



HIV 101: Overview of HIV Epidemiology, Treatment and Outcome

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Learning Objectives:

- 1. Identify the virus that causes HIV
- 2. Explain the natural course of HIV infection
- 3. Understand the treatments associated with HIV
- 4. Discuss trends in epidemiology associated with HIV

No disclosures

HIV: what is it?

Human immunodeficiency virus



HIV: what is it?

- Human immunodeficiency virus
- Infects only humans
 - \rightarrow Research Studies Limited
 - \rightarrow Exposure Risks

HIV Transmission

- The body fluids that transmit HIV
 - Blood
 - Semen
 - Vaginal Fluids
 - Breast Milk
- High Risk Behaviors
 - Unprotected sex
 - Injection Drug use
- Mother to child during pregnancy, delivery, and breast feeding

HIV: what it is

- Human immunodeficiency virus
- Disease is caused by weakened immune system
- Immune system protects from infections
 Increased risk of infections

HIV: what it is

- Human immunodeficiency virus
- Like all viruses, it needs living cells to make copies of itself and spread
 - → Not hardy outside the body
- The type of virus called a retrovirus
 - Viral RNA works with cells DNA to make more Viral RNA, and packages it as new HIV viruses
 - \rightarrow Long latent phase
 - \rightarrow Potential for increased risk cancers

Life Cycle of HIV-1



Natural History of HIV Infection



What is the difference between HIV and AIDS ?

- Human Immunodeficiency Virus
- AIDS: Acquired Immunodeficiency Disease Syndrome, the disease caused by the virus
 - → Acquired = from an other human
 - → Immunodeficiency = weakened immune system CD4 Cell count is 200 or less
 - → Syndrome: meeting the CDC Criteria for an "AIDSdefining illness"





- Candidiasis of bronchi, trachea, or lungs
- Candidiasis, esophageal
- Coccidioidomycosis, disseminated or extrapulmonary
- Cryptococcosis, extrapulmonary
- Cryptosporidiosis, chronic intestinal
- Cytomegalovirus disease
- Encephalopathy, HIV related
- Herpes simplex: chronic ulcers, pneumonitis, or esophagitis
- Histoplasmosis, disseminated or extrapulmonary
- Isosporiasis, chronic intestinal
- Kaposi sarcoma
- Lymphoid interstitial pneumonia
- Lymphoma, Burkitt or immunoblastic
- Lymphoma, primary, of brain
- Mycobacterium avium complex or M. kansasii, disseminated or extrapulmonary
- Mycobacterium tuberculosis, of any site
- Pneumocystis jirovecii pneumonia (PCP)
- Pneumonia, recurrent
- Progressive multifocal leukoencephalopathy
- Salmonella septicemia, recurrent
- Toxoplasmosis of brain
- Wasting syndrome due to HIV

Historic Milestones in Treatment of HIV

1983Isolation of the virus

1985

1995

- AZT and nucleoside RT inhibitors
- 1990 Non-nucleoside RT inhibitors
- **1990's Combination NRTI therapy**
- **1993 Protease inhibitors**

Combination of 2+ drug classes, aka: HAART- Highly Active Antiretroviral therapy cART- Combination Antiretroviral therapy "the cocktail"

Goals of Therapy



Increase or Maintain:

- Viral suppression
- CD4 cell count
- Good Health
- Quality of life

Decrease:

- Toxicity
- Drug resistance
- Overall cost of care

Life Cycle of HIV-1 and Mechanisms of Action of Antiretroviral Drugs



Reverse Transcriptase Inhibitors

Year approved*	Generic Name	Trade Name
1987	Zidovudine	Retrovir
1991	Didanosine	Videx
1992	Zalcitabine, Hivid (ddC, dideoxycytic	dine) by Roche: Mar
1994	Stavudine	Zerit
1995	Lamivudine	Epivir
1997	Zidovudine/Lamivudine	Combivir
1998	Abacavir	Ziagen
2000	Zidovudine/Lamivudine/Abacavir	Trizivir
2001	Tenofovir	Viread
2003	Emtricitabine	Emtriva
2004	Abacavir/Lamivudine	Epzicom
2004	Emtricitabine/Tenofovir	Truvada

NRTI

NNRTI

1996	Nevirapine	Viramune
1997	Delavirdine	Rescriptor
1998	Efavirenz	Sustiva
2008	Etravirine	Intelence
2011	Rilpivirine	Edurant

Protease Inhibitors

1995	Saquinavir	Invirase	SQV	
1996	Ritonavir	Norvir	RTV	
1996	Indinavir	Crixivan	IDV	
1997	Nelfinavir	Viracept	NFV	
1997	Saquinavir	Fortovase Manufacture	Fortovase Manufacture discontinued in 2006; Roche	
1999	Amprenavir	Agenerase Manufactur	Agenerase Manufacture discontinued in 2007	
2000	Lopinavir/ritonavir	Kaletra, Aluvia	LPV	
2003	Atazanavir	Reyataz	ATV	
2003	Fosamprenavir	Lexiva	FPV	
2005	Tipranavir	Aptivus	TPV	
2006	Darunavir	Prezista	DRV	

Integrase Inhibitors

2007 Raltegravir	Isentress	RGV
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Fusion Inhibitors & Chemokine Coreceptor Antagonists

2003	Enfuvirtide	Fuzeon	T-20
2007	Maraviroc	Selzentry, Celsentri	MVC

Metabolic and Morphologic Complications of HIV and HAART

Morphologic

- Fat accumulation
- Fat loss

<u>Others</u>

- Osteoporosis
- Osteopenia
- Osteonecrosis

Metabolic

- Dyslipidemias
 - hypercholesteremia
 - hypertriglyceridemia
- Impaired glucose tolerance/ Diabetes
- Lactic acidosis

Viral Suppression and Adherence



PI adherence, % (electronic bottle caps)

Paterson, et al. 2000

Reasons for Non-Adherence: Clinician vs Patient Views



Chesney M. Adherence to antiretroviral therapy. 12th World AIDS Conference, 1998; Geneva. Lecture 281

Probability of ADI without ART by Baseline CD4 and Viral Load

CD4	HIV RNA	3 years	9 years
<350	<20,000	8%	66%
<350	>55,000	73%	96%
350-500	<20,000	6%	61%
350-500	>55,000	48%	94%
>500	<20,000	7%	50%
>500	>55,000	33%	76%

The Balance of Antiretroviral Therapy

Reduce Viral Load Increase CD4 cells Prevent AIDS Prolong life Reduce transmission Not cure HIV infection Drug resistance Toxicities / side-effects Access to medication & care Monitoring Cost

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%

Slide courtesy A. Justice, ART-CC, Lancet 2008;372:293-99

Epidemiology of HIV: the Changing Face of the Epidemic



The NEW ENGLAND JOURNAL of MEDICINE



Perspective

HIV and Aging - Preparing for the Challenges Ahead

Edward J. Mills, Ph.D., Till Bärnighausen, M.D., Sc.D., and Joel Negin, M.I.A. N Engl J Med 2012; 366:1270-1273 April 5, 2012

Article

By 2015, half the U.S. population living with human immunodeficiency virus (HIV) infection will be older than 50 years of age. As antiretroviral therapy (ART) coverage continues to expand worldwide, this aging of the HIV epidemic will be mirrored in developing countries. In sub-Saharan Africa, ART has already reduced mortality rates, with 320,000 (or 20%) fewer people dying of HIV-related causes in 2009 than in 2004.1 Currently, HIV-infected Ugandans in their 40s who are receiving ART can expect to live well into their 60s.2 The increased life expectancy of HIV-infected persons will lead to increases in HIV prevalence among older adults. Approximately 1 in 8 HIV-infected adults and 1 in 10 patients receiving ART in sub-Saharan Africa are older than 50 years of age,3 and these ratios are likely to increase manyfold in the coming decades

Proportion Patients With HIV 50+ Years of Age in United States 2001-2017



State of Care for Veterans with HIV/AIDS: 2009 Report

Figure 5. HIV Infected Veterans in VHA Care 2008 - Age by Decade of Life



State of Care for Veterans with HIV/AIDS: 2009 Report

Figure 4a/b. Comparison of Race/Ethnicity for New AIDS Cases Reported to the CDC in 2007 and All Veterans with HIV/AIDS in Care in 2008.





>50% of Deaths Attributed to Non-AIDS Events



Cumulative Mortality by COD Among Those on cART (1996-2006) ART-CC, CID 2010: 1387-1396

Incident Non AIDS Events by HIV Group



1995: 46 year old man diagnosed HIV 6 more years injection drug use



2001: Depression and SA treatment 2003: 54 yo after 2 yrs HAART



2004: 55 yo with DM, obesity



2005: healthy and fit 56 year old



Healthy Aging with HIV



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- 1. Identify the virus that causes HIV
- \rightarrow Requires body fluid for transmission
- \rightarrow Infection and destruction of CD4 cells
- \rightarrow HIV vs. AIDS: infection vs disease

- 2. Explain the natural course of HIV infection
- → Long latency period
- \rightarrow CD4 Cells below- loss of viral control
- → AIDS Defining Illness

- 3. Understand the treatments associated with HIV
- \rightarrow Multiple drug classes in effective combination
- \rightarrow Side effects and metabolic toxicity
- → Adherence is key to prevent drug resistance
- → Individualized balancing Act

- 4. Discuss trends in epidemiology associated with HIV
- → Increased Life Expectancy
- \rightarrow Age as the new wrinkle in the epidemic
- → Non-HIV related morbidity and mortality

www.HIV.va.gov

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	HEALTH CARE PROVIDERS	The Image Library More than 200 images of dinical symptoms of HV infection Go to library 5 HEALTH CARE PROVIDERS For VETERANS and the PUBLIC

Additional Web resources

 Patient education material aidsinfonet.org aidsmeds.com

aidsinfo.nih.gov

See education/materials/glossary

- aids-ed.org
- hivinsite.org
- hivandhepatitis.com