# The History of HIV

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Conflicts of interest: None

Pneumocystis Pneumonia --- Los Angeles

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Weekly

June 5, 1981 / 30(21);1-3

#### **Epidemiologic Notes and Reports**

#### Pneumocystis Pneumonia --- Los Angeles

In the period October 1980-May 1981, 5 young men, all active homosexuals, were treated for biopsy-confirmed *Pneumocystis carinii* pneumonia at 3 different hospitals in Los Angeles, California. Two of the patients died. All 5 patients had laboratory-confirmed previous or current cytomegalovirus (CMV) infection and candidal mucosal infection. Case reports of these patients follow.

Patient 1: A previous associated with eleva \* The patient's condi and postmortem exar

CDC: MMWR June 5, 1981 9 MSM with PCP

CDC: MMWR July 3, 1981
MSM with PCP or KS

2-month history of fever as 256; in may 1981 it was 32. d acyclovir. He died May 3,

#### The Science-Director of NIAID made this a new almost singular priority

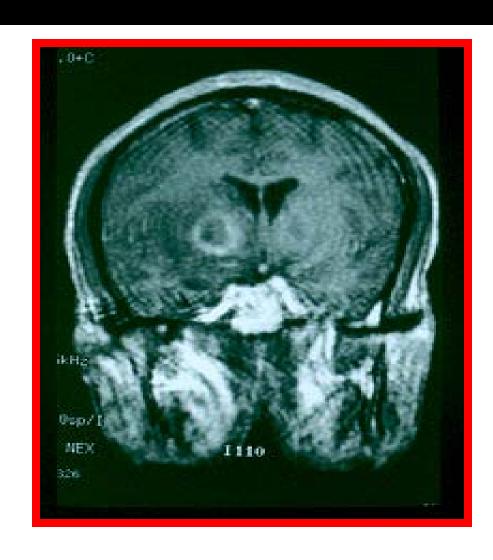


"Goose pimples" 1981

### PNEUMOCYSTIS PNEUMONIA



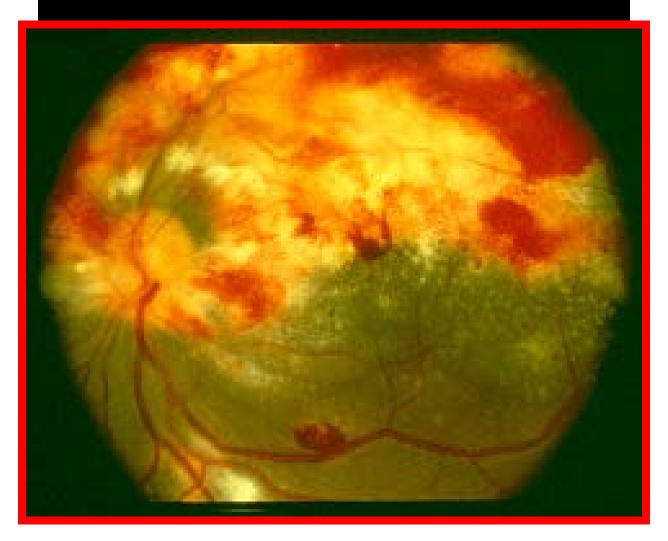
### **TOXOPLASMOSIS**



## **KAPOSI SARCOMA**



## **CMV RETINITIS**



# WHO WINNING PHOTO AWARD: AIDS – THE HUMAN TRAGEDY



## The stigma- Ryan White



**Child with** hemophilia - HIV discovered 12/17/84 banned from school and given 6 months to live. Died 5 years later, 4 months before passage of the Ryan White Care Act

# Government leadership C. Everett Koop

"The Letter" 1987



**Surgeon General under President Reagan** Champion of -Sex education **Condoms** Handicapped children Cigarette harm "The Letter" was an

AIDS wakeup call

# The Response: AIDS COALITION to UNLEASH POWER: 1987 ACT-UP



Civil disobedience: Wall Street 1987, NY Post Office 1987, St. Patrick's Cathedral 1989, NIH 1990 (Now TAG)

# Martin Delaney HIV Activist 1945-2008



Founded Project Inform "for medically supervised guerrilla trials" - Major force in NIAID, FDA and ethics ("First in the tent")

#### **MAGIC JOHNSON**

Olympic hero & greatest guard in NBA history
The Announcement: 1991



"I have AIDS"

Trumped efforts of millions to destigmatize HIV infection

#### The beginning of ART: Jerome Horwitz 1919-2012

Discovered AZT (1964) for cancer Then ddl & d4T → failed & shelved "Great drugs waiting for the right disease"

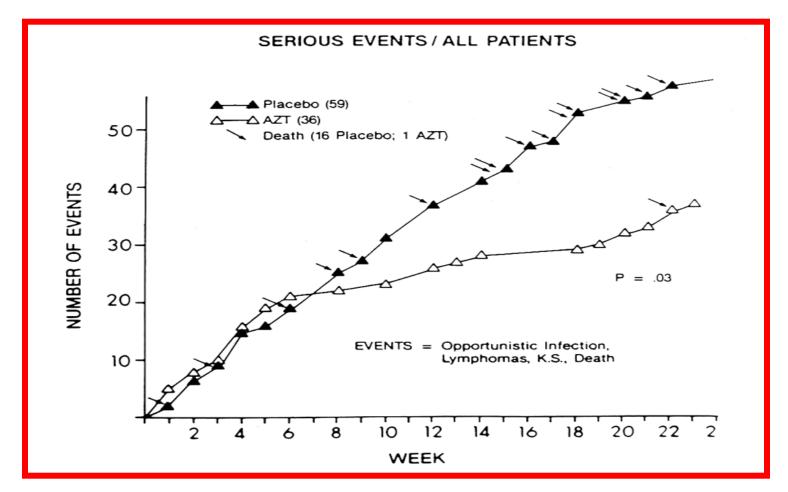


Broder, Mitsuya and Yarchoan (NCI) showed anti-HIV action in CD4 cells



**AZT trial** → **First HIV** → **FDA** record for approval: one cohort, 282 patients, stopped at 6 months





The Trial: AZT vs. placebo (n=282).

DSMB stopped study: 19 deaths
(placebo) vs 1 (AZT)

(Fischl MA. NEJM 1987;317:185)

#### Agency Hero: Jeff Murray, MD

Deputy Director
Division of Antimicrobial Drug Products for FDA



Major facilitator of HIV drugs (n=28) and, more recently HCV (#37 approved or in trials)

# First ART Success: ACTG 076 AZT to prevent perinatal transmission



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#### ORIGINAL ARTICLE

Volume 331:1173-1180 November 3, 1

**November 3, 1994** Number 18

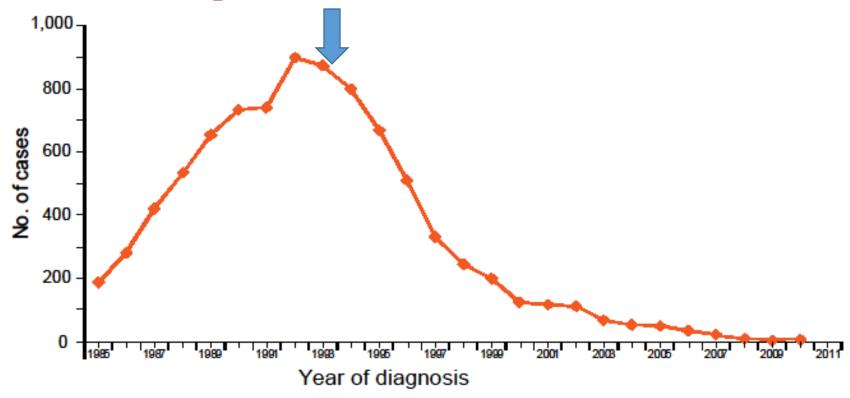
Next ►

#### Reduction of Maternal-Infant Transmission of Human Immunodeficiency Virus Type 1 with Zidovudine Treatment

Edward M. Connor, Rhoda S. Sperling, Richard Gelber, Pavel Kiselev, Gwendolyn Scott, Mary Jo O'Sullivan, Russell VanDyke, Mohammed Bey, William Shearer, Robert L. Jacobson, Eleanor Jimenez, Edward O'Neill, Brigitte Bazin, Jean-Francois Delfraissy, Mary Culnane, Robert Coombs, Mary Elkins, Jack Moye, Pamela Stratton, James Balsley, for The Pediatric AIDS Clinical Trials Group Protocol 076 Study Group

Perinatal tx rates: Placebo-23%; AZT-8% (p=0.0006)

# Estimated Numbers of Perinatally Acquired AIDS Cases by Year of Diagnosis, 1985–2010 — United States and Dependent Areas

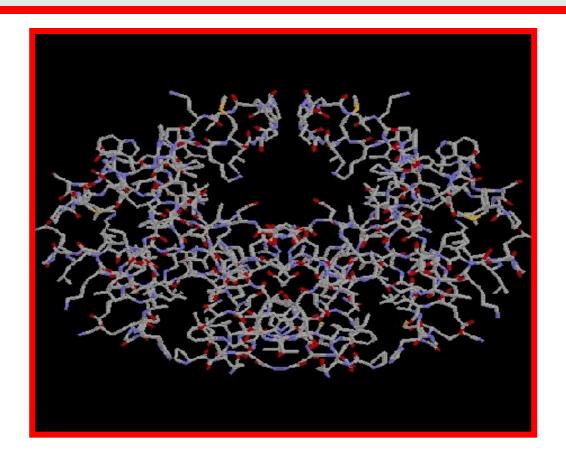


Note: Data have been adjusted for reporting delays and missing risk-factor information.





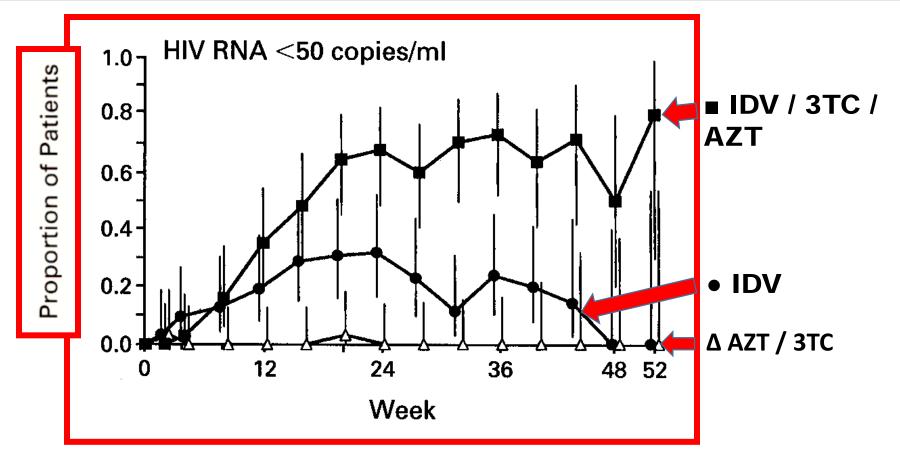
### PROTEASE 1996



The development of protease inhibitors and beginning of HAART

#### THE STUDY THAT CHANGED AIDS

Gulick RM. Merck 035 NEJM 1997;337:734



**HAART: IDV/AZT/3TC** 

Results for NDV at 52 weeks - 80% vs. zero

# David Ho: Post Vancouver IAS Meeting



"Hit hard and hit early"
(3 ART agents for all with HIV starting at diagnosis)

Time Magazine:

Man-of-the-year, 1996

	NRTI	NNRTI	PI	El	CCR5	Ш
1987	AZT					
1991-92	ddl, ddC					
1995	d4T					
1996	3TC		SQV			
1997		NVP	RTV, IDV			
1997		DLV	NFV			
1998		EFV				
1999	ABC		APV			
2000			LPV			
2001	TDF					
2003	FTC		ATV	ENF		
2005			TPV			
2006			DRV			
2007					MVC	RAL
2008		ETR				
2011		RPV				
2012						EVG
2013						DTG

# RECOMMENDATIONS FOR WHEN TO START ART

(DHHS Panel on ART 1996-2012)

Year	Who by CD4 count		
1996	CD4 <500: Merck 035		
2000	CD4 <200: Toxicity/Resis		
2006	CD4 <350: Half between		
2009	CD4 <500: 14 ART drugs		
2012	All pts; Any CD4 count		

# The START Trial

Trial: Start ART at <CD4 350 vs >500

Sample size: 4400 in 230 clinics, 36 countries

Started: 1/1/14; first data review 2016

Web statement: Early start is based on

"embarrassingly uninterpretable studies like NA-ACCORD"

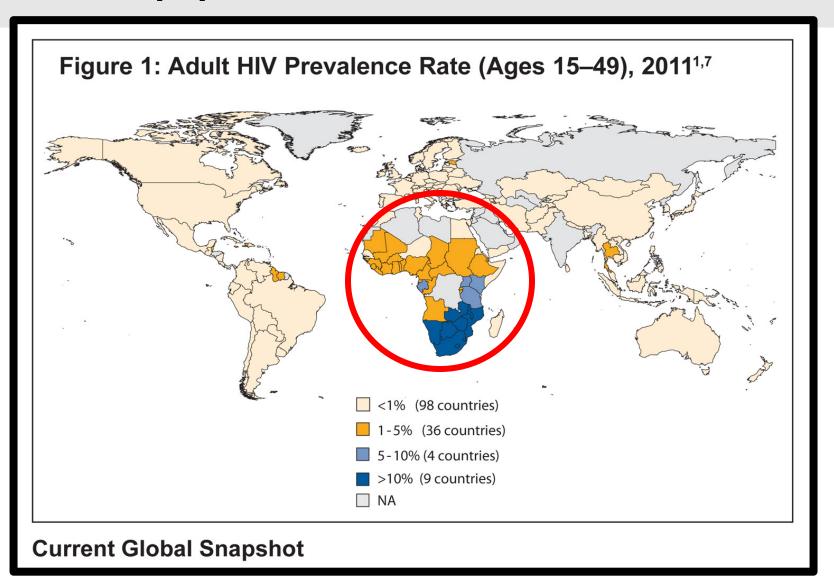
**Early review, May 2015**: Early ART clearly superior (53% reduction in HIV complications)

# Average duration of life after AIDS diagnosis in a 25 year old man

Year	Age of Death	Added years
1995	28 years	3
1997-99	48 years	25
2000-05	64 years	39
2011-12	73 years*	48

<sup>\*</sup> Moore Clinic

# GLOBAL HIV- Subsaharan Africa:12% global population 69% of HIV cases



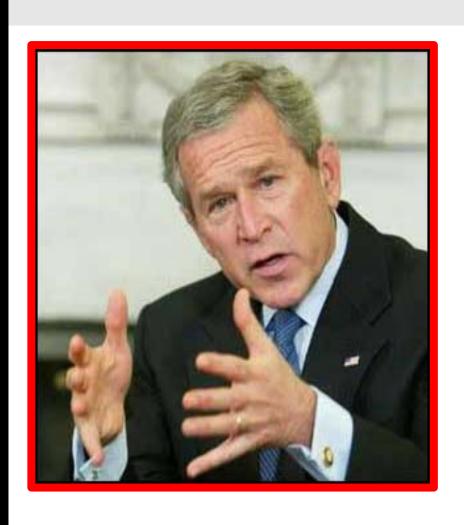
## Zimbabwe: Lifetime Probability of AIDS- 60%!



# White House meeting of President Bush & Joe O'Neill



# President George W. Bush PEPFAR: 2002



**Bush:** Do those HIV drugs work?

O'Neil: "Let me tell you about my morning in the Moore Clinic"

Bush: "Get Tony, we need to do something. Think big - really big"

# President George W. Bush



Fauci: "Thinking big"

**Collaborators** -

Dybul, Mugyeni, Goosby, Pape

Plan Proposed
Would be
considered

**January 2003**:

"Seldom have we been offered the opportunity to do so much for so many"

#### Peter Mugyeni and Laura Bush State of the Union Address



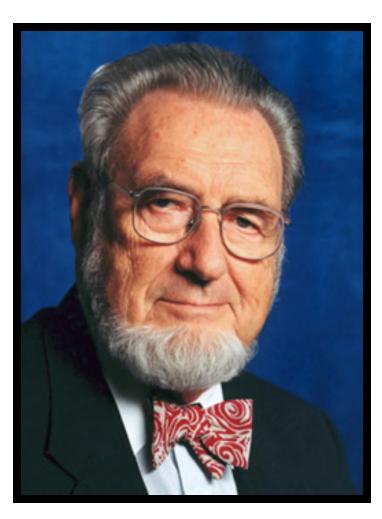
Bush: "\$15 billion for AIDS in Africa" Mugyeni: Jumped up → cheered → hugged the First Lady

#### Tony Fauci - Medal of Freedom



President Bush awarding Dr. Fauci the Presidential Medal of Freedom - June 18, 2008

# Everett Koop Award "The boys from Brooklyn"



2011

First C. Everett Koop Award - Dr. Fauci (Forum, 2011)

2012

Second Award - from Dr. Fauci to Dr. Koop (Forum, 2012)

#### JB- Personal HIV History at Hopkins

1980: recruited to Hopkins to start an ID Division and continue C difficile work 1983: Frank Polk changed career-irreversible passion for starting an HIV program 1983: HIV program launched: Moore Clinic quickly filled & hidden in basement 1986: Hospital funded HIV in-pt Osler- 8 1988: Cardiology fellow- needlestick AIDS 1991: Hopkins surgeon dies with HIV; Test 1000: 1 positive; chair incident

2014: O8 closed & converted to Ebola Center

# What was good about it

Filled a desperate need in tradition of Osler & J Hopkins

1984: Dr. Heyssel requested AIDS Program plan & budget

1984-94: US News & World Report- Hopkins always #2

**1985**: Lambert

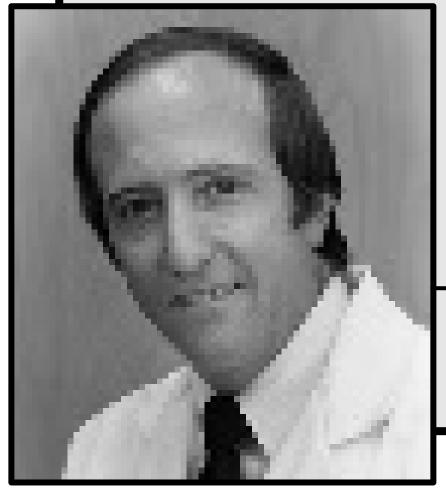
1989: AIDS Busters Softball team won the Hopkins league beating the Maintenance men in the playoffs 4-2

1993: 10th anniversary – Bo Diddly celebration

1983-2006: 24 faculty promoted-Prof based on work in 14 depts-Ob, Neuro, Basic, Card, Psych, Onc, Peds, Pharm, TB, HCV

### FRANK POLK, MD

The 1983 Career change



"There's this new disease. Someone will find the virusnot me. BUT **Epidemiology-**That's for me" Started MACS and **ALIVE cohorts:>1000** publications

# Dr. Robert Heyssel:



President JHH-Birth of Osler 8 "It doesn't matter (an AIDS Ward). It's a moral imperative" 1986



#### Osler 8 nurses:

1986-2013: Perennial best nursing unit-JHH. Unique service for knowledge of HIV and this patient population

# Thomas Quinn, MD Director of Johns Hopkins Center for Global Health.



Author of > 1000 publications

34 global HIV Centers, 450 faculty, 710 projects in 110 countries funded at \$75M/yr

# David Thomas, MD



CHIEF, Infectious Disease Division & international leader- HCV basic science

### Mark Sulkowski, MD

National leader in HCV and HIV/HCV treatment



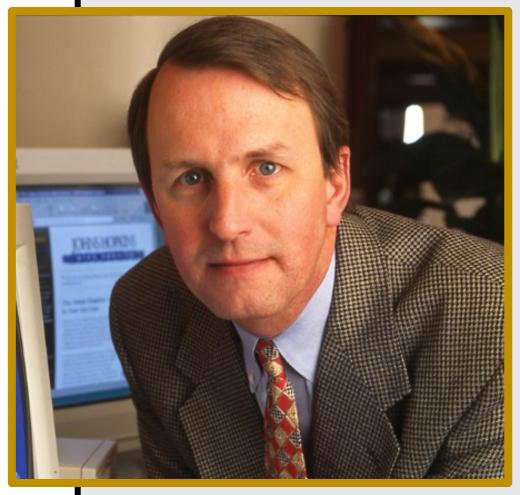
Director, Hopkins Center for Viral Hepatitis

#### Joel Gallant, MD



(Arguably) HIV's best clinician; National and international HIV guidelines, **President HIV Med Assoc** 

# Richard Chaisson, MD



Co- PI JHU CFAR; **InternationaL** leader- TB & HIV \* Chair-NIH TB Clin TRIAL Group \*TB trials-INDIA Brazil, Africa \*CHAIR-CREATE (TB-GATES)

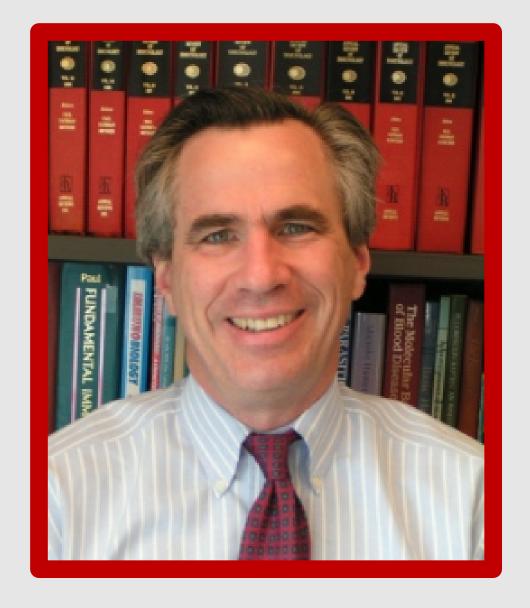
# RICHARD MOORE, MD Internal Medicine



Master of design, management and analysis of cohorts.

PI of NA-ACCORD, NIH- sponsored, 8 yrs, 25 cohorts, >200 sites, 150,000 patients

## Robert Siliciano, MD, PhD



H. Hughes Investigator, Leading scientist in HIV basic research and "the Cure"

### WHAT HAS HIV TAUGHT US

- Science leadership: A. Fauci
- Pharma leadership: Merck Indinavir
- Oslerian Medicine: HIV providers < 1997</li>
- Greatest humanitarian effort: PEPFAR (Bono)
- Government leadership: E. Koop
- Implementation science: 076

# "AIDS Busters" Softball Championship vs Maintenance Department, 1989



**Big Challenges Build Strong Bonds**