Aging and HIV Infection: Epidemiology and Clinical Issues

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2010 Success

- A single combination pill, taken once daily is well tolerated, and achieves viral suppression in >70%
- In many settings >80% of patients are on this treatment and remain on it for extended periods
- Median CD4 counts are increasing and viral load are declining in most clinics
- AIDS defining events are increasingly rare

Life Expectancy on HAART

At HAART	CD4 Cell Count (mm ³⁾		
Initiation	<100	100-199	<u>></u> 200
A 20 yr old will live to (years)	52	62	70
A 35 yr old will live to (years)	62	65	72
% Remaining Life Lost (all ages)	46%	27%	14%

Adapted from ART-CC, Lancet 2008;372:293-99

Epidemiology

Who is living and dying with HIV?

Projected Proportion of those Living With HIV in United States 50+ Years* 2001-2017



*Data from 2008, onward projected based on 2001-2007 trends (calculated by author), 2001-2007 data from CDC Surveillance Reports 2007

Risk Category of Those 50+ Years Diagnosed with HIV in 2007, NY City



As reported to the New York City Department of Health and Mental Hygiene by September 30, 2008.

HIV Among 50+ in NYC 2004-2008

Death Rates⁴ by Age and Race/Ethnicity⁵



HIV Epidemiology & Field Services Semiannual Report, NYCDOH. April 2010

Clinical Issues

New Infections Timing and Response to HAART Non AIDS Conditions and Individualized Care

Sex is Not Only for the Young



Lindau ST. et al. ... Sexuality and Health among Older Adults in the US NEJM 2007 357(8):762–774

Sexual Risks Specific to Older Adults

- Newly single (widowed/divorced) status
- Ratio of men to women increasingly skewed
- Less likely to use condoms
 - Post menopausal women pregnancy no longer possible
 - Men may have erectile dysfunction complicating condom use
- Lower estrogen lead to vaginal dryness and increased risk of virus entering blood stream

New HIV Diagnoses Over Age 50 Years



Diagnoses of HIV Infection and AIDS in the US and Dependent Areas, 2008. HIV surveillance Report, Volume 20, CDC (2010).

Clinical Issues

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Delayed Presentation By Age (NA ACCORD)



Althoff KN. et al. under review, AIDS Research Therapy, presented HIV and Aging Meeting, Baltimore Oct 5 2010

12 Months after HIV Diagnosis by Age, 2007 (37 states with confidential name-based HIV infection reporting)



HIV surveillance Report, Volume 20, CDC (2010)

Immunity in HIV and Aging

- <u>Synergistic</u> effects of HIV and Aging lead to:
 - Reduced naïve T cell numbers
 - Increased levels of "senescent" T cells
 - Reduced naïve CD4 diversity
- These changes accompanied by:
 - Low level immune activation and inflammation
 - In gut , loss of mucosal integrity/microbial translocation, contributes to inflammation

CD4 Response to cART by Age



Althoff KN et al. AIDS 2010 24:2469-2479

Clinical Issues

New Infections Timing and Response to HAART Non AIDS Conditions and Individualized Care

Non AIDS Events Are Associated with HIV Disease Progression*

	Treatment Sparing	Treatment Intensive	Total
All Cause Death	55	30	85
Serious OI	13	2	15
Nonserious OI	63	18	81
Major CAD, Renal,	65	39	104
or Liver Disease			

*More AIDS <u>and</u> "Non-AIDS" Events Among Treatment Sparing Arm (HR 1.7 in SMART) *NEJM 2006;355:2283-96*

HANA Events Among HIV Infected and Demographically Matched Uninfected



Likely To Increase with Aging

- Effects of chronic "inflammation"
 - Liver disease, lung disease, bone marrow suppression
 - Vascular disease, renal disease
 - Cancer
- Multimorbidity
 - Adverse events from poly pharmacy
 - Organ system injury from multiple causes (frailty)
 - Increased demand for supportive care services
- Need to prioritize care based on individual risk, patient preferences, and likely effectiveness

Final Common Pathway



Justice AC. HIV and Aging: Time for a New Paradigm. Curr HIV/AIDS Rep. 2010 May;7(2):69-76

Veterans Aging Cohort Study Risk Index (VACS Index)

- Composed of age and laboratory tests currently recommended for clinical management
 - <u>HIV Biomarkers</u>: HIV-1 RNA, CD4 Count, AIDS defining conditions
 - "non HIV Biomarkers": Hemoglobin, hepatitis C, composite markers for liver and renal injury
- Developed in US veterans, validated in Europe and North America



VACS Index Highly Predictive of Long Term (5 Year) All Cause Mortality



Justice, AC. et. al, HIV Med. 2010 Feb;11(2):143-51. Epub 2009 Sep 14.

Justice AC. HIV and Aging: Time for a New Paradigm. Curr HIV/AIDS Rep. 2010 May;7(2):69-76

MICU Admission Over 6 Years

Kaplan-Meier Survival Estimates



Akgun K. et al. American Thoracic Society 2010 . A5199

Clinical Summary

- HIV, age, and substance use increase risk of 'non AIDS' conditions
- What is common for those aging with HIV is not identical to uninfected individuals
- Guidelines for aging-related non-AIDS condition require adaptation for those with HIV
 - HANA may justify earlier or more aggressive HAART
 - HAART may cause some conditions, but effects are often less than those of HIV itself
 - Some primary care guidelines may be contra-indicated due to reduced life expectancy and polypharmacy

What Can We Do Now?

- Universal HIV screening and early treatment
- Study joint effects of aging, HIV, substance use to identify ways to intervene
 - Compare effectiveness of intervention on overall risk
 - Tailor HIV treatment and primary care accordingly
- Train those who work with:
 - the aging on special issues surrounding HIV
 - with HIV on special issues surrounding aging



Veterans Aging Cohort Study

- <u>PI and Co-PI</u>: AC Justice, DA Fiellin
- Scientific Officer (NIAAA): K Bryant
- <u>Participating VA Medical Centers</u>: Atlanta (D. Rimland), Baltimore (KA Oursler, R Titanji), Bronx (S Brown, S Garrison), Houston (M Rodriguez-Barradas, N Masozera), Los Angeles (M Goetz, D Leaf), Manhattan-Brooklyn (M Simberkoff, D Blumenthal, H Leaf, J Leung), Pittsburgh (A Butt, E Hoffman), and Washington DC (C Gibert, R Peck)
- <u>Core Faculty</u>: K Akgun, S Braithwaite, C Brandt, K Bryant, R Cook, K Crothers, J Chang, S Crystal, N Day, R Dubrow, M Duggal, J Erdos, M Freiberg, M Gaziano, M Gerschenson, A Gordon, J Goulet, N Kim, M Kozal, K Kraemer, V LoRe, S Maisto, K Mattocks, P Miller, P O'Connor, C Parikh, C Rinaldo, J Samet
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- <u>Major Collaborators</u>: VA Public Health Strategic Healthcare Group, VA Pharmacy Benefits Management, Massachusetts Veterans Epidemiology Research and Information Center (MAVERIC), Yale Center for Interdisciplinary Research on AIDS (CIRA), Center for Health Equity Research and Promotion (CHERP), ART-CC, NA-ACCORD, HIV-Causal
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Reference List for bar graphs of HANA incidence rates

(1) Bedimo RJ, McGinnis KA, Dunlap M, Rodriguez-Barradas MC, Justice AC. Incidence of Non-AIDS-Defining Malignancies in HIV-Infected Versus Noninfected Patients in the HAART Era: Impact of Immunosuppression. J Acquir Immune Defic Syndr. 2009.

(3) Lucas GM, Mehta SH, Atta MG, Kirk GD, Galai N, Vlahov D et al. End-stage renal disease and chronic kidney disease in a cohort of African-American HIV-infected and at-risk HIV-seronegative participants followed between 1988 and 2004. AIDS. 2007;21:2435-43.

(4) Fischer MJ, Wyatt CM, Gordon K, Gibert CL, Brown ST, Rimland D et al. Hepatitis C and the risk of kidney disease and mortality in veterans with HIV. J Acquir Immune Defic Syndr. 2010;53:222-26.

(5) Justice AC, Zingmond DS, Gordon KS, Fultz SL, Goulet JL, King JT, Jr. et al. Drug toxicity, HIV progression, or comorbidity of aging: does tipranavir use increase the risk of intracranial hemorrhage? Clin Infect Dis. 2008;47:1226-30.

(6) Sico, J., Chang, CC, Freiberg, M., Hylek, E, Butt, A., Gibert, C., Goetz, M. B., Rimland, D, Kuller, L., Justice AC, and for the VACS Project Team. HIV Infection and the Risk of Ischemic Stroke in the VACS VC. SGIM 2010 Poster . 4-28-2010.

Ref Type: Abstract

(7) Womack, J., Goulet, J., Gibert, C., Brandt, C., Mattocks, K., Rimland, D, Rodriquez-Barradas, M. C., Tate, J., Yin, M., Justice, A., and and VACS Project Team. HIV Infection and Fragility Fracture risk among Male Veterans. CROI 2010 Presentation (Abstract #129). 2-18-2010.

Ref Type: Abstract

(8) Thorpe J. et al. CROI 2010 absract # 683

(9) Thein HH. Et al. Natural history of Hepatitis C virus infection in HIV-infected individuals and the impact of HIV in te era of HAART: a meta-analysis. AIDS 2008;22:1979-91.

(10) Crothers K, Huang L, Goulet J, Goetz MB, Brown S, Rodriguez-Barradas M et al. HIV Infection and Risk for Incident Pulmonary Diseases in the Combination Antiretroviral Therapy Era. AJRCCM. 2010; [In press].