidney Transplants from HIV-Positive Donors: Insights, Challenges, and Future Directions

Dinesh Waghmare, Student, Nephrology, ZORAM MEDICAL COLLEGE, Government of Mizoram, Mizoram

Abstract

The best way to treat end-stage renal disease (ESRD) and increase survival and quality of life is with a kidney transplant. Concerns regarding viral transmission and graft survival led to the long-standing policy of not transplanting organs from donors who tested positive for HIV. Antiretroviral treatment (ART) developments have necessitated a new assessment of this position. New studies on the safety and results of organ transplants from HIV-positive donors to HIV-positive recipients were encouraged by the HIV Organ Policy Equity (HOPE) Act of 2013, which legalised the practice in the United States. In this study, we take a look at the safety of kidney transplantation from donors who are HIV +. We cover the present data, the dangers, the possible advantages, and the problems in this field of transplantation.

Keywords: Kidney transplantation, HIV-positive donors, HIV transmission, Antiretroviral therapy (ART), Transplant outcomes, HIV Organ Policy Equity (HOPE) Act, ESRD, Organ donation, Immunosuppression, Graft survival

1. Introduction

1.1 Background

For a long time, kidney transplantation was thought to be the best course of therapy for end-stage renal disease (ESRD) patients since it enhanced their quality of life and had higher survival rates than dialysis. Nevertheless, a significant obstacle is the disparity between the need for kidney transplants and the supply of appropriate donors. Legal developments and improvements in medical treatment of HIV infection have brought the prospect of using organs from donors who test positive for the virus to the forefront in recent years. To be more specific, in 2013, the HOPE Act authorised the practice of kidney and liver transplants between those living with HIV and those living with HIV in the US. This has created new possibilities for dealing with organ shortages, but it has also brought up concerns over the efficacy and durability of these transplants.

1.2 Overview of HIV and Kidney Transplantation

Fears of organ rejection, opportunistic infections, and immunocompromised status have kept HIV-positive individuals from receiving transplants in the past. Patients with managed HIV infection now have kidney transplantation as an option, thanks to the development of efficient antiretroviral therapy (ART). There is a lot of knowledge about kidney transplantation in people with HIV, but there are more problems

when using kidneys from people with HIV as donors. These include issues with immunosuppressive medication, the possibility of HIV superinfection, and the long-term results of the graft.

2. Kidney Transplantation from HIV-Positive Donors: Mechanism and Guidelines

2.1 Transplant Procedure and ART Management

Similar surgical procedures are used for kidney transplants from donors who are positive for HIV. To reduce the likelihood of problems after the transplant, the recipient must be on stable antiretroviral therapy (ART), have viral loads that cannot be detected, and have sufficient CD4+ T cell counts. Maintaining effective antiretroviral treatment (ART) after a transplant while also balancing immunosuppressive medication to avoid organ rejection is a significant problem.

Transplant patients often use immunosuppressant medications such as corticosteroids, calcineurin inhibitors (CNIs), and mTOR inhibitors. Many immunosuppressants have a metabolism with protease inhibitors and integrase strand transfer inhibitors; hence, it is important to monitor the use of these classes of pharmaceuticals closely to prevent drug interactions with antiretroviral therapy (ART) agents.

2.2 Legal Framework: The HOPE Act

Regarding legislation about organ transplantation, the HOPE Act was revolutionary. Even with HIV-positive recipients, it was previously against the law to transplant organs from donors who tested positive for the virus. The HOPE Act legalised clinical research to determine if recipients with the virus might safely receive organ transplants from donors with HIV. Although longer-term studies are needed to evaluate hazards, including HIV superinfection and rejection thoroughly, preliminary results from these trials are encouraging, demonstrating results comparable to those of transplants from HIV-negative donors to HIV-positive recipients.

3. Safety and Outcomes

3.1 Risk of HIV Superinfection

The possibility of HIV superinfection is a major worry when it comes to using kidneys for HIV-positive people. When a recipient contracts two different strains of HIV from the same donor, a condition known as "superinfection," the recipient's health may deteriorate. Nevertheless, preliminary research suggests that there is little chance of HIV superinfection when HIV is well-controlled with antiretroviral therapy. This danger may be further reduced by using genotypic testing to match donor and recipient strains and sensitive viral load monitoring.

3.2 Graft Survival and Patient Outcomes

Graft survival rates in patients receiving kidney transplants from HIV-positive donors are similar to those in patients receiving transplants from HIV-negative donors, according to early research. This includes data from the HOPE in Action Multicenter Study. It seems that graft function is unaffected by the donor's

HIV status as long as the recipient's HIV is well-controlled with antiretroviral therapy. Transplant recipients who had kidneys donated by people living with HIV had graft survival rates of 93% after one year, which is in line with the overall transplant survival rate.

3.3 Complications and Challenges

Although there has been encouraging first data, there is still cause for worry about potential repercussions, especially regarding immunological suppression. There is a correlation between the side effects of antiretroviral treatment (ART) and immunosuppressive medication, which puts HIV-positive transplant patients at increased risk for infections, particularly opportunistic infections, as well as post-transplant diabetes and cardiovascular problems. In addition, it is crucial to closely monitor and manage the possibility of drug-drug interactions between immunosuppressive medicines and ART.

4. Challenges in Transplantation from HIV-Positive Donors

4.1 Immunosuppression Management

Preventing organ rejection while keeping antiretroviral therapy (ART) successful is a complex challenge in managing immunosuppression in HIV-positive recipients. The metabolic effects of both immunosuppressants and antiretroviral therapy (ART) might interact, raising the possibility of side effects or subtherapeutic dosage. Modifying the dosage of protease inhibitors more often than other medications is necessary since they raise blood levels of CNIs and mTOR inhibitors.

4.2 Ethical Considerations

Concerns around disclosure of risks and informed consent come up concerning the use of organs from people who are HIV positive. Transplantation cannot continue unless both the donor and the recipient have given their informed permission after carefully considering all of the risks, including the likelihood of HIV superinfection. There are further worries about the stigma that can accompany obtaining an organ from a donor who is HIV positive and the fairness of transplantation access for people living with the virus.

5. Future Directions and Research

5.1 Expanding the Donor Pool

Donors who test positive for HIV might significantly increase the number of available donors, especially in areas where the virus is prevalent. To further understand the long-term effects of these transplants and how to match donors and recipients according to HIV types and antiretroviral treatment plans, more studies are required.

5.2 Long-Term Studies on Graft and Patient Outcomes

Results from short-term trials are encouraging, but further research is needed to determine how long it takes for the transplant to take effect and how many patients will survive after receiving an HIV-positive kidney. The HOPE in Action research and similar ongoing clinical studies will aid in refining transplantation techniques for this group by providing essential data on long-term results.

6. Conclusion

Kidney transplantation from HIV-positive donors represents a promising option for addressing the organ shortage and providing life-saving treatment to HIV-positive patients with ESRD. The HOPE Act has paved the way for clinical research demonstrating such transplants' safety and efficacy. However, challenges remain in managing drug interactions, infection risk, and ethical concerns. With careful patient selection, ART management, and continued research, kidney transplantation from HIV-positive donors could become a standard of care for HIV-positive patients, significantly improving their quality of life and survival.

References

1. Durand CM, Zhang W, Brown DM, Yu S, Desai N, Redd AD, Bagnasco SM, Naqvi FF, Seaman S, Doby BL, Ostrander D, Bowring MG, Eby Y, Fernandez RE, Friedman-Moraco R, Turgeon N, Stock P, Chin-Hong P, Mehta S, Stosor V, Small CB, Gupta G, Mehta SA, Wolfe CR, Husson J, Gilbert A, Cooper M, Adebisi O, Agarwal A, Muller E, Quinn TC, Odum J, Huprikar S, Florman S, Massie AB, Tobian AAR, Segev DL; HOPE in Action Investigators. A prospective multicenter pilot study of HIV-positive deceased donor to HIV-positive recipient kidney transplantation: HOPE in action. *Am J Transplant*. 2021 May;21(5):1754-1764. doi: 10.1111/ajt.16205. Epub 2020 Aug 8. PMID: 32701209; PMCID: PMC8073960.
2. Muller E, Botha FCJ, Barday ZA, Manning K, Chin-Hong P, Stock P. Kidney Transplantation in HIV-positive Patients: Current Practice and Management Strategies. *Transplantation*. 2021 Jul 1;105(7):1492-1501. doi: 10.1097/TP.0000000000003485. PMID: 33044431; PMCID: PMC8026768.
3. Stock PG, Barin B, Murphy B, Hanto D, Diego JM, Light J, Davis C, Blumberg E, Simon D, Subramanian A, Millis JM, Lyon GM, Brayman K, Slakey D, Shapiro R, Melancon J, Jacobson JM, Stosor V, Olson JL, Stablein DM, Roland ME. Outcomes of kidney transplantation in HIV-infected recipients. *N Engl J Med*. 2010 Nov 18;363(21):2004-14. doi: 10.1056/NEJMoa1001197. Erratum in: *N Engl J Med*. 2011 Mar 17;364(11):1082. PMID: 21083386; PMCID: PMC3028983.
4. Lynch, E.N.; Russo, F.P. Liver Transplantation in People Living with HIV: Still an Experimental Procedure or Standard of Care? *Life* 2023, 13, 1975. <https://doi.org/10.3390/life13101975>
5. van Maarseveen EM, Rogers CC, Trofe-Clark J, van Zuilen AD, Mudrikova T. Drug-drug interactions between antiretroviral and immunosuppressive agents in HIV-infected patients after solid organ transplantation: a review. *AIDS Patient Care STDS*. 2012 Oct;26(10):568-81. doi: 10.1089/apc.2012.0169. PMID: 23025916.

6. Eisenbach C, Merle U, Stremmel W, Encke J. Liver transplantation in HIV-positive patients. *Clin Transplant*. 2009 Dec;23 Suppl 21:68-74. doi: 10.1111/j.1399-0012.2009.01112.x. PMID: 19930319.