



HHS Public Access

Author manuscript

J Acquir Immune Defic Syndr. Author manuscript; available in PMC 2016 January 01.

Published in final edited form as:

J Acquir Immune Defic Syndr. 2015 January 1; 68(1): e9–10. doi:10.1097/QAI.0000000000000385.

Test, Treat and Cure

Douglas F. Nixon¹, Gary L. Simon², and Rui André Saraiva Raposo¹

¹Department of Microbiology, Immunology & Tropical Medicine, The George Washington University, Washington DC, 20037

²Division of Infectious Diseases, Department of Medicine, The George Washington University, Washington DC, 20037

The HIV/AIDS research field was galvanized with the apparent functional cure of Timothy Brown, the Berlin patient [1]. The potential second case of the Mississippi baby brought additional excitement; however, the emergence of virus in this infant has given pause to hopes for additional functional cures [2]. At the recent IAS conference on HIV/AIDS, Zaunders and colleagues presented on patient C135, one of the delta nef deleted HIV virus recipients in the Sydney blood bank cohort, who is not on antiretroviral medications and in whom infectious virus cannot now be detected [3]. Whether this person is functionally cured or in remission is unknown. C135 has some genetic biomarkers that are associated with slower disease progression: HLA-B*57 positivity and heterozygosity for the CCR5Delta 32 mutation. As HIV researchers develop strategies for regimens designed towards functional cure, we propose that in HIV infected persons in whom functional cure regimens are being considered, and in whom therapeutic vaccination is a proposed intervention, immunogenomic approaches are taken into account to prioritize initial studies. For example, those with HLA or KIR alleles associated with slow progression, such as HLA-B*57 [4]; Delta 32CCR5 heterozygosity [5]; overexpression of intrinsic resistance genes [6]; and others, could be candidates prioritized for these studies. We call this identification of “remission ready” patients. Oncologists stage patients based on a number of biomarkers for entry into differential clinical practices. HIV health practitioners enrolling subjects into functional cure regimens that include therapeutic immunizations should consider taking immunogenomics into account when identifying remission ready patients.

While the ultimate goal for HIV research is a universal cure and effective vaccine, this Sydney blood bank cohort subject has shown that genetics is a powerful indicator of viral suppression and potential remission. Immunogenomic profiling can help identify other such “remission ready patients”. This strategy could be described as “test” (immunogenomic profiling), “treat” (provide anti-retroviral therapy), and “cure” (augmentative therapies designed for functional cures).

Acknowledgements

We thank the District of Columbia Developmental Center for AIDS Research (P30AI087714)

Conflicts of Interest: None to declare

References

1. Hutter G, Nowak D, Mossner M, et al. Long-term control of HIV by CCR5 Delta32/Delta32 stem-cell transplantation. *N Engl J Med*. 2009; 360:692–698. [PubMed: 19213682]
2. Ledford H. HIV rebound dashes hope of 'Mississippi baby' cure. *Nature News*. 2014
3. Zaunders J, Dyer WB, Churchill M, et al. Possible clearance of transfusion-acquired nef-deleted attenuated HIV-1 infection by a long-term non-progressor with CCR5 Delta32 heterozygous and HLA-B*57/DR13 genotype. *AIDS*. 2014 Abstract TUA0105.
4. Pereyra F, Jia X, McLaren PJ, et al. The major genetic determinants of HIV-1 control affect HLA class I peptide presentation. *Science*. 2010; 330:1551–1557. [PubMed: 21051598]
5. Huang Y, Paxton WA, Wolinsky SM, et al. The role of a mutant CCR5 allele in HIV-1 transmission and disease progression. *Nat Med*. 1996; 2:1240–1243. [PubMed: 8898752]
6. Raposo RA, Abdel-Mohsen M, Holditch SJ, et al. Increased expression of intrinsic antiviral genes in HLA-B*57-positive individuals. *J Leukoc Biol*. 2013; 94:1051–1059. [PubMed: 23929683]