

# HIV ve AIDS

## Geeçmişten Günümüze

Deniz Gökengin  
Ege Üniversitesi Tıp Fakültesi  
Enfeksiyon Hastalıkları ve Klinik  
Mikrobiyoloji Anabilim Dalı

# 1981



CDC Home | Search | Health Topics A-Z

# MMWR<sup>®</sup>

Weekly

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

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## 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 previously healthy 33-year-old man developed *P. carinii* pneumonia and oral mucosal candidiasis in March 1981 after a 2-month history of fever associated with elevated liver enzymes, leukopenia, and CMV viremia. The serum complement-fixation CMV titer in October 1980 was 256; in May 1981 it was 32.\* The patient's condition deteriorated despite courses of treatment with trimethoprim-sulfamethoxazole (TMP/SMX), pentamidine, and acyclovir. He died May 3, and postmortem examination showed residual *P. carinii* and CMV pneumonia, but no evidence of neoplasia.

Patient 2: A previously healthy 30-year-old man developed *P. carinii* pneumonia in April 1981 after a 5-month history of fever each day and of elevated liver-function tests, CMV viremia, and documented seroconversion to CMV, i.e., an acute-phase titer of 16 and a convalescent-phase titer of 28\* in anticomplement immunofluorescence tests. Other features of his illness included leukopenia and mucosal candidiasis. His pneumonia responded to a course of intravenous TMP/SMX, but, as of the latest reports, he continues to have a fever each day.

Patient 3: A 30-year-old man was well until January 1981 when he developed esophageal and oral candidiasis that responded to Amphotericin B treatment. He was hospitalized in February 1981 for *P. carinii* pneumonia that responded to TMP/SMX. His esophageal candidiasis recurred after the pneumonia was diagnosed, and he was again given Amphotericin B. The CMV complement-fixation titer in March 1981 was 8. Material from an esophageal biopsy was positive for CMV.

# 1981

## RARE CANCER SEEN IN 41 HOMOSEXUALS

Outbreak Occurs Among Men  
in New York and California  
—8 Died Inside 2 Years

By LAWRENCE K. ALTMAN.

Doctors in New York and California have diagnosed among homosexual men 41 cases of a rare and often rapidly fatal form of cancer. Eight of the victims died less than 24 months after the diagnosis was made.

The cause of the outbreak is unknown, and there is as yet no evidence of contamination. But the doctors who have made the

**"What's it like  
to have  
Kaposi's  
sarcoma?"**

***It's a bummer."***

It's a bummer being thirty years old and having a rare, often-fatal cancer seeping. Friends often visit and die. It's a bummer going through the rounds of tests and treatments. Doctors use to diagnose and treat cancer. It's a bummer running up a bill of \$10,000 or more for a month of chemo. It's a bummer not knowing what caused this cancer or if it can be cured.

It's a bummer you can't move well. I don't feel sick. My cancer hasn't spread. I still function pretty much.

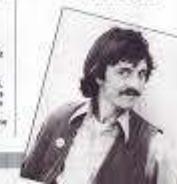
Also, I have a good support system — a lover, a therapist, understanding parents, lots of friends. That's important insurance and disability insurance.

Even so, I sometimes get real depressed. This thing could kill you. You could die alone, alone, and hundreds of other brothers that I don't know personally. I don't know if I'll live to be 40.

Am I thinking, "This can't happen to me?" I didn't think it could happen to me, either. But it did.

The last thing that underlines AIDS and other "immune diseases" is that the public's immune system these cold fight off disease has somehow been weakened. No one knows for sure why this is happening.

—Bobby Campbell



A20

L

## RARE CANCER SEEN IN 41 HOMOSEXUALS

Outbreak Occurs Among Men

**A Pneumonia  
That Strikes  
Gay Males**

A mysterious outbreak of a sometimes fatal pneumonia among gay men has occurred in San Francisco and several other major cities. It was revealed yesterday.

**24 Eylül 1982**

ilk kez **AIDS** terimi kullanıldı

# 1983



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## MMWR<sup>®</sup>

Weekly

January 07, 1983 / 31(52):697-8

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## Epidemiologic Notes and Reports Immunodeficiency among Female Sexual Partners of Males with Acquired Immune Deficiency Syndrome (AIDS) -- New York

CDC has received reports of two females with cellular immunodeficiency who have been steady sexual partners of males with the acquired immune deficiency syndrome (AIDS).

Case 1: A 37-year-old black female began losing weight and developed malaise in June 1982. In July, she had oral candidiasis and generalized lymphadenopathy and then developed fever, non-productive cough, and diffuse intestinal pulmonary infiltrates. A transbronchial biopsy revealed *Pneumocystis carinii* pneumonia (PCP). Immunologic studies showed elevated immunoglobulin levels, lymphopenia, and an undetectable number of T-helper cells. She responded to antimicrobial therapy, but 3 months after hospital discharge had lymphadenopathy, oral candidiasis, and persistent depletion of T-helper cells.

The patient had no previous illnesses or therapy associated with immunosuppression. She admitted to moderate alcohol consumption, but denied intravenous (IV) drug abuse. Since 1976, she had lived with and had been the steady sexual partner of a male with a history of IV drug abuse. He developed oral candidiasis in March 1982 and in June had PCP. He had laboratory evidence of immune dysfunction typical of AIDS and died in November 1982.

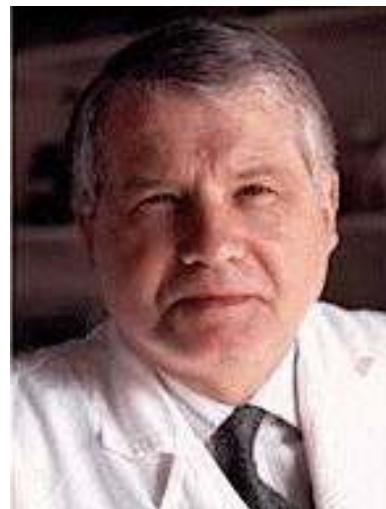
Case 2: A 23-year-old Hispanic female was well until February 1982 when she developed generalized lymphadenopathy. Immunologic studies showed elevated immunoglobulin levels, lymphopenia, decreased T-helper cell numbers, and a depressed T-helper/T-suppressor cell ratio (0.82). Common infectious causes of lymphadenopathy were excluded by serologic testing. A lymph node biopsy showed lymphoid hyperplasia. The lymphadenopathy has persisted for almost a year; no etiology for it has been found.

The patient had no previous illnesses or therapy associated with immunosuppression and denied IV drug abuse. Since the summer of 1981, her only sexual partner has been a bisexual male who

# 1983



Robert Gallo  
Institute of Human Virology



Luc Montagnier  
Pasteur Enstitüsü



Francoise Barre Sinoussie  
Pasteur Enstitüsü

20 Mayıs 1983'de Luc Montagnier ve Francoise Barre Sinoussie LAV'ı izole ettiler

Kasım 1983'de Robert Gallo AIDS etkenini izole etti (daha sonra HTLV III olarak adlandırıldı)

# 1984

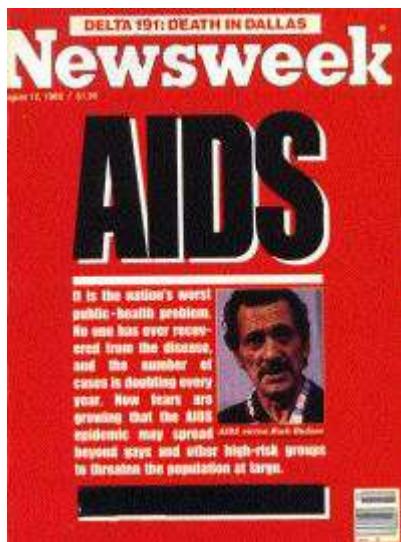
- Robert Gallo ve ekibi HTLV'yi saptayacak tanı testini keşfetti
- Robert Gallo ve Luc Montagnier ortak bir toplantı yaparak buldukları virüslerin birbirinin aynı olduğunu açıkladılar.

# 1985



## 1. Uluslararası AIDS Konferansı Atlanta-Georgia

CDC kanların taranması konusunda bir kılavuz yayımladı.



FDA ilk ticari ELISA testine onay verdi.

Rock Hudson AIDS nedeniyle öldü

# 1986

AIDS etkeni virüsün adı **HIV** olarak belirlendi

# 1987



İlk antiretroviral ilaç FDA onayı aldı

WB tanı kitı FDA onayı aldı

AIDS Anma Battaniyesi Washington DC'de bir alışveriş merkezinde sergilendi  
(1920 panel; günümüzde >48.000)

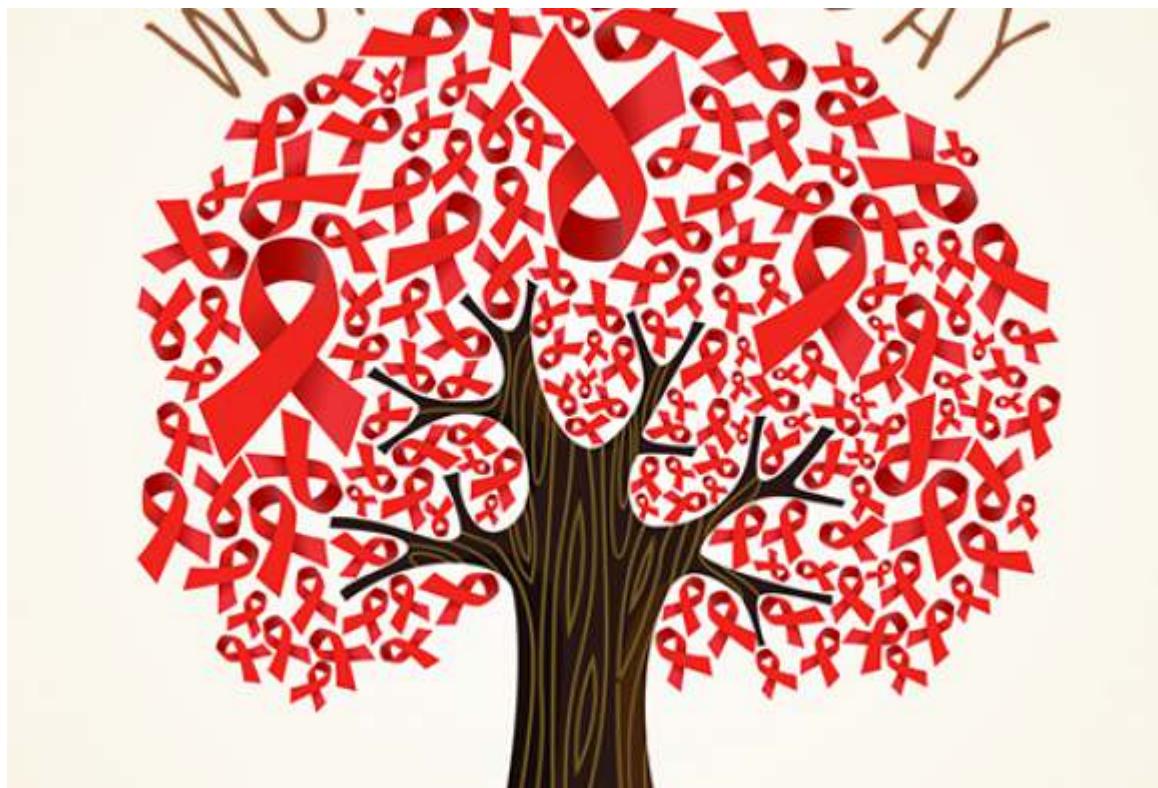


AIDS'e karşı ilk aşısı insanlar üzerinde denenmeye başlandı.

[www.aids.gov](http://www.aids.gov)

[www.aidsquilt.org](http://www.aidsquilt.org)

**1988**



**1 Aralık  
Dünya AIDS Günü**

# 1989



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## MMWR

Supplements

June 16, 1989 / 38(5-5);1-9

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## Guidelines for Prophylaxis Against *Pneumocystis carinii* Pneumonia for Persons Infected with Human Immunodeficiency Virus



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## MMWR

Supplements

June 23, 1989 / 38(5-6);3-37

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## Guidelines for Prevention of Transmission of Human Immunodeficiency Virus and Hepatitis B Virus to Health-Care and Public-Safety Workers A Response to P.L. 100-607 The Health Omnibus Programs Extension Act of 1988

The material in this report was developed by the National Institute for Occupational Safety and Health in collaboration with the Center for Infectious Diseases, Centers for Disease Control.

Introduction

# 1990



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## MMWR

Recommendations and Reports

January 26, 1990 / 39(01);1-14

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## Public Health Service statement on management of occupational exposure to human immunodeficiency virus, including considerations regarding zidovudine postexposure use

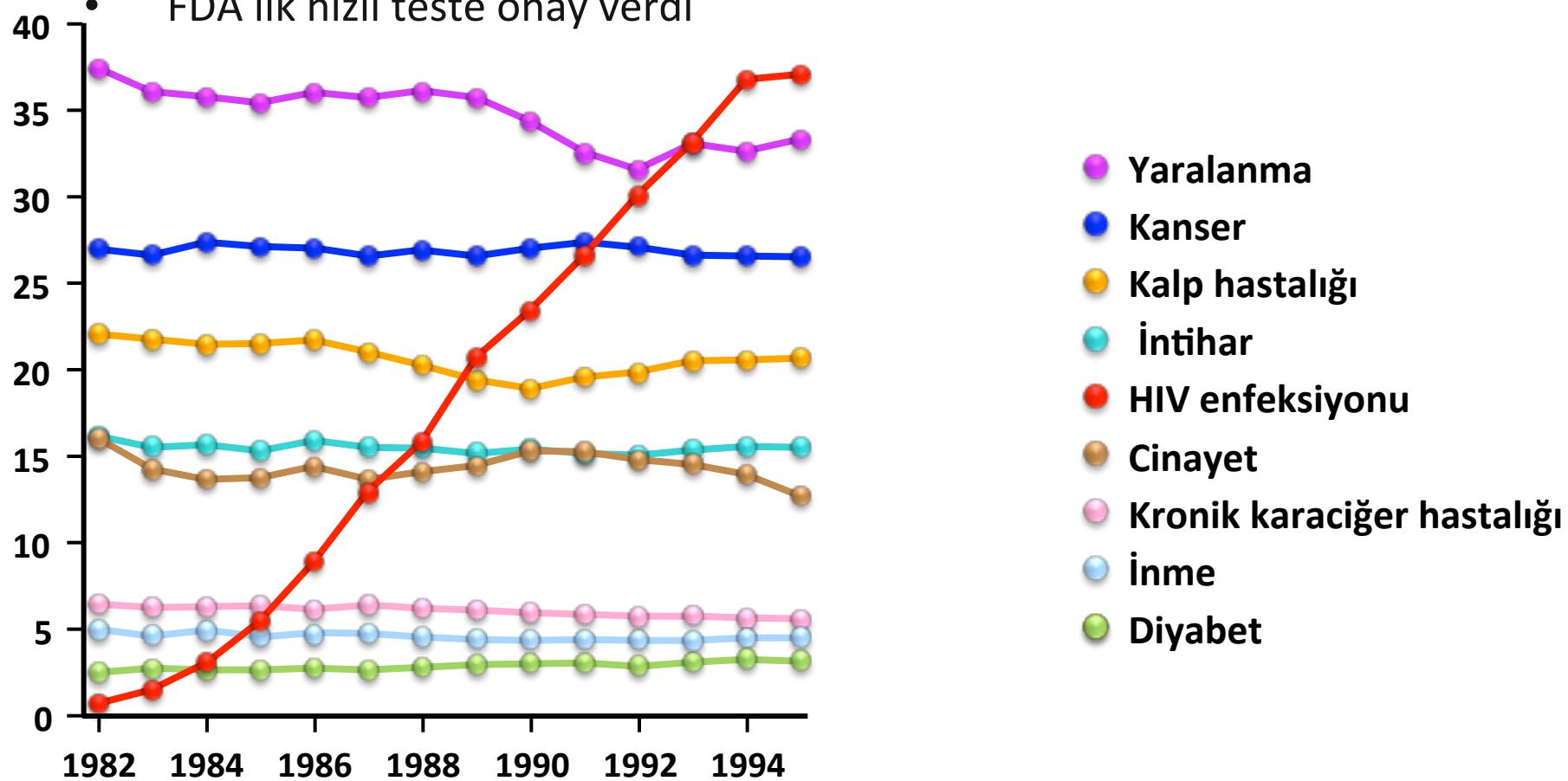
The PHS staff members listed below served as authors of this document. CENTERS FOR DISEASE CONTROL Coordinators Jacquelyn A. Polder, B.S.N., M.P.H. David M. Bell, M.D. Edward Baker, M.D., M.P.H. Kenneth Castro, M.D. Mary Chamberland, M.D., M.P.H. James Curran, M.D., M.P.H. Thomas Folks, Ph.D. Julia Garner, R.N., M.N. James Hughes, M.D. Harold Jaffe, M.D. William Jarvis, M.D. Ruthanne Marcus, M.P.H. William Martone, M.D., M.Sc. Robert Mullin, M.D. Gerald Schochetman, Ph.D. NATIONAL INSTITUTES OF HEALTH David Henderson, M.D. Deborah Katz, R.N., M.S. Charles Litterst, Ph.D. John McGowan, Ph.D. Linda ReckJack Whitescarver, Ph.D. FOOD AND DRUG ADMINISTRATION Janet Arrowsmith, M.D. Paul Beninger, M.D. HEALTH RESOURCES AND SERVICES ADMINISTRATION Samuel Matheny, M.D. INTRODUCTION



[www.aids.gov](http://www.aids.gov)

# 1992

- ABD'de AIDS 25-44 yaş arasındaki **erkeklerde** bir numaralı ölüm nedeni oldu
- FDA ilk hızlı teste onay verdi



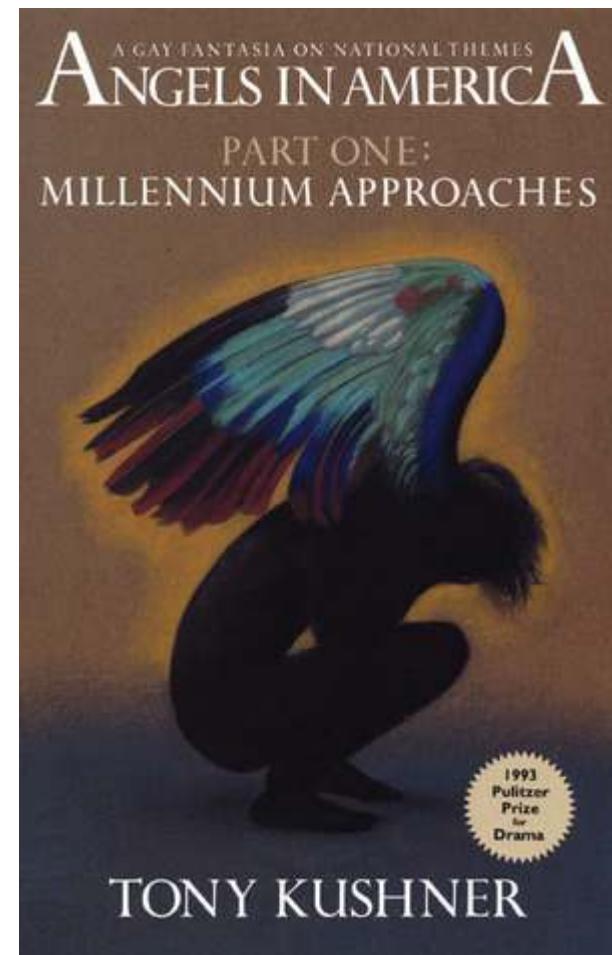


1993





# 1993



AIDS konulu ilk film  
Philadelphia

[www.aids.gov](http://www.aids.gov)

# 1994

AIDS ABD'de yaşayan 25-44 yaş arasındaki **tüm bireylerde** ölümün onde gelen nedeni oldu

Perinatal bulaşmanın önlenmesi için AZT kullanımı önerildi

İlk tükürük testi FDA tarafından onaylandı

The screenshot shows the CDC website header with the CDC logo, a search bar, and links for 'CDC Home' and 'Health Topics A-Z'. Below the header is the 'MMWR' logo and the text 'Recommendations and Reports'. Underneath that, it says 'May 20, 1994 | 43(RR-8);1-17'. A red banner at the bottom left states: 'The content, links, and pdfs are no longer maintained and might be outdated.' Below this, two bullet points provide information about the page's purpose and where to find current information.

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## Guidelines for Preventing Transmission of Human Immunodeficiency Virus Through Transplantation of Human Tissue and Organs

The following CDC staff members prepared this report:

Morris F. Rogers, M.D.; Robert J. Simonds, M.D.; Key R. Lantam, R.N.; M.N.; Robin R. Mowley, M.A.T. National Center for Infectious Diseases

Wanda K. Jones, Dr.P.H. Office of the Associate Director for HIV/AIDS

# 1995



İlk proteaz inhibitörü FDA tarafından onaylandı

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## USPHS/IDSA Guidelines for the Prevention of Opportunistic Infections in Persons Infected with Human Immunodeficiency Virus: A Summary

The following CDC staff member prepared this report:

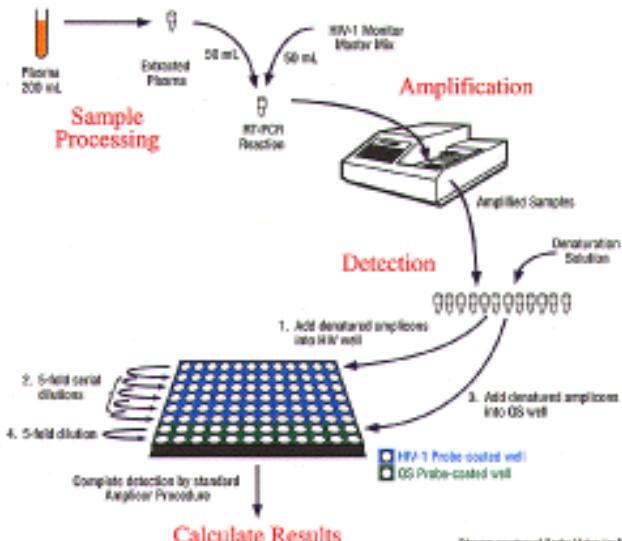
Jonathan E. Kaplan, M.D. National Center for Infectious Diseases National Center for HIV/STD/TB Prevention

# 1996



[www.aids.gov](http://www.aids.gov)  
[www.unaids.org](http://www.unaids.org)

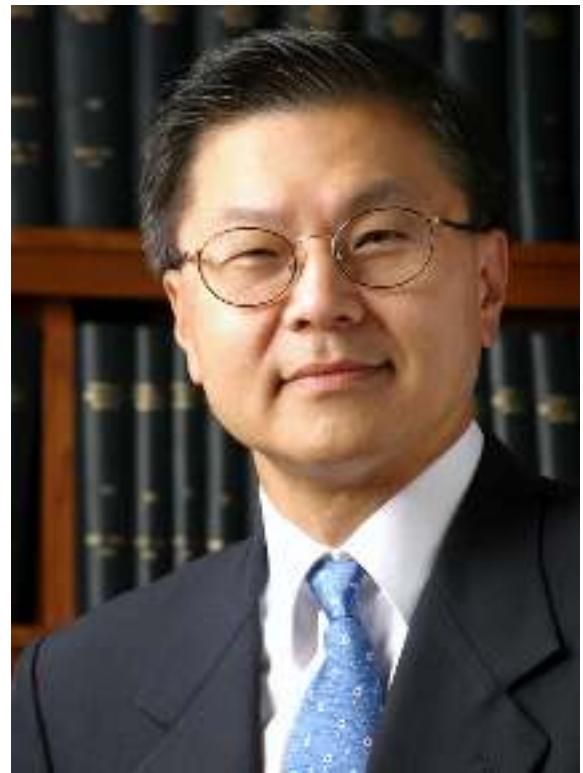
#### How Viral Load Is Measured by HIV RNA Assay



# 1996



## Karma ART



Dr. David Ho

Hit early, hit hard

# 1997

Karma ART standart tedavi yaklaşımı oldu



İlk karma preparat (Combivir) FDA onayı aldı

CDC ABD'de ölümlerin azaldığını bildirdi

Proteaz inhibitörlerine karşı direnç  
gelişmeye başladı

**Dünya üzerinde HIV ile yaşayan 30 milyon  
insan olduğu tahmin ediliyor**

[www.aids.gov](http://www.aids.gov)

[www.unaids.org](http://www.unaids.org)

**1998**



**Guidelines for the Use of Antiretroviral Agents in  
HIV-1-Infected Adults and Adolescents**

[aidsinfo.nih.gov](http://aidsinfo.nih.gov)

# 1999



HIV dünyada ölüm nedenleri arasında  
dördüncü, Afrika'da birinci sırada

Vaxgen aşısı çalışması Tayland'da başlatıldı

# 2000



Develop a global  
partnership for  
development



Eradicate extreme  
poverty and hunger



Achieve universal  
primary education



Ensure environmental  
sustainability

# 2015

## MILLENIUM DEVELOPMENT GOALS



Promote gender  
equality and  
empower women



Combat HIV/Aids,  
malaria and other  
diseases



Improve maternal  
health



Reduce child mortality

# Declaration of Commitment on HIV/AIDS

UNITED NATIONS GENERAL ASSEMBLY  
SPECIAL SESSION ON HIV/AIDS  
25 - 27 JUNE 2001



# 2001

1. İnsanların enfeksiyondan korunmak için ne yapmaları gerektiğini bilmelerini sağlamak
2. Anneden bebeğe bulaşı engellemek
3. Enfekte olan herkesin tedaviye ulaşmasını sağlamak
4. Hastalığa çare ve aşısı bulmak için iki kat daha fazla çalışmak
5. Yaşamları AIDS nedeniyle sarsılmış tüm insanlara yardım etmek

# 2001

- Jenerik ilaç üreticilerinin az gelişmiş ülkeler için düşük fiyatlı ilaç üretimi
- Diğer firmaların az gelişmiş ülkeler için fiyatları düşürmeleri
- Doha Deklarasyonu— gelişmekte olan ülkelere jenerik ilaç satın alma ve üretme hakkının verilmesi



**2002**



# **The Global Fund**

To Fight AIDS, Tuberculosis and Malaria

Hükümetler, sivil toplum örgütleri, özel sektör ve etkilenen toplumların ortaklığı ile kuruldu

# 2002



Parmak kanında hızlı test kiti FDA onayı aldı

Yan etkiler ve artan ilaç direnci nedeniyle  
«hit early hit hard» stratejisi sorgulanmaya başlandı

# 2003



5 yıllık 15 milyon dolarlık plan



[www.aids.gov](http://www.aids.gov)  
[www.who.int](http://www.who.int)

DSÖ üyesi 192 ülkenin işbirliği ile başlatıldı

# 2003

- Vaxgen Faz III çalışması sonlandı
  - Tayland'da 2546 DİİB
  - Kuzey Amerika ve Avrupa'da 5417 eşcinsel
- Plaseboya göre HIV enfeksiyonunda %3,8 oranında azalma
- Aşının koruyucu hiçbir etkisi yok

<http://www.vaxreport.org/Back-Issues/Pages/VaxGenreleasesresultsofThaiPhaseIIIttrial.aspx>

Daniel J. DeNoon. Unsurprising and Surprising Results of VaxGen's HIV Vaccine Trial: An Expert Interview With Mark Feinberg, MD, PhD. Medscape. Feb 28, 2003.

# 2004

- Hızlı testte tükürük kullanımı FDA tarafından onaylandı
- FDA sabit dozlu kombinasyonların onay sürecini hızlandırdı

**2005**



**The Global Fund**  
To Fight AIDS, Tuberculosis and Malaria



World Health  
Organization



**UNAIDS**

**Tedaviye ulaşan insan sayısı  
2004 sonu itibarıyle**

**700 000**

[www.theglobalfund.org](http://www.theglobalfund.org)  
[www.who.int](http://www.who.int)  
[www.unaids.org](http://www.unaids.org)

# 2006

# HIV/AIDS'in 25. yılı



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## MMWR

Recommendations and Reports

September 22, 2006 / 55(RR14);1-17

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## Revised Recommendations for HIV Testing of Adults, Adolescents, and Pregnant Women in Health-Care Settings

Prepared by

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<sup>2</sup>Division of STD Prevention, National Center for HIV/AIDS, Viral Hepatitis, STD, and TB Prevention (proposed) and University of Washington, Seattle, Washington

<sup>3</sup>Northrup Grumman Information Technology (contractor with CDC)

## HIV-1 Target Cells in Foreskins of African Men With Varying Histories of Sexually Transmitted Infections

Betty A. Donoval,<sup>1</sup> Alan L. Landay, PhD,<sup>2</sup> Stephen Moses, MD, PhD,<sup>3</sup> Kawango Agot, PhD,<sup>4</sup> J.O. Ndinya-Achola, MBchB,<sup>5</sup> Edith A. Nyagaya,<sup>4</sup> Ian MacLean, PhD,<sup>3</sup> and Robert C. Bailey, PhD<sup>1</sup>

**Key Words:** HIV-1; Foreskin; Circumcision; Immunohistochemistry

DOI: 10.1309/JVHQWDYKM58EPH

CYBE olan (s:20) ve olmayan (s: 19) erkeklerin sünnet derileri incelenmiş CYBE öyküsü olanlarda Langerhans hücreleri ve makrofajların sayısı daha yüksek Sünnet derisinin mukozal yüzeyinde bulunan bu hücreler kişiyi HIV'e daha duyarlı kılıyor

# 2007

**HIV/AIDS Programme**  
Strengthening health services to fight HIV/AIDS

**GUIDANCE ON PROVIDER-INITIATED  
HIV TESTING AND COUNSELLING  
IN HEALTH FACILITIES**

 World Health Organization

 UNAIDS  
Joint United Nations Programme on HIV/AIDS

[www.who.int](http://www.who.int)  
[www.unaids.org](http://www.unaids.org)

# 2008



5 yıl daha uzatıldı

[www.aids.gov](http://www.aids.gov)

Son 10 yılda  
yeni enfeksiyonlarda  
%17 azalma

Doğu Asya'da %25 artış



# 2009



# The NEW ENGLAND JOURNAL of MEDICINE

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## Long-Term Control of HIV by CCR5 Delta32/Delta32 Stem-Cell Transplantation

Gero Hütter, M.D., Daniel Nowak, M.D., Maximilian Mossner, B.S., Susanne Ganepola, M.D., Arne Müßig, M.D., Kristina Allers, Ph.D., Thomas Schneider, M.D., Ph.D., Jörg Hofmann, Ph.D., Claudia Kücherer, M.D., Olga Blau, M.D., Igor W. Blau, M.D., Wolf K. Hofmann, M.D., and Eckhard Thiel, M.D.

N Engl J Med 2009; 360:692-698 | February 12, 2009 | DOI: 10.1056/NEJMoa0802905

# 2010



## NIH Public Access Author Manuscript

*Science*. Author manuscript; available in PMC 2011 September 3.

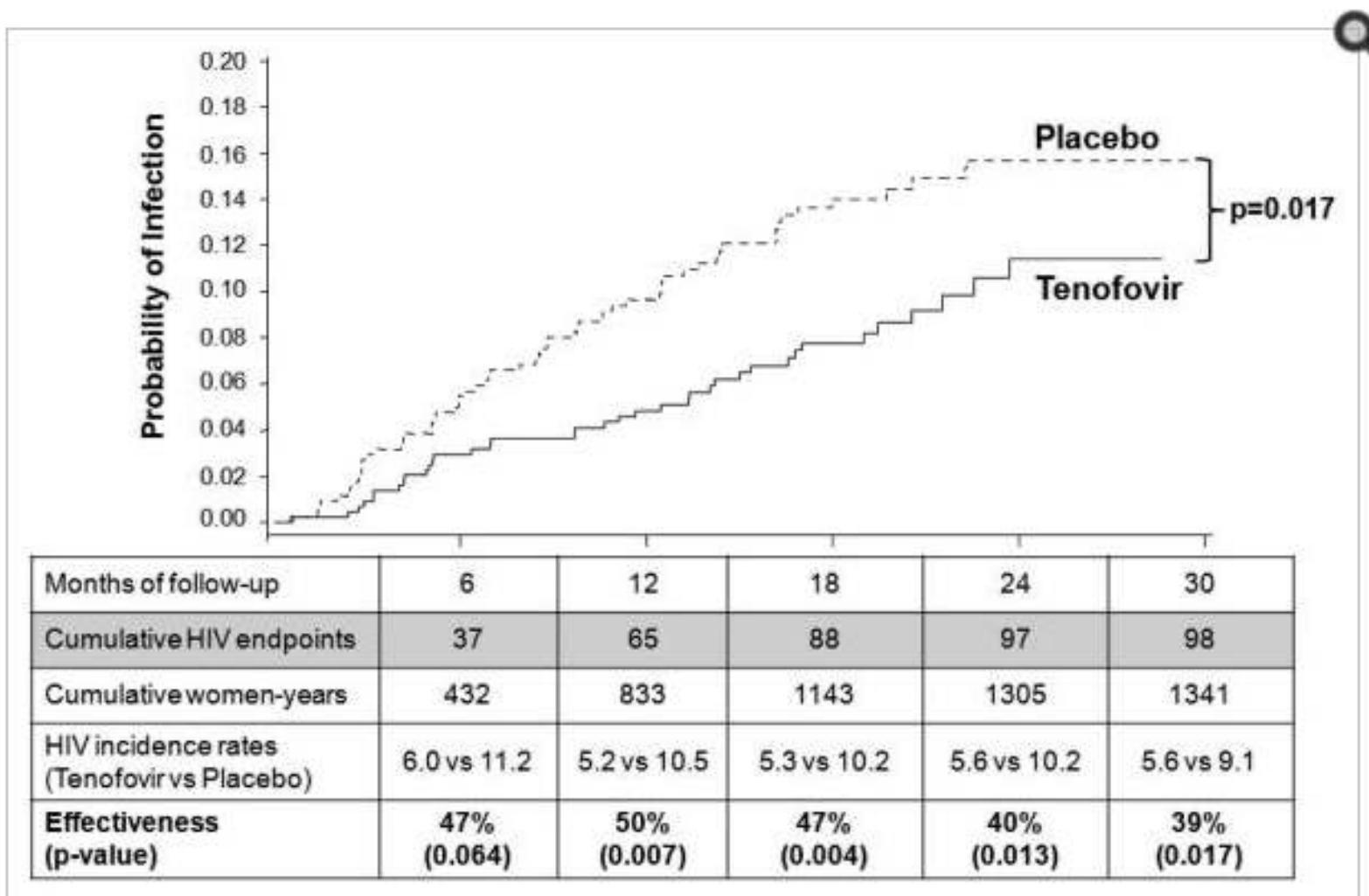
Published in final edited form as:

*Science*. 2010 September 3; 329(5996): 1168–1174. doi:10.1126/science.1193748.

### **Effectiveness and Safety of Tenofovir Gel, an Antiretroviral Microbicide, for the Prevention of HIV Infection in Women**

Quarraisha Abdool Karim<sup>\*1,2</sup>, Salim S. Abdool Karim<sup>\*1,2,3</sup>, Janet A. Frohlich<sup>1</sup>, Anneke C. Grobler<sup>1</sup>, Cheryl Baxter<sup>1</sup>, Leila E. Mansoor<sup>1</sup>, Ayesha B.M. Kharsany<sup>1</sup>, Sengeziwe Sibeko<sup>1</sup>, Koleka P. Mlisana<sup>1</sup>, Zaheen Omar<sup>1</sup>, Tanuja N Gengiah<sup>1</sup>, Silvia Maarschalk<sup>1</sup>, Natasha Arulappan<sup>1</sup>, Mukelisiwe Mlotshwa<sup>1</sup>, Lynn Morris<sup>4</sup>, and Douglas Taylor<sup>5</sup> on behalf of the CAPRISA 004 Trial Group

## HIV'in edinilme riskinde %39 (tam uyumla kullananlarda %54) azalma Figure 2



İnsidans TDF kolunda 5.6/100 kadın yılı, placebo kolunda 9.1/100 kadın yılı

# 2010



## NIH Public Access Author Manuscript

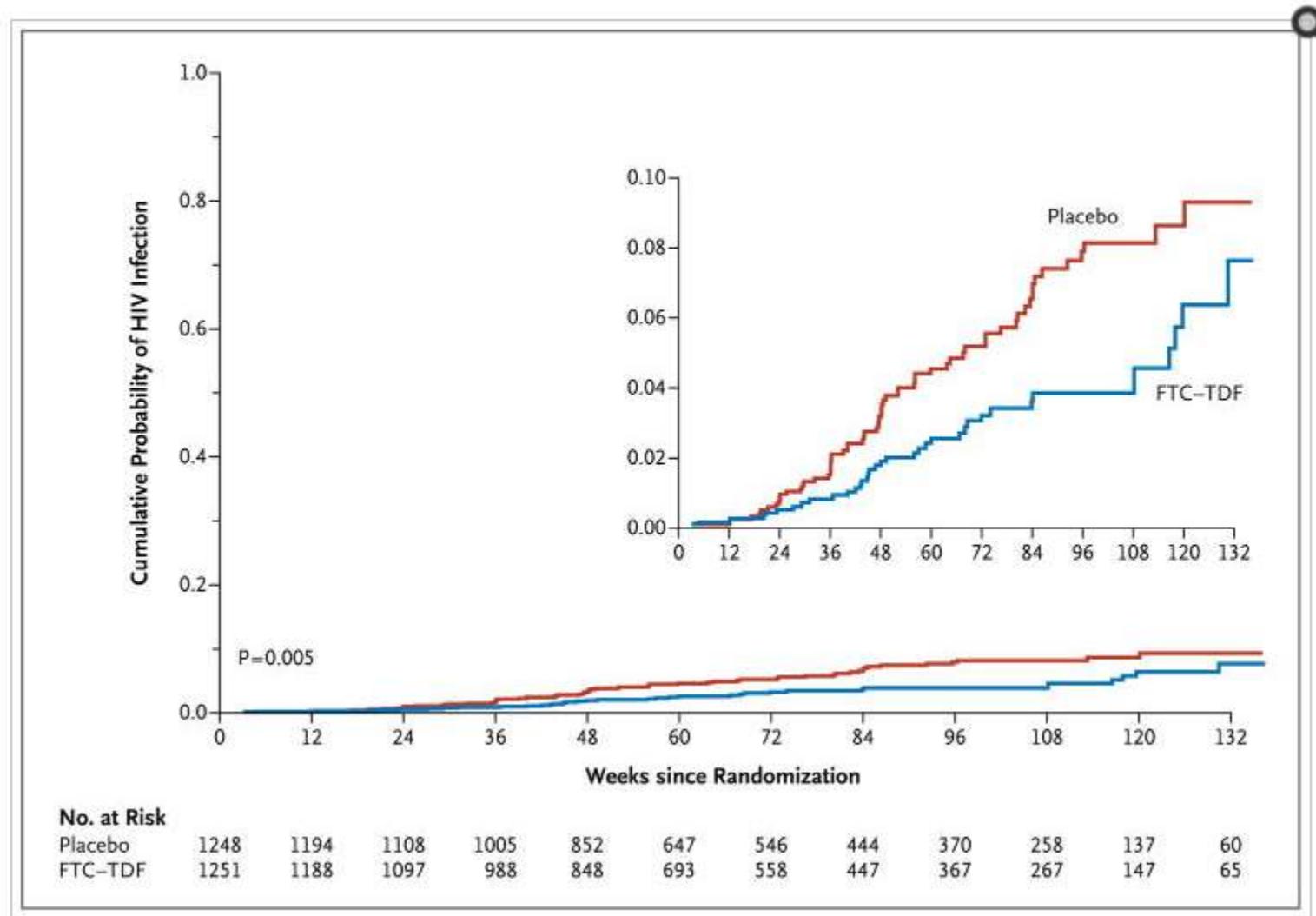
*N Engl J Med.* Author manuscript; available in PMC 2011 June 30.

Published in final edited form as:

*N Engl J Med.* 2010 December 30; 363(27): 2587–2599. doi:10.1056/NEJMoa1011205.

### Preexposure Chemoprophylaxis for HIV Prevention in Men Who Have Sex with Men

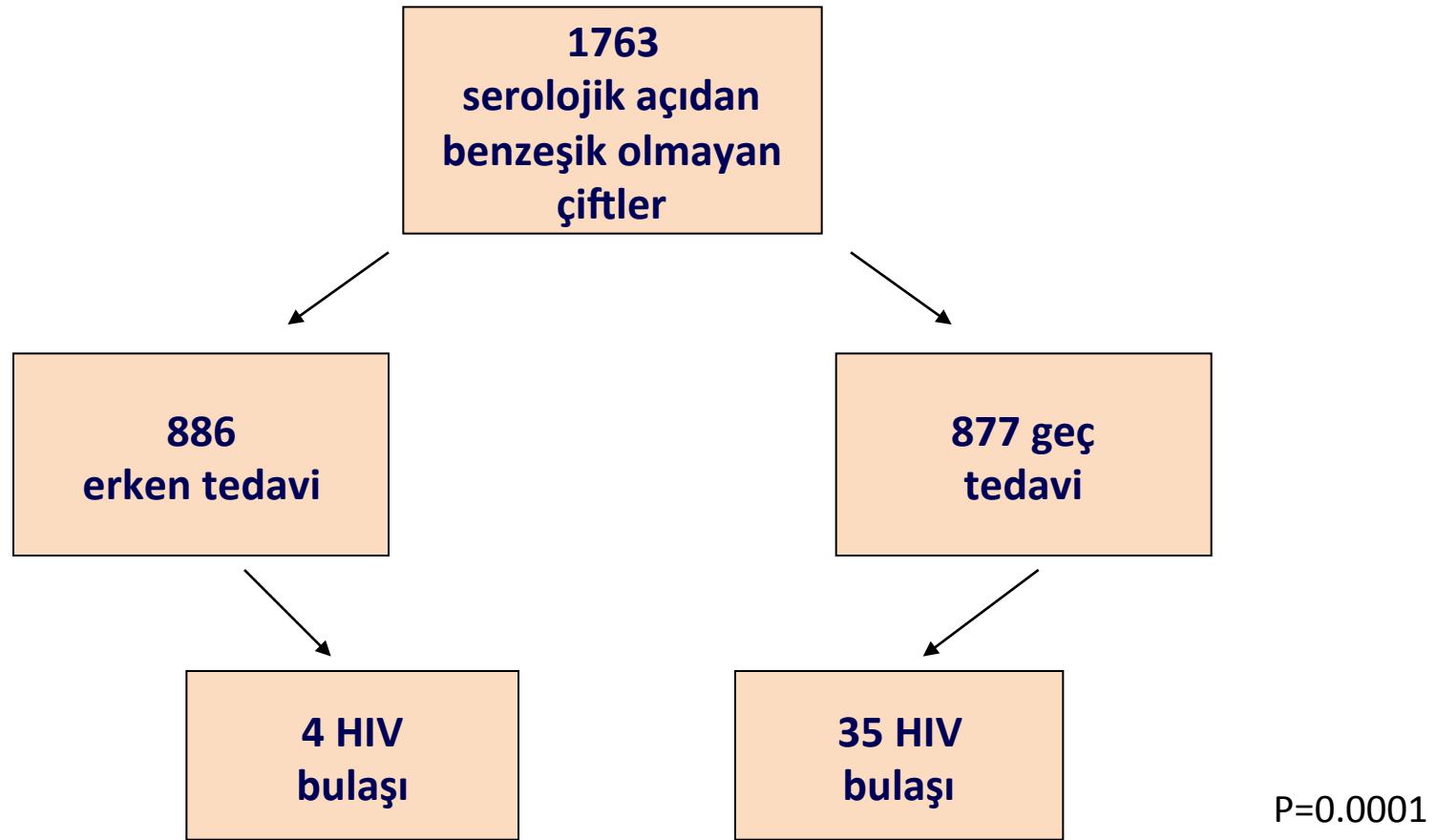
Robert M. Grant, M.D., M.P.H., Javier R. Lama, M.D., M.P.H., Peter L. Anderson, Pharm. D., Vanessa McMahan, B.S., Albert Y. Liu, M.D., M.P.H., Lorena Vargas, Pedro Goicochea, M.Sc., Martín Casapia, M.D., M.P.H., Juan Vicente Guanira-Carranza, M.D., M.P.H., María E. Ramírez-Cardich, M.D., Orlando Montoya-Herrera, M.Sc., Telmo Fernández, M.D., Valdilea G. Veloso, M.D., Ph.D., Susan P. Buchbinder, M.D., Suwat Chariyalertsak, M.D., Dr. P.H., Mauro Schechter, M.D., Ph.D., Linda-Gail Bekker, M.B., Ch.B., Ph.D., Kenneth H. Mayer, M.D., Esper Georges Kallás, M.D., Ph.D., K. Rivet Amico, Ph.D., Kathleen Mulligan, Ph.D., Lane R. Bushman, B. Chem., Robert J. Hance, A.A., Carmela Ganoza, M.D., Patricia Defechereux, Ph.D., Brian Postle, B.S., Furong Wang, M.D., J. Jeff McConnell, M.A., Jia-Hua Zheng, Ph.D., Jeanny Lee, B.S., James F. Rooney, M.D., Howard S. Jaffe, M.D., Ana I. Martinez, R. Ph., David N. Burns, M.D., M.P.H., and David V. Glidden, Ph.D. for the iPrEx Study Team\*

**Figure 2****Kaplan-Meier Estimates of Time to HIV Infection (Modified Intention-to-Treat Population)**

TDF-FTC alan grupta HIV enfeksiyonunda %44 azalma

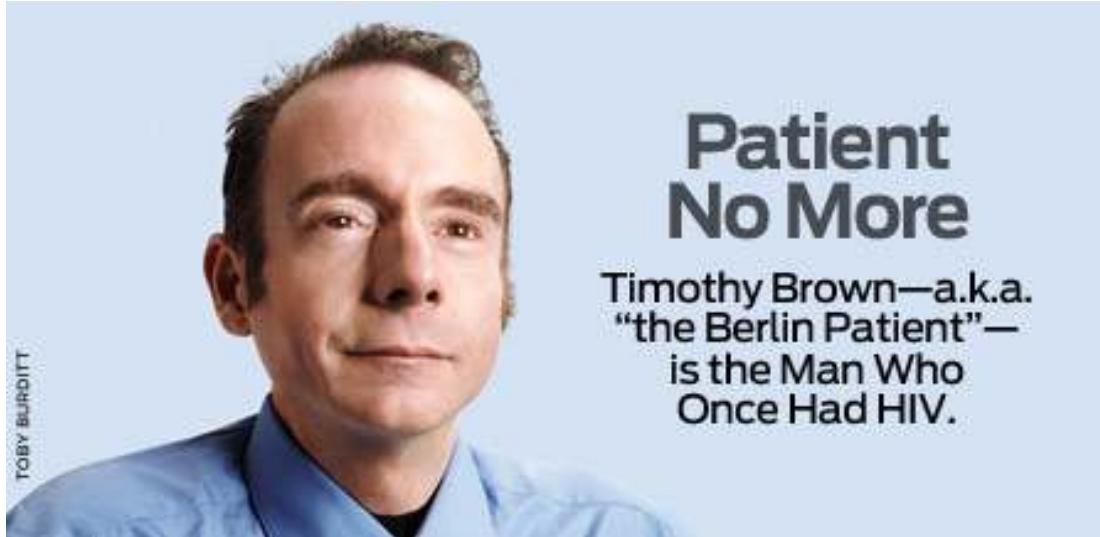
# 2011

## HPTN 052 çalışması



# (HPTN 052)

	Erken (s=886)	Geç (s=877)	
Başlangıçtaki CD4 ortanca (ÇDG)	442 (373–522)	428 (357–522)	
Başlangıçtaki viral yük ortanca ÇDG	4.4 (3.8–4.9)	4.4 (3.9–4.9)	
Yaş (indeks eş)	33	32	
Evli	%94	%95	
Korunmasız ilişki	%6	%8	
Bağlantılı geçişler (s)	1	27	HR 0.04 (%95 GA: 0.01–0.28)
DSÖ Evre IV olaylar, pulmoner TB, ağır bakteriyel enf. veya ölüm (s=hasta)	40 (her HY için 2.4)	65 (her HY için 4.0)	HR 0.59, %95 GA: (0.40, 0.88), p=0.01
TB (s=olay)	17	33	
Ekstrapulmoner TB (s)	3	17	p<0.002
Ölüm	10	13	HR 0.77, %95 GA: (0.34, 1.76), p>0.5
İstenmeyen olay	%24	%5	



Toby Burritt

# Patient No More

Timothy Brown—a.k.a.  
“the Berlin Patient”—  
is the Man Who  
Once Had HIV.

2011

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## CLINICAL TRIALS AND OBSERVATIONS

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### Evidence for the cure of HIV infection by CCR5Δ32/Δ32 stem cell transplantation

Kristina Allers,<sup>1</sup> Gero Hütter,<sup>2</sup> Jörg Hofmann,<sup>3</sup> Christoph Loddenkemper,<sup>4</sup> Kathrin Rieger,<sup>2</sup> Eckhard Thiel,<sup>2</sup> and Thomas Schneider<sup>1</sup>

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# 2012



## NIH Public Access Author Manuscript

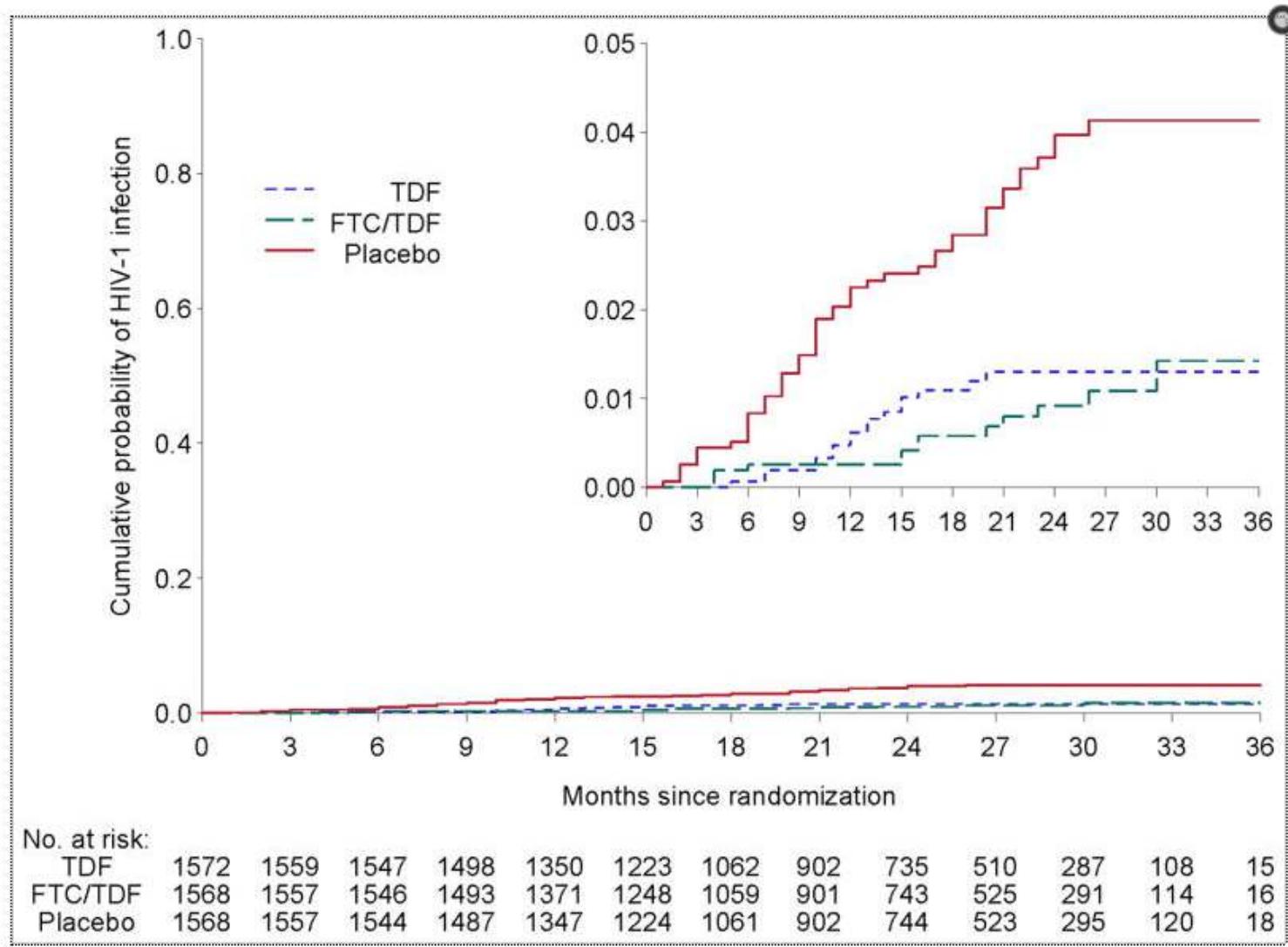
*N Engl J Med* Author manuscript; available in PMC 2013 September 11.

Published in final edited form as:

*N Engl J Med.* 2012 August 2; 367(5): 399–410. doi:10.1056/NEJMoa1108524.

### Antiretroviral Prophylaxis for HIV-1 Prevention among Heterosexual Men and Women

Jared M. Baeten, M.D., Ph.D., Deborah Donnell, Ph.D., Patrick Ndase, M.B., Ch.B., M.P.H., Nelly R. Mugo, M.B., Ch.B., M.P.H., James D. Campbell, M.D., M.S., Jonathan Wangisi, M.B., Ch.B., M.Sc., Jordan W. Tappero, M.D., M.P.H., Elizabeth A. Bukusi, M.B., Ch.B., Ph.D., Craig R. Cohen, M.D., M.P.H., Elly Katabira, M.B., Ch.B., Allan Ronald, M.D., Elioda Tumwesigye, M.B., Ch.B., M.S., Edwin Were, M.B., Ch.B., M.P.H., Kenneth H. Fife, M.D., Ph.D., James Kiarie, M.B., Ch.B., M.P.H., Carey Farquhar, M.D., M.P.H., Grace John-Stewart, M.D., Ph.D., Aloysious Kakia, M.B., Ch.B., Josephine Odoyo, M.P.H., Akasiima Mucunguzi, M.B., Ch.B., Edith Nakku-Joloba, M.B., Ch.B., Ph.D., Rogers Tswesigye, M.B., Ch.B., M.P.H., Kenneth Ngure, M.P.H., Cosmas Apaka, B.Sc., Harrison Tamoooh, M.B., Ch.B., Fridah Gabona, M.B., Ch.B., Andrew Mujugira, M.B., Ch.B., M.Sc., Dana Panteleeff, B.S., Katherine K. Thomas, M.S., Lara Kidoguchi, M.P.H., Meighan Krows, B.A., Jennifer Revall, B.A., Susan Morrison, M.D., M.P.H., Harald Haugen, M.S., Mira Emmanuel-Ogier, B.A., Lisa Ondrejcek, M.A., Robert W. Coombs, M.D., Ph.D., Lisa Frenkel, M.D., Craig Hendrix, M.D., Namandjé N. Bumpus, Ph.D., David Bangsberg, M.D., M.P.H., Jessica E. Haberer, M.D., M.P.H., Wendy S. Stevens, M.D., F.C.Path., Jairam R. Lingappa, M.D., Ph.D., and Connie Celum, M.D., M.P.H. for the Partners PrEP Study Team\*



Kaplan-Meier curve for the primary modified intention-to-treat analysis

Plaseboya göre HIV-1 insidansında TDF ile %67, TDF-FTC ile %75 azalma; fark anlamlı değil

# 2012

## Key Updates to Existing Sections

Following are key updates to existing sections of the guidelines.

### ***Initiating Antiretroviral Therapy in Treatment-Naive Patients***

The Panel updated its recommendations on initiation of ART in treatment-naive patients. The changes are primarily based on increasing evidence showing the harmful impact of ongoing HIV replication on AIDS and non-AIDS disease progression. In addition, the updated recommendations reflect emerging data showing the benefit of effective ART in preventing secondary transmission of HIV. The updated section includes more in-depth discussion on the rationale for these recommendations and on the risks and benefits of long-term ART.

The Panel's recommendations are listed below.

- ART is recommended for all HIV-infected individuals. The strength of this recommendation<sup>a</sup> varies on the basis of pretreatment CD4 cell count:
  - CD4 count <350 cells/mm<sup>3</sup> (**AI**)
  - CD4 count 350 to 500 cells/mm<sup>3</sup> (**AII**)
  - CD4 count >500 cells/mm<sup>3</sup> (**BIII**)
- Regardless of CD4 count, initiation of ART is strongly recommended for individuals with the following conditions:
  - Pregnancy (**AI**) (see [perinatal guidelines](#) for more detailed discussion)
  - History of an AIDS-defining illness (**AI**)
  - HIV-associated nephropathy (HIVAN) (**AII**)
  - HIV/hepatitis B virus (HBV) coinfection (**AII**)

# 2012

TDF+FTC temas öncesi korunmada  
kullanılmak üzere FDA onayı aldı



# 2013

Dünyada 35 milyon kişinin HIV ile yaşadığı  
tahmin ediliyor

2001-2013 arasında yeni HIV  
enfeksiyonları %38 azaldı

## 2020'ye dek

90%

of all



living with HIV will  
know their HIV  
status

90%

of all



living with HIV will  
receive sustained  
antiretroviral  
therapy

90%

of all



receiving  
antiretroviral therapy  
will have durable viral  
suppression

2015

# The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

AUGUST 27, 2015

VOL. 373 NO. 9

## Initiation of Antiretroviral Therapy in Early Asymptomatic HIV Infection

The INSIGHT START Study Group\*

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### ABSTRACT

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#### BACKGROUND

Data from randomized trials are lacking on the benefits and risks of initiating antiretroviral therapy in patients with asymptomatic human immunodeficiency virus (HIV) infection who have a CD4+ count of more than 350 cells per cubic millimeter.

#### METHODS

We randomly assigned HIV-positive adults who had a CD4+ count of more than 500 cells per cubic millimeter to start antiretroviral therapy immediately (immediate-initiation group) or to defer it until the CD4+ count decreased to 350 cells per cubic millimeter or until the development of the acquired immunodeficiency syndrome (AIDS) or another condition that dictated the use of antiretroviral therapy (deferred-initiation group). The primary composite end point was any serious AIDS-related event, serious non-AIDS-related event, or death from any cause.

#### RESULTS

A total of 4685 patients were followed for a mean of 3.0 years. At study entry, the median HIV viral load was 12,759 copies per milliliter, and the median CD4+ count was 651 cells per cubic millimeter. On May 15, 2015, on the basis of an interim analysis, the data and safety monitoring board determined that the study question had been answered and recommended that patients in the deferred-initi-

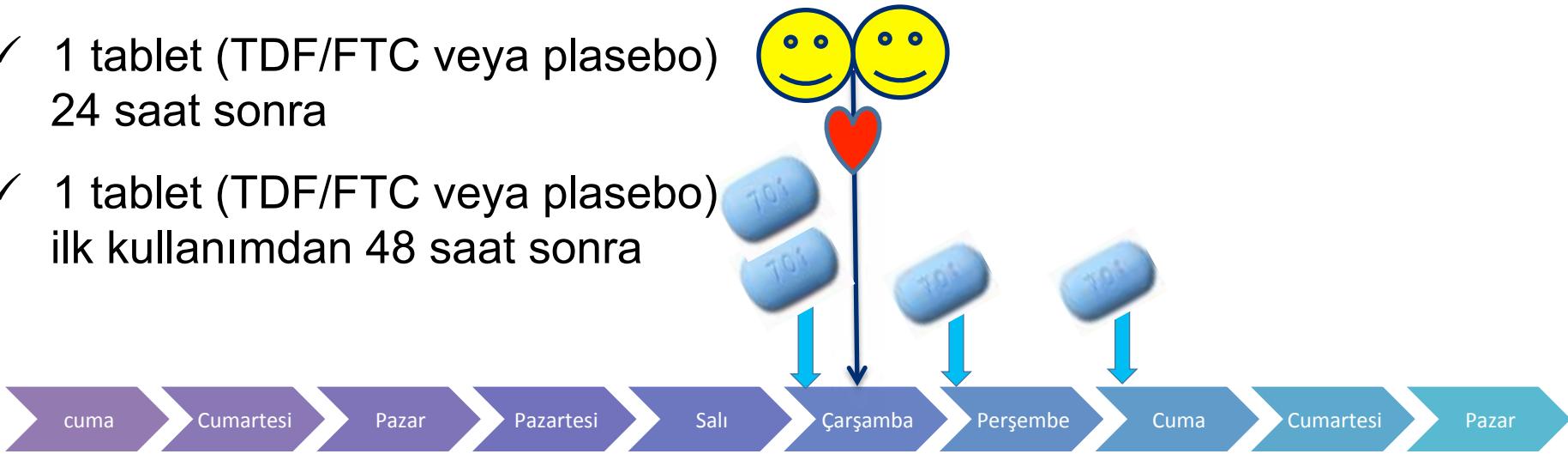
The members of the writing group (Jens D. Lundgren, M.D. [cochair], Abdel G. Babiker, Ph.D. [cochair], Fred Gordin, M.D. [cochair], Sean Emery, Ph.D., Birgit Grund, Ph.D., Shweta Sharma, M.S., Anchalee Avihingsanon, M.D., David A. Cooper, M.D., Gerd Fätkenheuer, M.D., Josep M. Llibre, M.D., Jean-Michel Molina, M.D., Paula Munderi, M.D., Mauro Schechter, M.D., Robin Wood, M.D., Karin L. Klingman, M.D., Simon Collins, H. Clifford Lane, M.D., Andrew N. Phillips, Ph.D., and James D. Neaton, Ph.D. [INSIGHT PI]) of the INSIGHT START Study Group assume responsibility for the overall content and integrity of this article. The affiliations of the members of the writing group are listed in the Appendix. Address reprint requests to Dr. Lundgren at the Department of Infectious Diseases, Rigshospitalet, University of Copenhagen, Blegdamsvej 9, 2100 Copenhagen Ø, Denmark, or at jens.lundgren@regionh.dk.

End Point	Immediate-Initiation Group (N = 2326)		Deferred-Initiation Group (N = 2359)		Hazard Ratio (95% CI)†	P Value
	no.	no./100 person-yr	no.	no./100 person-yr		
Composite primary end point	42	0.60	96	1.38	0.43 (0.30–0.62)	<0.001
Components of the primary end point						
Serious AIDS-related event	14	0.20	50	0.72	0.28 (0.15–0.50)	<0.001
Serious non-AIDS-related event	29	0.42	47	0.67	0.61 (0.38–0.97)	0.04
Death from any cause	12	0.17	21	0.30	0.58 (0.28–1.17)	0.13
Tuberculosis	6	0.09	20	0.28	0.29 (0.12–0.73)	0.008
Kaposi's sarcoma	1	0.01	11	0.16	0.09 (0.01–0.71)	0.02
Malignant lymphoma	3	0.04	10	0.14	0.30 (0.08–1.10)	0.07
Cancer not related to AIDS	9	0.13	18	0.26	0.50 (0.22–1.11)	0.09
Cardiovascular disease	12	0.17	14	0.20	0.84 (0.39–1.81)	0.65
Other secondary end points						
Grade 4 event‡	73	1.06	73	1.05	1.01 (0.73–1.39)	0.97
Unscheduled hospitalization§	262	4.02	287	4.40	0.91 (0.77–1.08)	0.28
Grade 4 event, unscheduled hospitalization, or death from any cause	283	4.36	311	4.78	0.91 (0.77–1.07)	0.25
Most common grade 4 events, unscheduled hospitalization, or death from any cause¶						
Bacterial infectious disorder	14	0.20	36	0.52	0.38 (0.20–0.70)	0.002
Bone or joint injury	17	0.24	11	0.16	1.55 (0.73–3.31)	0.26
Depressed mood disorder or disturbance	12	0.17	9	0.13	1.34 (0.57–3.19)	0.50
Infection with unspecified pathogen	64	0.93	65	0.94	0.99 (0.70–1.40)	0.96
Injury not elsewhere classified	11	0.16	22	0.31	0.50 (0.24–1.03)	0.06
Suicidal or self-injurious behavior not elsewhere classified	27	0.39	24	0.34	1.15 (0.66–1.99)	0.63
Viral infectious disorder	12	0.17	15	0.21	0.81 (0.38–1.72)	0.58
Grade 4 event, unscheduled hospitalization, or primary end point	295	4.56	355	5.52	0.82 (0.71–0.96)	0.01

# 2015

# IPERGAY Cinsel eylemle bağlantılı TÖP

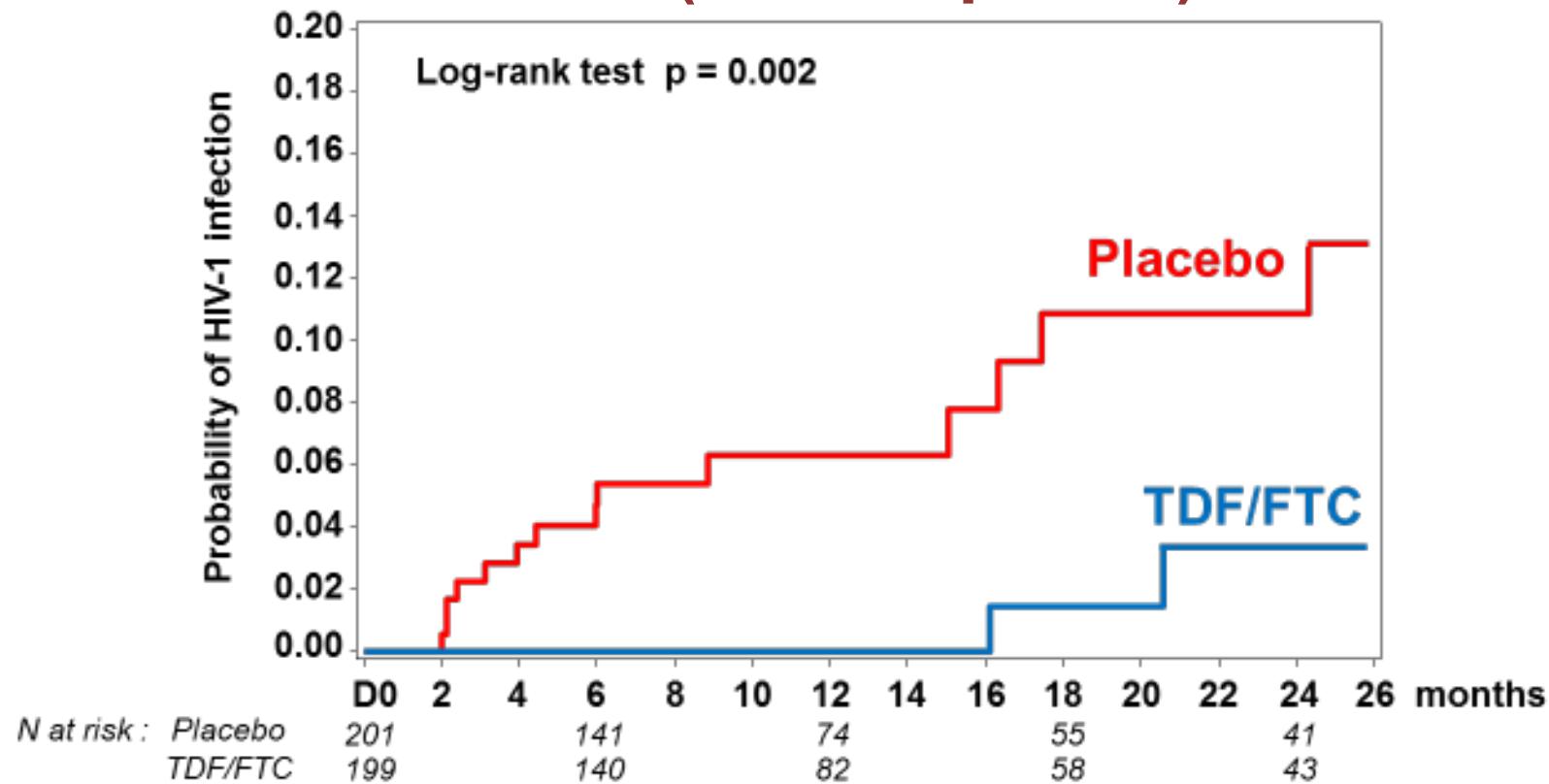
- ✓ 2 tablet (TDF/FTC veya placebo) seksten 2-24 saat önce
- ✓ 1 tablet (TDF/FTC veya placebo) 24 saat sonra
- ✓ 1 tablet (TDF/FTC veya placebo) ilk kullanımından 48 saat sonra



Tek cinsel ilişkiyi kapsamak için 3 gün içinde 4 tablet TDF/FTC



# HIV-1 enfeksiyonuna kadar geçen zaman için KM kestirimleri (mITT toplumu)



Ortalama izlem süresi 13 ay: 16 katılımcıda yeni enfeksiyon

**14 placebo kolunda** (insidans: 6,6 / 100 KY), **2 TDF/FTC kolunda** (insidans: 0,94 / 100 KY)  
Direnç mutasyonu saptanmamış

**HIV-1insidansında %86 görece azalma (%95 GA: 40-99, p=0,002)**

Bir enfeksiyonu önlemek için bir yıl boyunca tedavi edilmesi gereken kişi sayısı: **18**



Public Health  
England

2016



# PRe-exposure Option for HIV prevention in the UK: immediate or Deferred

<http://www.proud.mrc.ac.uk/>

# HIV İnsidansı

<b>Grup</b>	<b>Enf. sayısı</b>	<b>İzlem (KY)</b>	<b>İnsidans (her 100 KY)</b>	<b>%90 GA</b>
Tüm grup	22	453	4,9	3,4-6,8
Hemen	3 (+2*)	239	1,3	0,4-3,0
Ertelenen	19 (+1**)	214	8,9	6,0-12,7

**Etkinlik** =%86 (%90 GA: %58 – 96)

**P değeri** =0,0002

**Oran farkı** =7,6 (%90 GA: 4,1 – 11,2)

**Profilaksi alması gerekenler** =13 (%90 GA: 9–25)

\* 2 olgu ilk taramada pozitif bulunmuş

\*\* 1 olgu ilk taramada pozitif bulunmuş

# Then Now Future

Fifteen years of progress and hope. But miles to go to end the AIDS epidemic by 2030—new milestones to reach, barriers to break and frontiers to cross.

## People living with HIV on antiretroviral therapy

**1** million  
**15** million  
**All** people living with HIV

2001

2015

2030

## New HIV infections

**3** million  
**2** million  
**0.2** million

2001

2014

2030

## AIDS-related deaths

**2.0** million  
**1.2** million  
**0.2** million

2004

2014

2030

## Investments for AIDS response

**4.9** US\$ billion  
**21.7** US\$ billion  
**32** US\$ billion

2001

2015

2020