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Progress Realized: Trends in HIV-1 Viral Load and CD4 Cell Count in a Tertiary-Care Center from 1999 through 2011

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Abstract

Background: HIV-1 RNA and CD4 cell counts are important parameters for HIV care. The objective of this study was to assess the overall trends in HIV-1 viral load and CD4 cell counts within our clinic.

Methods: Patients with at least one of each test performed by the Infectious Diseases Laboratory from 1999 through 2011 were included in this analysis. By adapting a novel statistical model, log10 HIV-1 RNA means were estimated by month, and log10-transformed HIV-1 RNA means were estimated by calendar year. Geometric means were calculated for CD4 cell counts by month and calendar year. Log10 HIV-1 RNA and CD4 cell count monthly means were also examined with polynomial regression.

Results: There were 1,814 individuals with approximately 25,000 paired tests over the 13-year observation period. Based on each patient’s final value of the year, the percentage of patients with viral loads below the lower limit of quantitation rose from 29% in 1999 to 72% in 2011, while the percentage with CD4 counts <200 cells/μL fell from 31% to 11%. On average annually, the mean HIV-1 RNA decreased by 86 copies/mL and the mean CD4 counts increased by 16 cells/μL. For the monthly means, the correlations (R²) from second-order polynomial regressions were 0.944 for log10 HIV-1 RNA and 0.840 for CD4 cell counts.

Conclusions: Marked improvements in HIV-1 RNA suppression and CD4 cell counts were achieved in a large inner-city population from 1999 through 2011. This success demonstrates that sustained viral control with improved immunologic status can be a realistic goal for most individuals in clinical care.

Introduction

Care for HIV-infected patients has changed dramatically over the last three decades [1] largely due to advances in antiretroviral therapies, which have allowed improvements in HIV-1 viral loads and CD4 cell counts. Because of these gains, the life expectancy of individuals diagnosed with HIV who are able to maintain fully suppressive antiretroviral regimens, now approaches those without infection [2]. However, inner-city and veteran populations with serious co-morbidities can present special challenges to achieving these gains. These comorbidities [3–6] which often include alcohol and substance abuse [7–9] and mental illness [10–12] can lead to concurrent disease manifestations and drug-drug interactions. Compared to those without infection, HIV-infected patients also have higher rates of poor treatment adherence due to lack of family/social support, adverse drug effects, complex drug regimens, psychological distress, and low patient self-efficacy [13],[14]. In addition, combination antiretroviral therapy regimens have been associated with many adverse side effects including metabolic changes and drug toxicities [15] as well as development of drug resistance [16], thus leading to virologic failure and poor clinical outcomes [17].

In order to respond to these complexities, our medical center’s Infectious Diseases Clinic has provided HIV and primary care in a comprehensive model with an on-site, multidisciplinary team of nurses, physicians, social workers, pharmacists, and medical subspecialists. In this evaluation, we reviewed the HIV-1 viral loads and CD4 cell counts from 1999 through 2011 to determine the overall trends in viral load reduction and immune reconstitution across the entire spectrum of patients receiving HIV treatment in an inner-city setting. A novel statistical model was adapted to estimate the HIV-1 RNA values outside of the quantitative range.
Methods

We retrospectively evaluated every HIV-1 RNA and paired CD4 cell count performed by the Infectious Diseases Laboratory for all patients tested for both parameters at least once from January 1999 through December 2011 at the Washington DC Veterans Affairs Medical Center. This evaluation included all HIV-infected persons who received care at the clinic without regard to whether the person was prescribed antiretroviral therapy. No charts were reviewed.

The Infectious Diseases Laboratory performs the clinical HIV-1 RNA and CD4 cell counts for our medical center. Written consent was not needed from patients as these tests were performed for clinical indications and not specifically for research purposes. As a clinical laboratory, we maintain databases of these test results. For purposes of the present analyses, we de-identified the datasets. Because we used de-identified, limited datasets, our medical center’s IRB deemed this study to be exempt from the board’s review since it would pose minimal risk for patients’ privacy and data confidentiality.

Figure 1. Distribution of final HIV-1 RNA and CD4 cell counts. (A) Final HIV-1 RNA and (B) final CD4 cell counts by calendar year for 1,814 unique patients tested for both parameters at least once from 1999 through 2011. The number of individuals each year ranged from 575 in 1999 to 854 in 2011. The HIV-1 RNA lower limit of quantitation was 50 copies/mL from 1-1-99 to 10-17-02, 75 copies/mL from 10-18-02 to 3-4-08 and 40 copies/mL from 3-5-08 to 12-31-11.

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Figure 2. Annual and monthly means of HIV-1 RNA and CD4 cell counts. (A) Annual log_{10}-transformed HIV-1 RNA means and CD4 cell count geometric means calculated from each patient’s final values by calendar year. (B) Monthly log_{10} HIV-1 RNA means and (C) CD4 cell count geometric means calculated from all testing in a given month which represented 21–24% of that year’s number of patients (575 persons in 1999 to 854 in 2011). Two-limit Tobit, censored regression models were used to estimate the true mean log_{10} HIV-1 RNA values by month and year. CD4 cell counts of 0 were assigned the value of 1 cell/μL.

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Discussion

This evaluation demonstrates the profound impact on the course of HIV that can be achieved in a broadly diverse inner-city clinic. From the beginning of our observation period in 1999, HIV-1 viral loads improved remarkably such that 72% of our patients measured below the lower limit of quantitation by the end of 2011. Importantly, CD4 cell counts also improved such that 89% of our patients had ≥200 cells/μL by the end of 2011, above the generally recognized threshold for opportunistic infections [23]. Furthermore, our data includes information on persons who were not prescribed antiretroviral treatment based on CD4 cell count or patient-centered issues. Therefore, despite inclusion of these persons in the analysis, the viral burden of the overall clinic population markedly decreased over the 13 years of this evaluation. These gains are particularly meaningful given patients’ significant barriers to success [3–17]. In our clinic, more than half of our veterans have major medical, mental health or substance abuse issues.

Our study has the advantage of using a complete data set from a single inner-city setting to provide comprehensive trends for HIV-1 viral loads and CD4 cell counts within the same observation period and include data through 2011. Earlier analyses focused solely on viral loads [24–26] or CD4 cell counts [27],[28]. A recent publication [29] did analyze HIV-1 viral loads and CD4 cell counts trends for multiple U.S. sites through 2008 and examined median CD4 cell counts only at death. Our data on viral loads can be compared to the information on antiretroviral therapy in the Medical Monitoring Project in United States during 2008–2010 [30]. When using viral suppression as defined by a viral load of <200 copies/mL, 79% of our patients met this criterion, which was comparable to the estimated 77% of persons with HIV suppression in the Medical Monitoring study. However, this study did not include CD4 count monitoring.

The advantages of multidisciplinary teams providing medical and psychosocial support have been documented [14],[31]. Integrated care, pharmacists’ assistance, co-location of mental illness services, and psychosocial well-being have been important in retention to care [32–35], as more HIV-infected veterans reported a lower quality of life compared to non-infected veterans [36]. Our findings highlight that incorporating these approaches to HIV care, can help patients attain the benefits of highly effective antiretroviral treatment. Another valuable tool for us has been the robust electronic medical record of the Veterans Affairs healthcare system that allows the quick review of medication renewals in order to uncover adherence issues during clinic visits.

In conclusion, our clinic patients attained striking improvements in HIV-1 RNA suppression and CD4 cell counts from 1999 through 2011, demonstrating that these goals can be realistically achieved for most individuals in clinical care.

Supporting Information

Model S1 Tobit model description. (DOCX)

Table S1 Distribution of the final HIV-1 RNA for individual patients by calendar year. (DOC)
Table S2  Distribution of the final CD4 cell counts for individual patients by calendar year.

(DOC)

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Author Contributions

Collected and categorized the data: HBG. Interpretation of the data: HBG. Review of the manuscript: HBG. All authors read and approved the final manuscript.

References