

A Public Health Blueprint for Healthy Aging

Red Ribbons, Silver Threads:
Healthy Aging in the Era of HIV/AIDS

Linda P. Fried, M.D., M.P.H.

Longevity with HIV/AIDS:
Adding successful aging to clinical
and public health planning

Persons 50 and older living with HIV in New York State 2007

Male (50+)	30,716
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Female (50+)	11,462
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Total	42,178
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We are an aging society

- U.S. by 2030: 23% >65;
 - As many older adults as children
- Best educated, healthiest group of older adults in history
- Health status in last third of life critically important: to individual, to families, to cities, to society

Public Health Goals for an Aging Society

Life expectancy: 79 years

*Will people be living longer years
healthy or ill?*

Chronic Diseases Become the Norm

Adults 65 and older:
80% 1 or more diseases
50% 2 or more diseases
40% with disability in walking

“Compression of Morbidity”

- Improving healthy later years through delaying onset of morbidity and disability to latest points in the lives of older adults.
- *Public health and clinical goal for improving the health of an aging society.*

THE COMPRESSION OF MORBIDITY

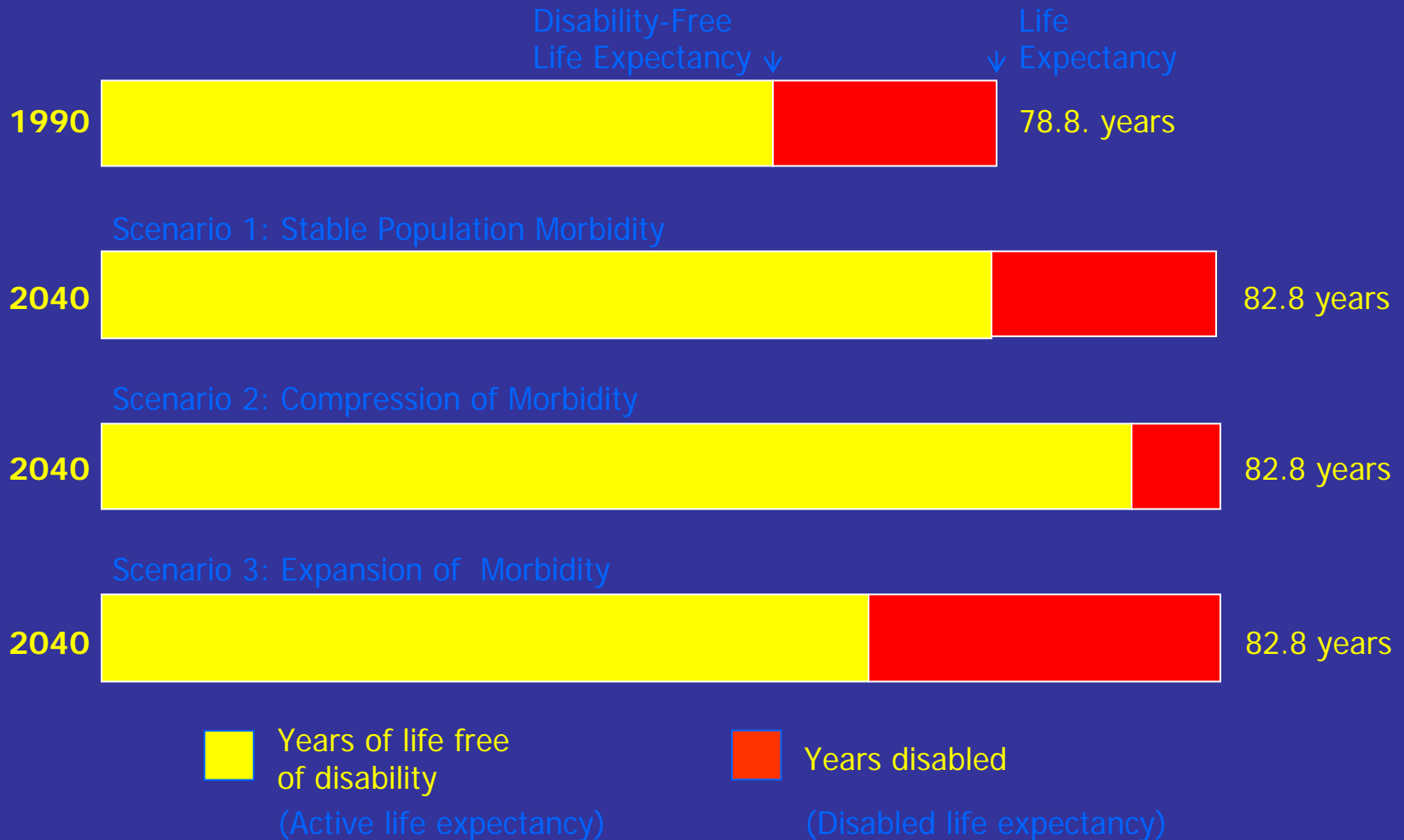


PROTOTYPIC LINGERING CHRONIC ILLNESS



EFFECTS OF THE POSTPONEMENT OF CHRONIC DISEASE

Scenarios for Change in Population Burden of Disability from 1990 to 2040



Changing realities

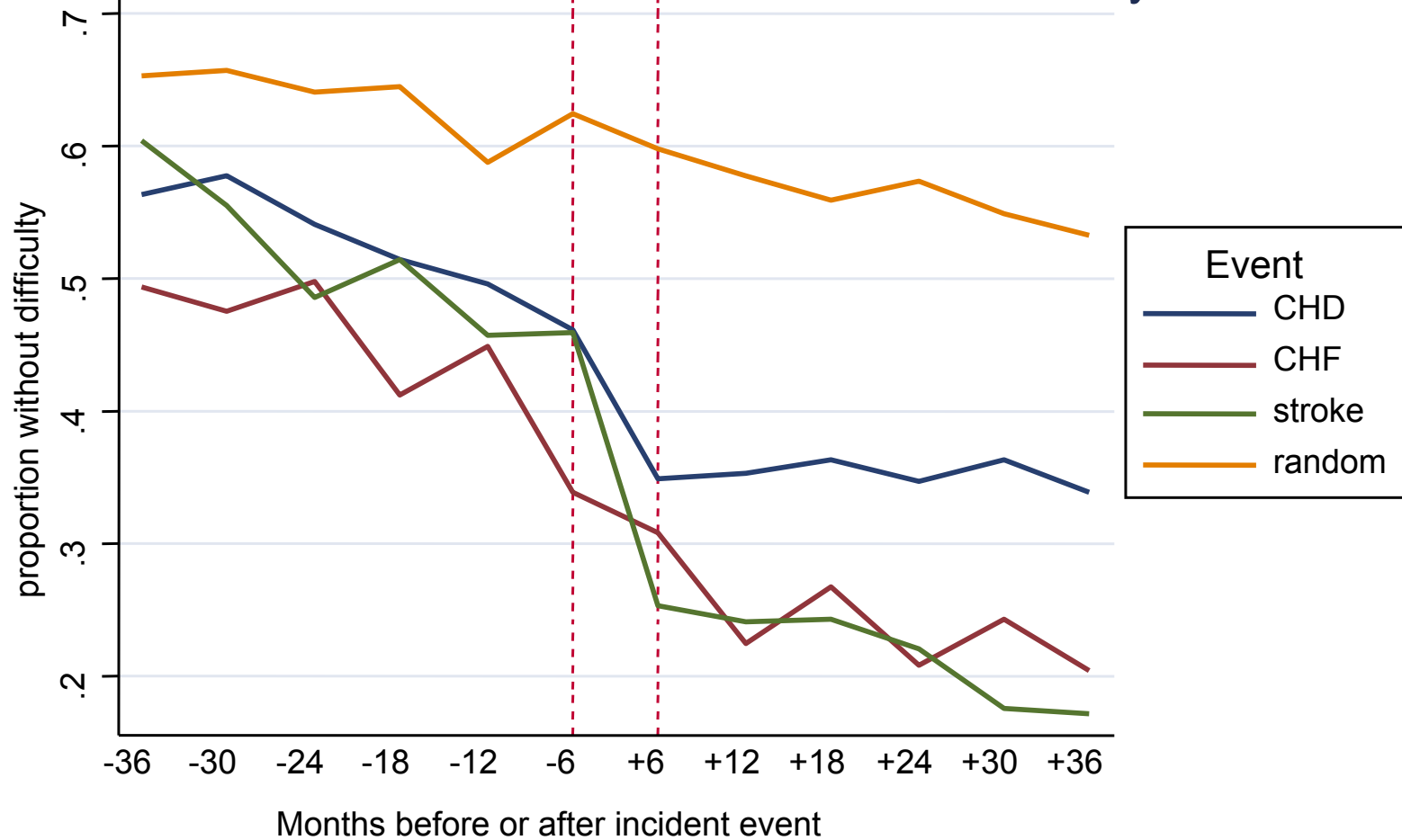
Age- and Disease-Associated Outcomes Emerge, 65+

- Chronic diseases (>80%)
- Multimorbidity (>50%)
- Frailty (7-10%)
- Falls (31%)
- Disability (40%)

Specific Diseases Cause Disability

- Heart disease
- Knee OA
- Hip fracture
- Diabetes
- Stroke
- Claudication
- COPD
- Visual Impairment
- Depression
- Cognitive Impairment

Onset of physical disability in high energy mobility tasks, men and women 65-101 in Cardiovascular Health Study



Health Disparities Persist

- Disability in Inner City African Americans 49-65 years:
 - Occurs 10 years earlier than in suburban AAs and whites
 - Prevalence: 60% with ≥ 1 disability (women > men)
 - Mediated by race and SES

Miller D et al, 2005

WHAT we DO Matters with aging: Improving Health Outcomes

- Physical Activity
- Cognitive Activity
- Social engagement:
 - Social networks and support; loneliness
 - Structured activities
 - Meaningful, productive roles
 - Leaving a legacy

Historic opportunity: Shifting life stages

- *In developed countries, people will be living $\geq 1/3$ of their lives after retirement*
- *Greater longevity has created a “third age” and a “fourth age” of aging, with new opportunities and needs*

Psychological Development Continues over the Life Course

(Erickson; Cohen G; others)

“generativity” is key to successful aging

- Leaving a legacy; Leaving the world better for future generations (Erikson)
 - Productive, meaningful engagement

Successful Aging is Multidimensional

- Avoidance of disease and disability
- Maintenance of high physical function
- Sustained engagement in social and productive activities

*Rowe and Kahn, 1998, from
MacArthur Network on Successful Aging*

Colin A Depp and Dilip V Jeste

Definitions and Predictors of Successful Aging

A Comprehensive Review of Larger Quantitative Studies

Predictors of Successful Aging

- Higher Income/ educations
 - Male sex (controversial)
 - White ethnicity
 - Current marriage/ large social network
 - Better cognitive performance and memory
 - **Better physical performance/ Muscle strength**
 - **No smoking/ moderate alcohol consumption**
 - **High physical activity**
 - Better self-rated health
 - Biomedical markers
 - High HDL cholesterol
 - **No overt obesity**
 - **No hypertension/diabetes/CHD/cancer/OA/stroke**
 - **No hearing problems**
 - **No depression/ Positive attitude toward aging**
 - High life satisfaction/ self-efficacy/ mastery
 - **Positive attitude toward Aging**
-

Potential for Prevention and Compression of Morbidity

We now know how to accomplish
healthy and successful aging –
enough to get started.....

Translation, and new knowledge, will
be key

Chronic Diseases

- Prevention and treatment make BIG differences into the oldest ages:
 - Primary prevention: disease **onset**
 - Secondary prevention: disease **remission** or prevention of progression
 - Tertiary prevention: *minimize symptoms, outcomes; maximize quality of life*

Approaches Demonstrated: Prevention or Treatment of Chronic Conditions

	Med Tx/Prev	Exercise
Arthritis	+	+
High Blood Pressure	+	+
Osteoporosis	+	+
Diabetes	+	+
Heart Attack	+	+
Angina	+	+
Cancer	+	
Lung Disease	+	
Stroke	+	+

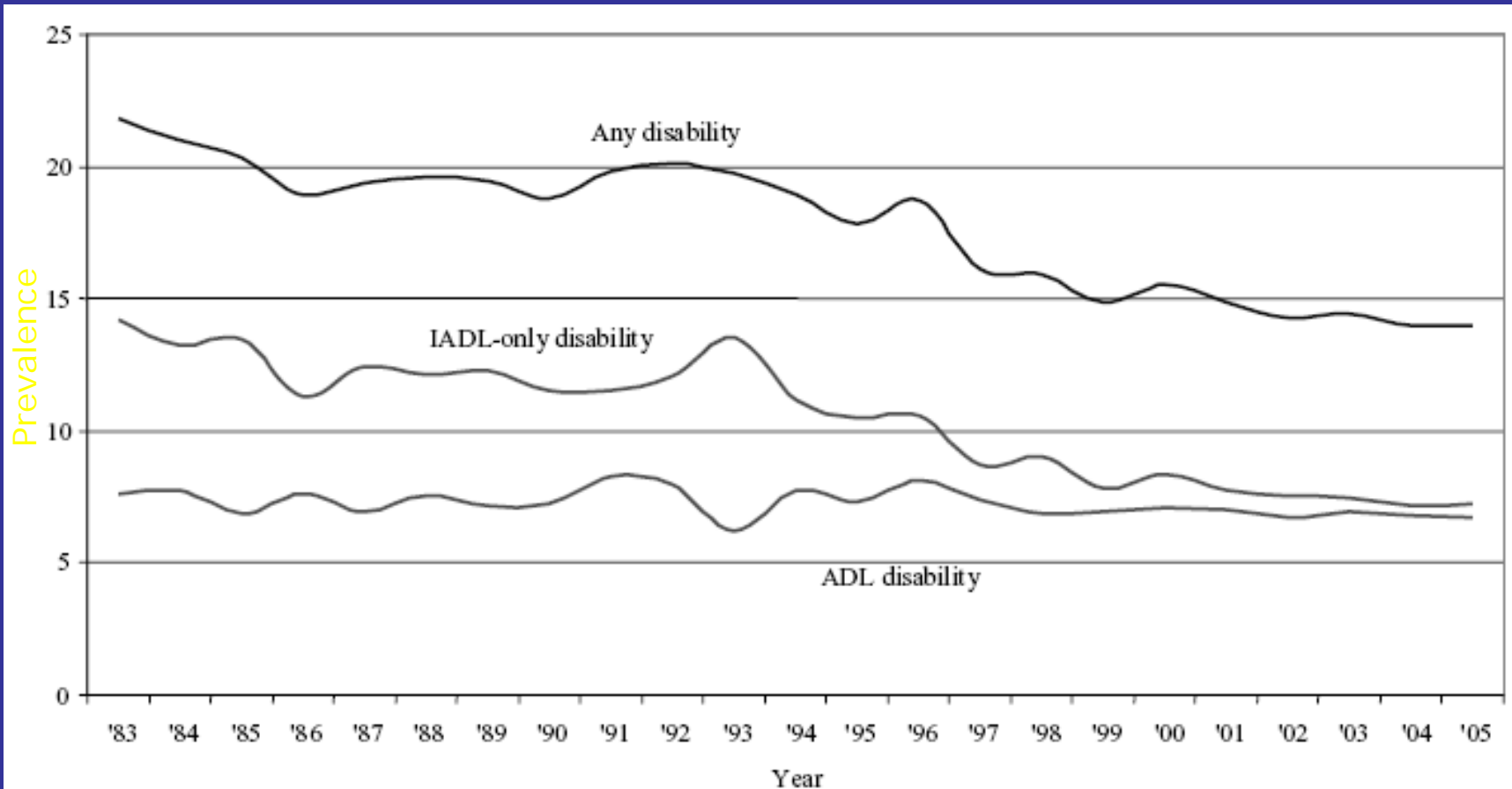
Impact of Effective Prevention: Change in U.S. death rates, 1950-1997

	Age Group	% change
Heart Disease	65-74	-58%
	75-84	-54%
	85+	-31%
Stroke	65-74	-76%
	75-84	-65%
	85+	-46%

Decline in CVD Mortality, U.S.

- Decline in CVD mortality, 1980-1990: 34% overall
- Contributing factors:
 - 25%: primary prevention of CVD
 - 29%: secondary prevention
 - 43%: improved medical care for patients with CVD

Disability Rates are Declining



Source: Authors' tabulations using the National Health Interview Survey.

FIGURE 1. Age-Adjusted Prevalence of Disability among the Noninstitutionalized Population 70 and Older, 1983 to 2005.

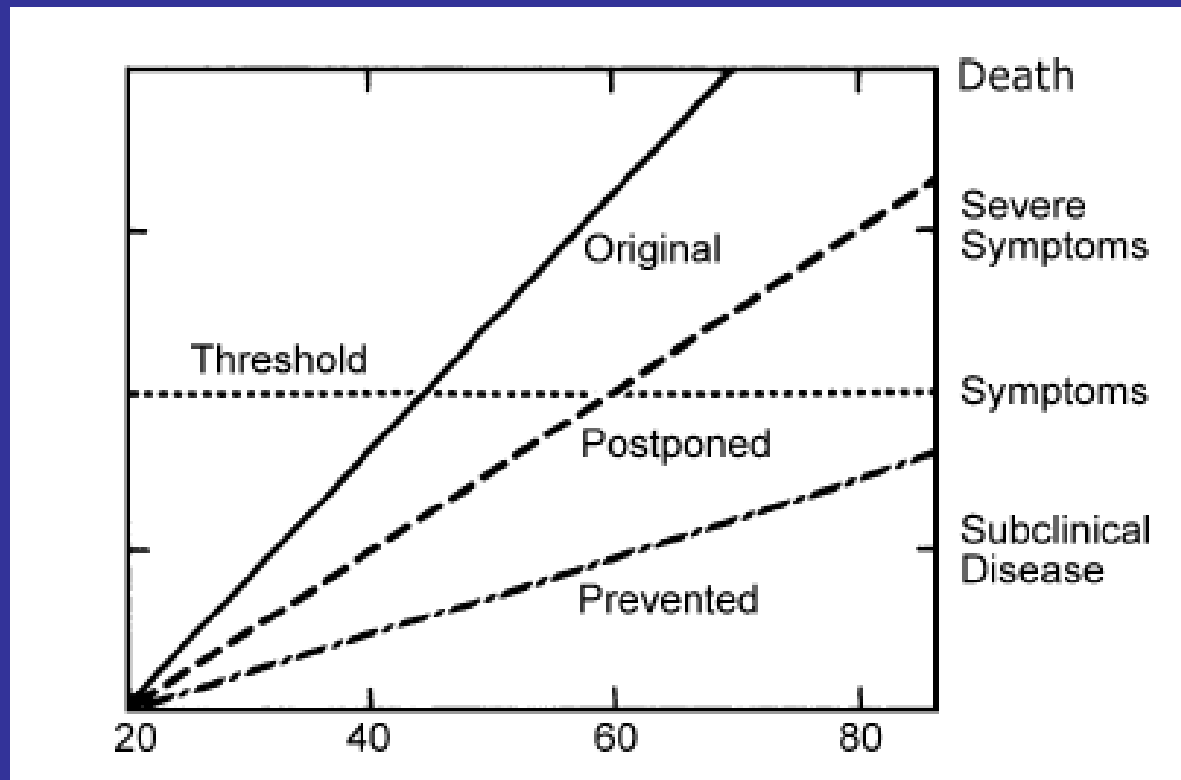
Freedman V, et al. *Milbank Quarterly*, 2008

What will healthy aging require?

1. Life course approach to Prevention of Chronic Conditions



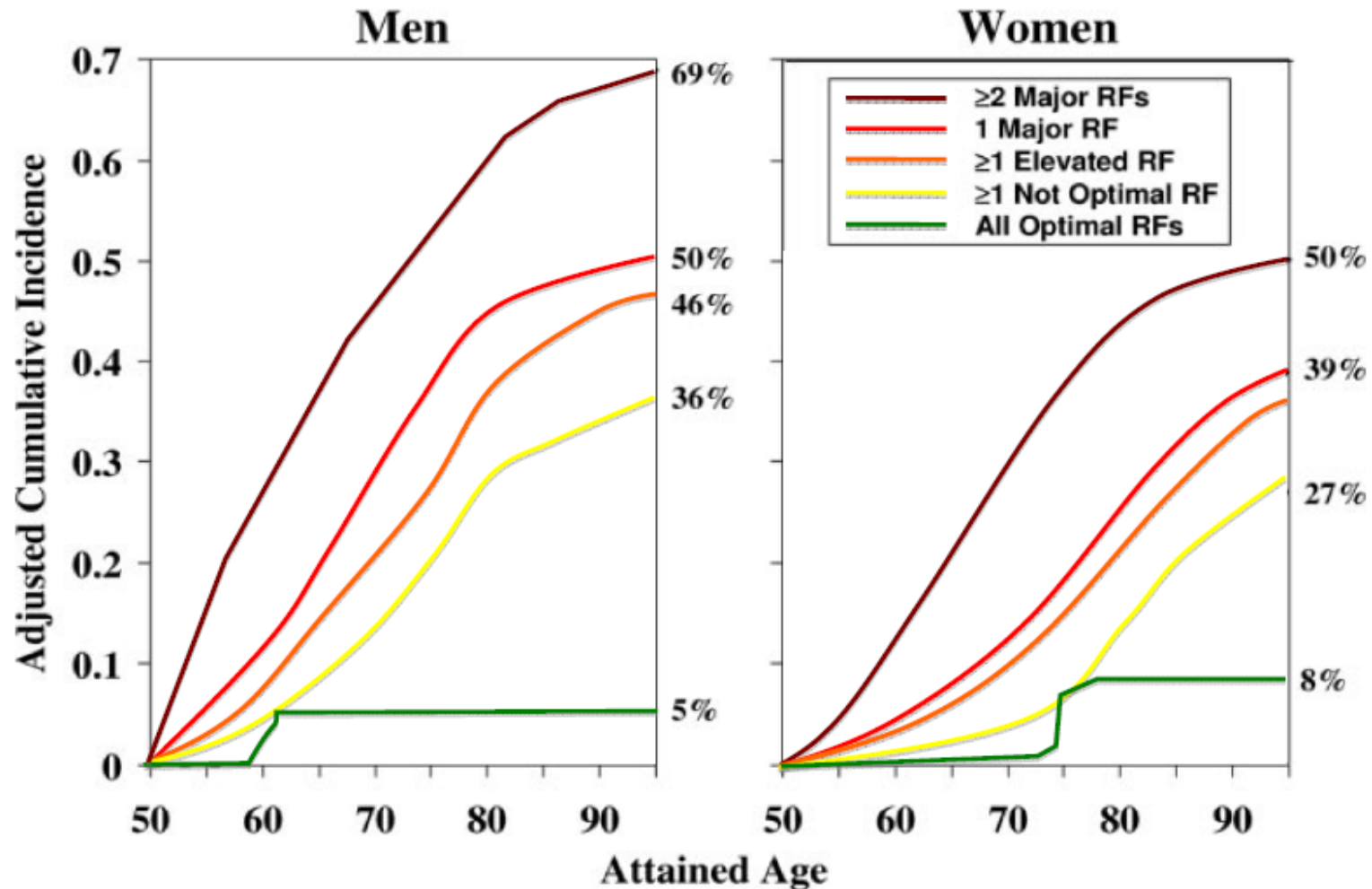
chronic diseases develop over many
years;
prevention is key



J. Fries. *The Milbank Quarterly*, 2005

Prediction of Lifetime Risk for CVD by Risk Factor Burden at 50 Years of Age

Donald M. Lloyd-Jones, MD, ScM; Eric P. Leip, PhD; Martin G. Larson, ScD; Ralph B. D'Agostino, PhD; Alexa Beiser, PhD; Peter W.F. Wilson, MD; Philip A. Wolf, MD; Daniel Levy, MD



Optimal risk factors: TC 180 mg/dL; BP 120/80 mm Hg, nonsmoker, and nondiabetic.
 Not optimal risk factors: TC 180 to 199 mg/dL, DBP 120-139 mm Hg, DBP 80-89 mm Hg.
 Elevated risk factors: TC 200-239 mg/dL, SBP 140-159 mm Hg, DBP 90-99 mm Hg.
 Major risk factors: TC >240 mg/dL, SBP >160 mm Hg, DBP >100 mm Hg, smoker, and diabetic

(*Circulation*. 2006;113:791-798.)

Predictors of cognitive decline and/or dementia in older ages

- Childhood exposures:
 - Education
 - Environmental risks and stressors
- Mid life:
 - Stroke risk factors and subclinical CVD in mid life predict cognitive decline
 - Mid life cognitive activity, joined with social activity, protective against onset of dementia in male twins; 25-44% decreased risk (Carlson et al 2008)
- Late life:
 - cognitive activity (Carlson) and physical activity (Kramer) associated with increased brain activation and cognitive function
 - Stroke predicts cognitive losses

Cumulative risk manifests at older ages

- Health disparities in older adults
 - African American older adults manifest
 - More chronic diseases
 - Disability 10 years earlier than whites

(Miller, Wolinsky, et al)

Excess disability in inner city African-Americans 49-65, compared with suburban sample

- Excess Disability:
 - 30% excess disability: walking one-half mile
 - 215% excess disability: managing medications

Miller D et al, J Ger: Med Sci, 2005

2. Knowledge of how health,
prevention and care needs change
with age

With longer age comes serious
health issues not found at younger
ages:

Geriatric conditions

Multimorbidity

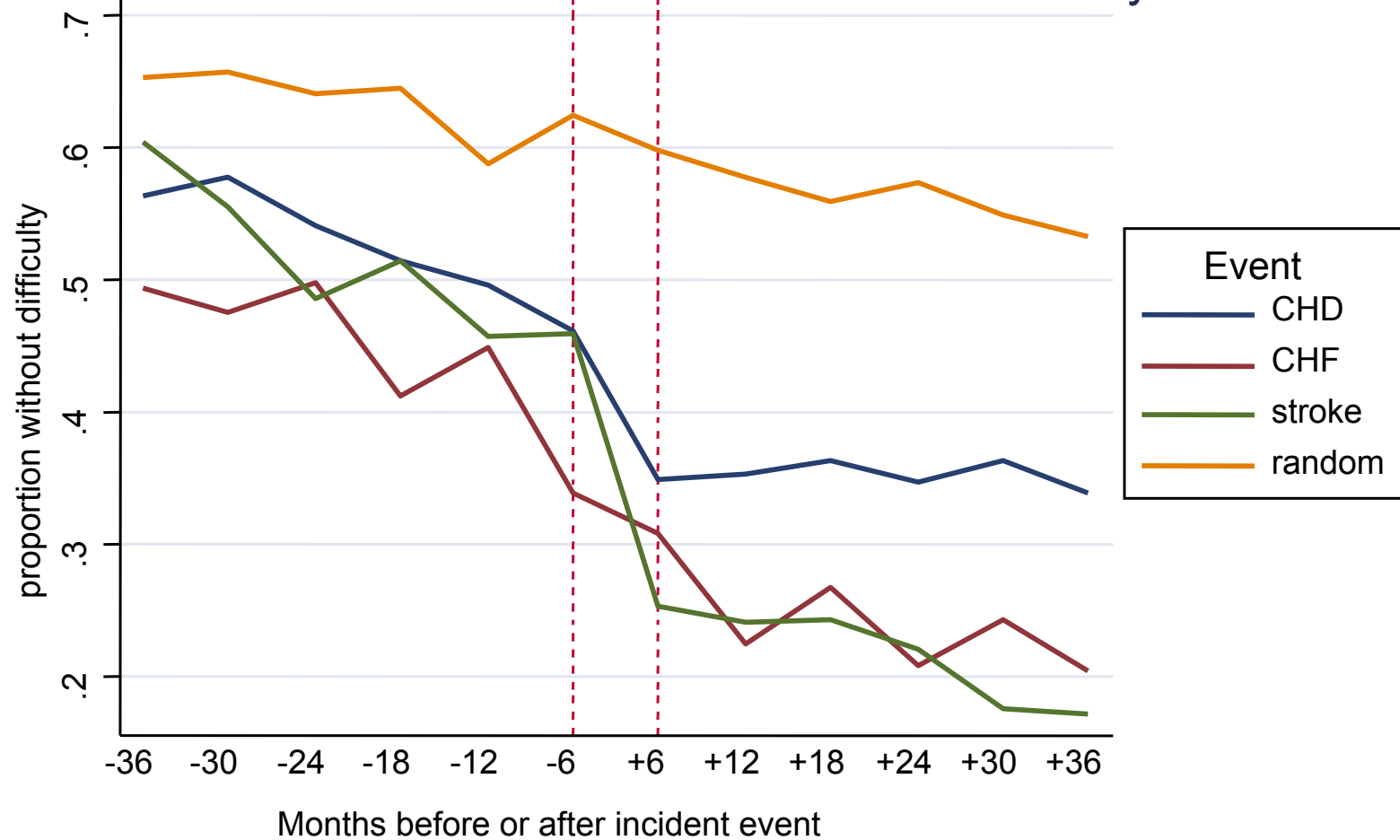
Frailty

Disability

Diseases/conditions can
contribute to other outcomes

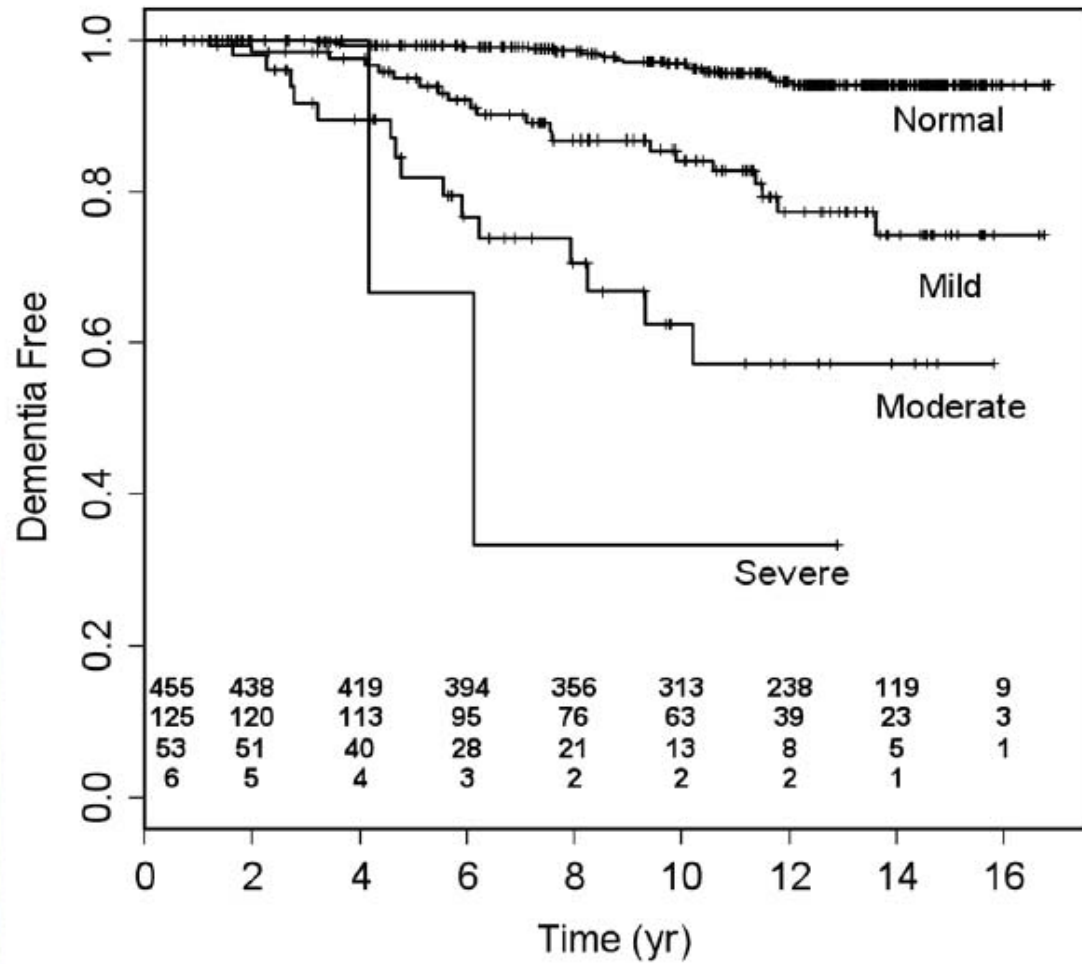
Prevention of distal outcomes is key

Onset of physical disability in high energy mobility tasks, men and women 65-101 in Cardiovascular Health Study



Hearing Loss and Incidence of Dementia

A longitudinal analysis in the *Baltimore Longitudinal Study of Aging*

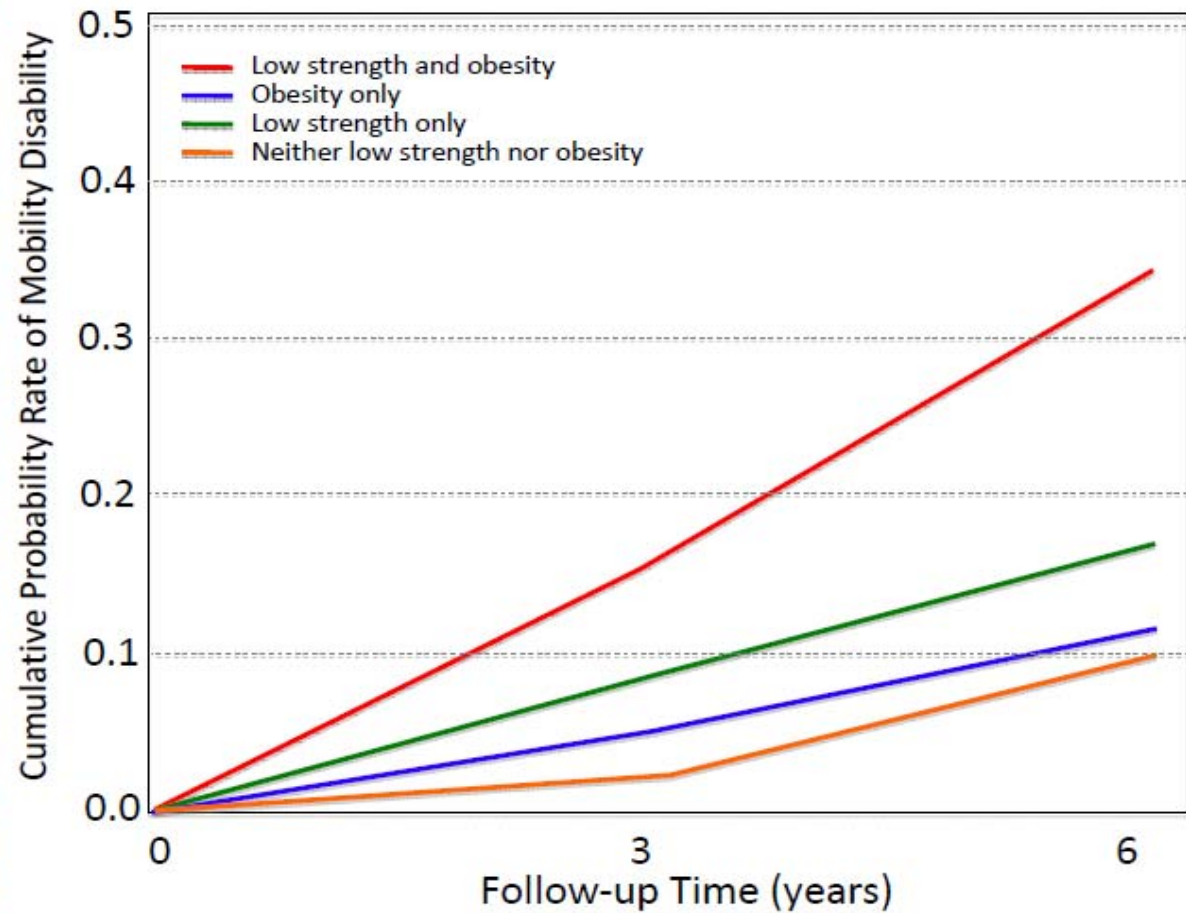


Submitted (not for citation)



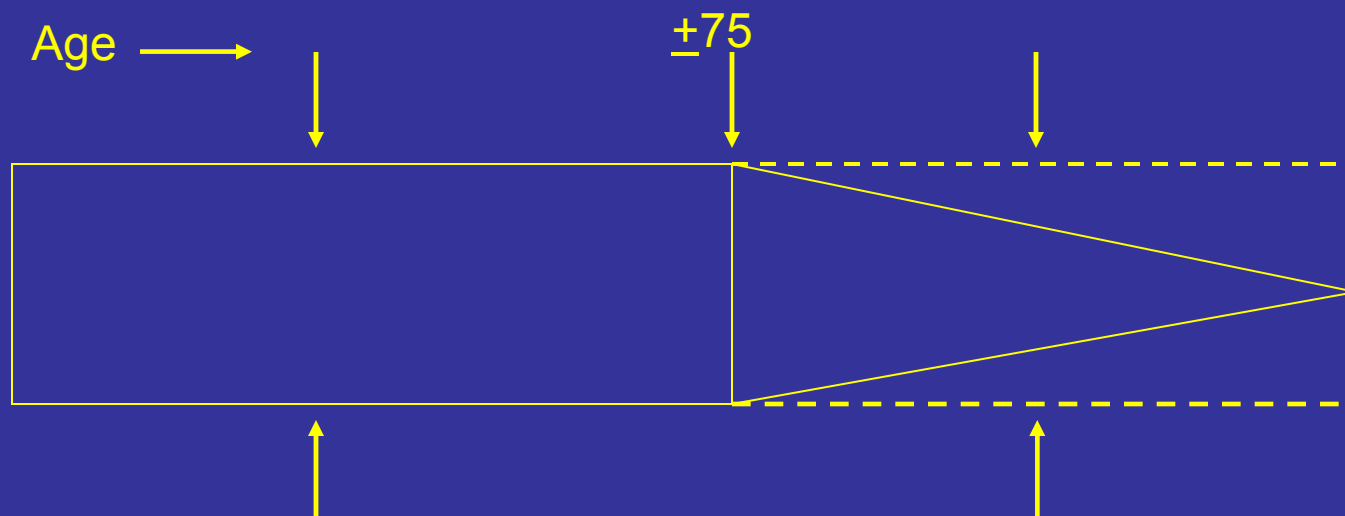
The Effect of Obesity Combined with Low Muscle Strength on Decline in Mobility in Older Persons: Results from the InCHIANTI Study

S Stenholm, D Alley, S Bandinelli, ME Griswold, S Koskinen, T Rantanen, JM Guralnik, and L Ferrucci.



Intrinsic physical and cognitive reserves evolve with aging

Baltes Hypothesis: loss of reserves with aging



- Decreased reserves necessitate:
- new approaches to understanding vulnerabilities in aging,
 - new approaches to prevention, care

Frailty Phenotype – Definition [1]

An individual is “frail” if ≥ 3 components - of 5 - are present

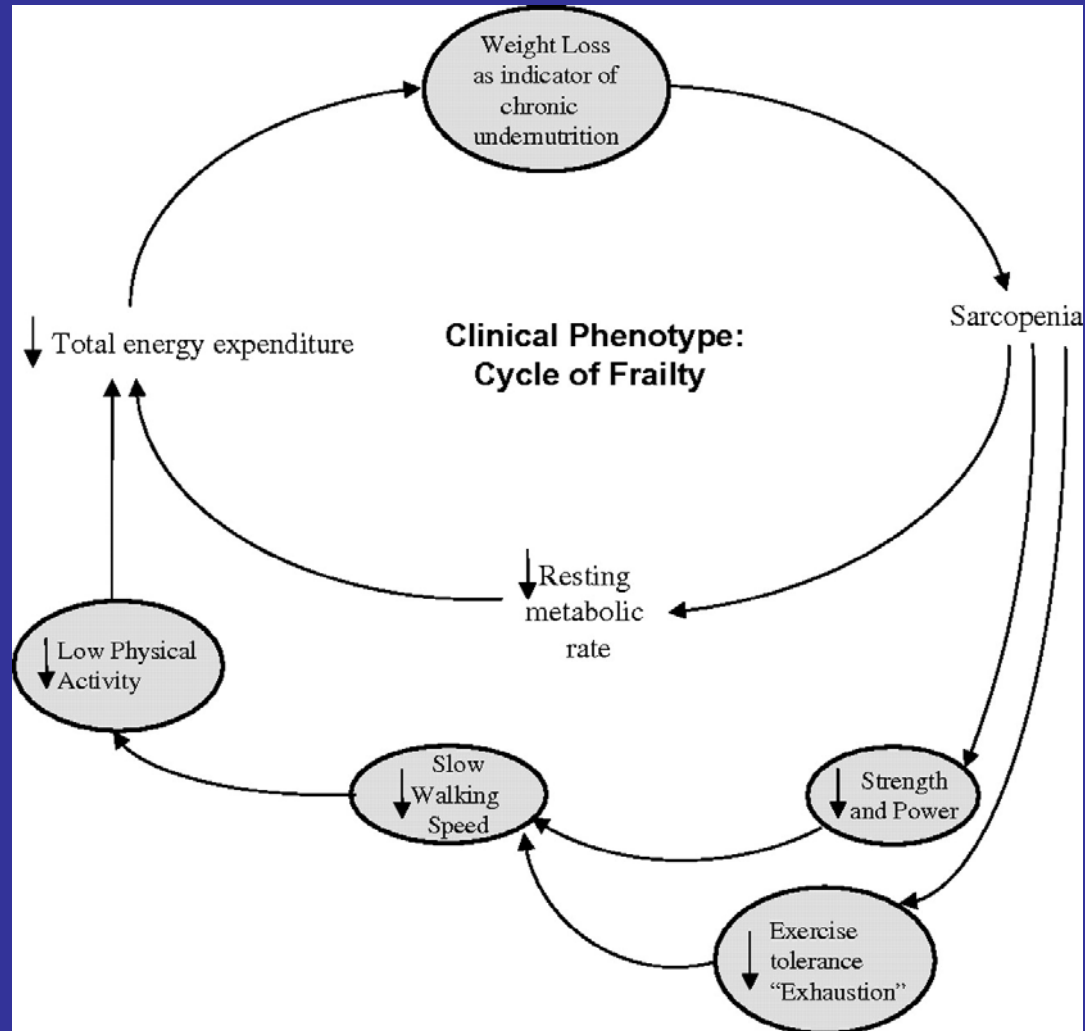
- **Physical shrinking (unintentional weight loss)**
- **Weakness (grip strength)**
- **Slowness (time to walk 15 feet)**
- **Low physical activity level (weighted score of kcal/week)**
- **Exhaustion (self-reported)**

Validated phenotype and medical syndrome [2]

[1] Fried LP, *et al.* J. Gerontol. A Biol. Sci. Med. Sci. 2001; 56:M146-56.

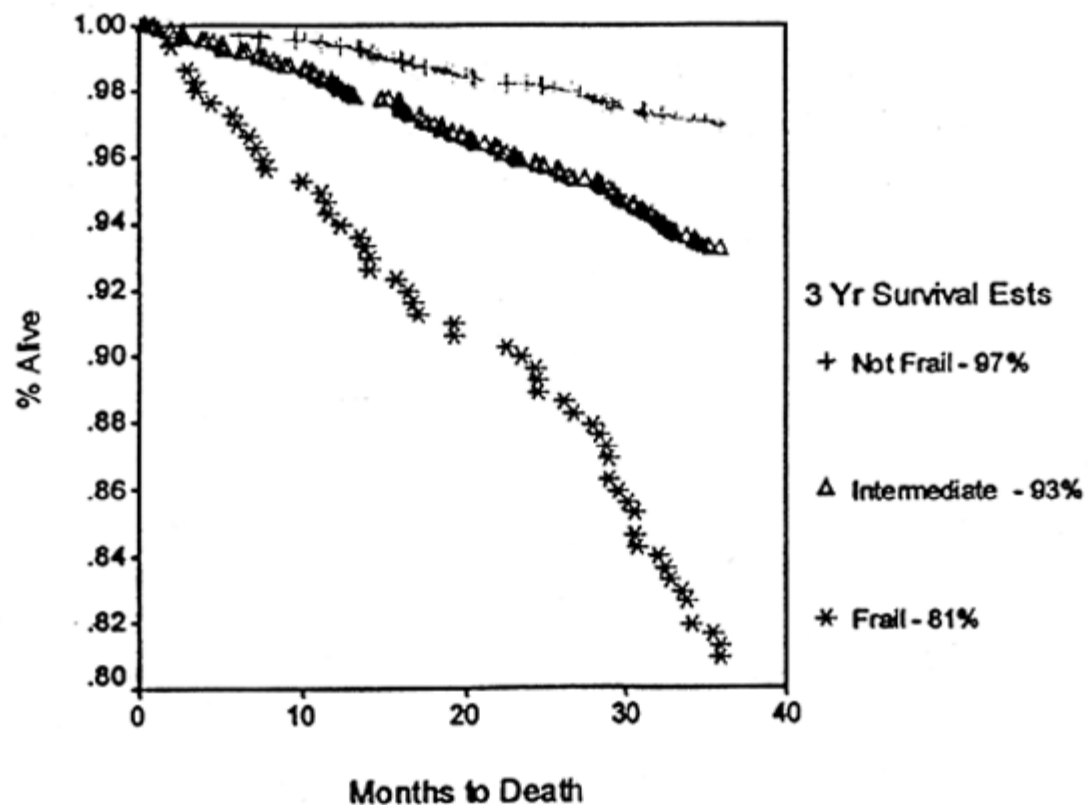
[2] Bandeen-Roche K, *et al.* J. Gerontol. A Biol. Sci. Med. Sci. 2006; 61:262-66.

Theorized and validated components of phenotype of frailty with aging, related in an adverse feedforward cycle



Frailty status predicts poor
outcomes

Unadjusted 3 Year Survival Estimates by Frailty Category



CHS: Baseline Frailty Status Predicts Adverse Outcomes Clinically Associated with Frailty

	Hazard Ratios* Estimated Over 3 Years
	Frail
Incident Fall	1.29
Worsening Mobility	1.50
Worsening ADL Disability	1.98
First Hospitalizations	1.29
Death	2.24

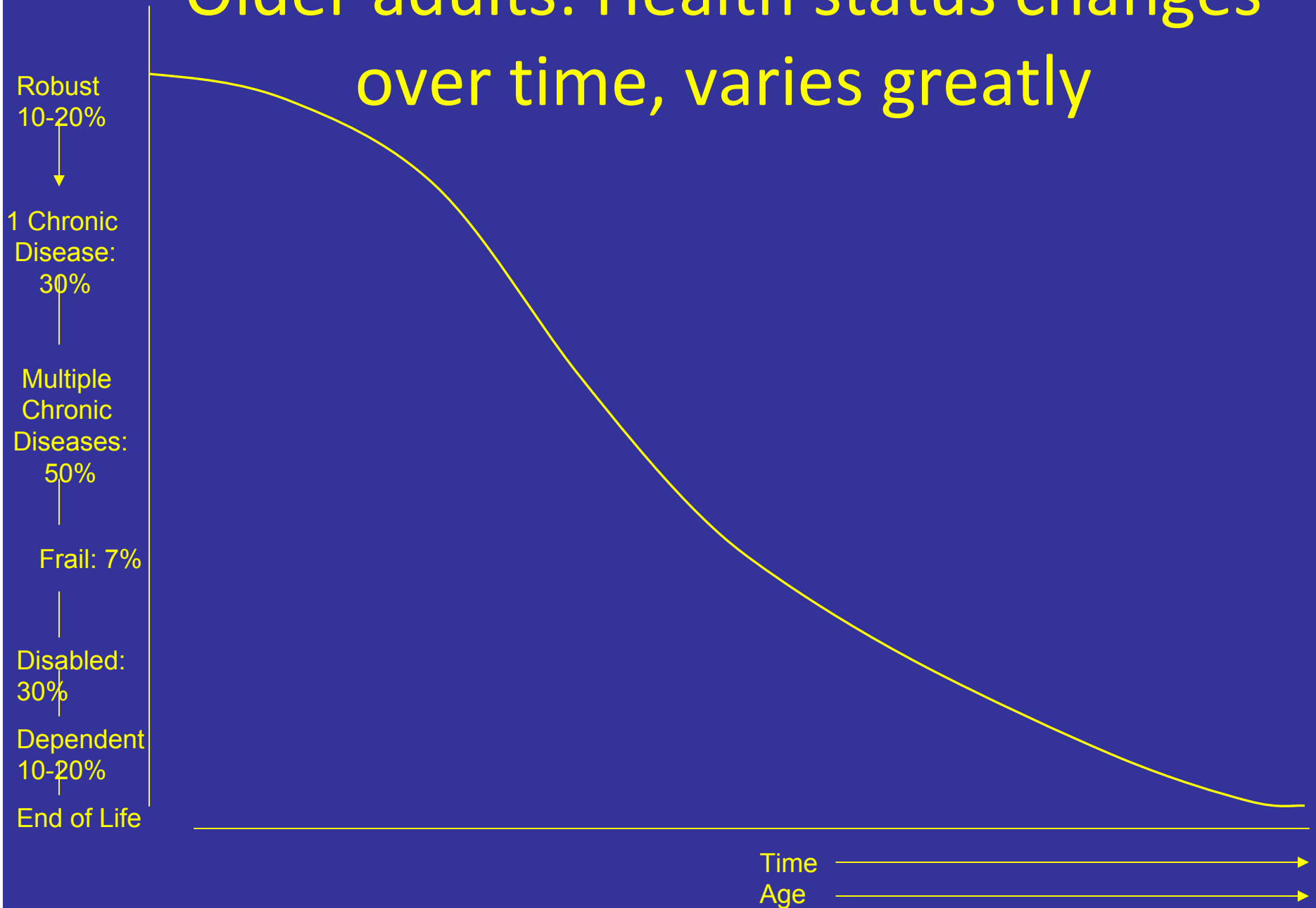
* Covariate Adjusted, $p \leq .05$

Fried et al, 2001

Theoretical Pathogenesis of Frailty [1,2]

- Aging-associated energy dysregulation syndrome
 - Phenotype is a vicious cycle
 - Triggered at multiple points in cycle
 - Triggered by many diseases as a final common pathway
 - E.g., CHF, COPD, diabetes, cancers, renal impairment, HIV, possibly others
 - Greater number of abnormal physiologic systems associated nonlinearly with increased risk of frailty (emergent property)

Older adults: Health status changes over time, varies greatly



Older age has many faces



3. Health promotion for those living with HIV

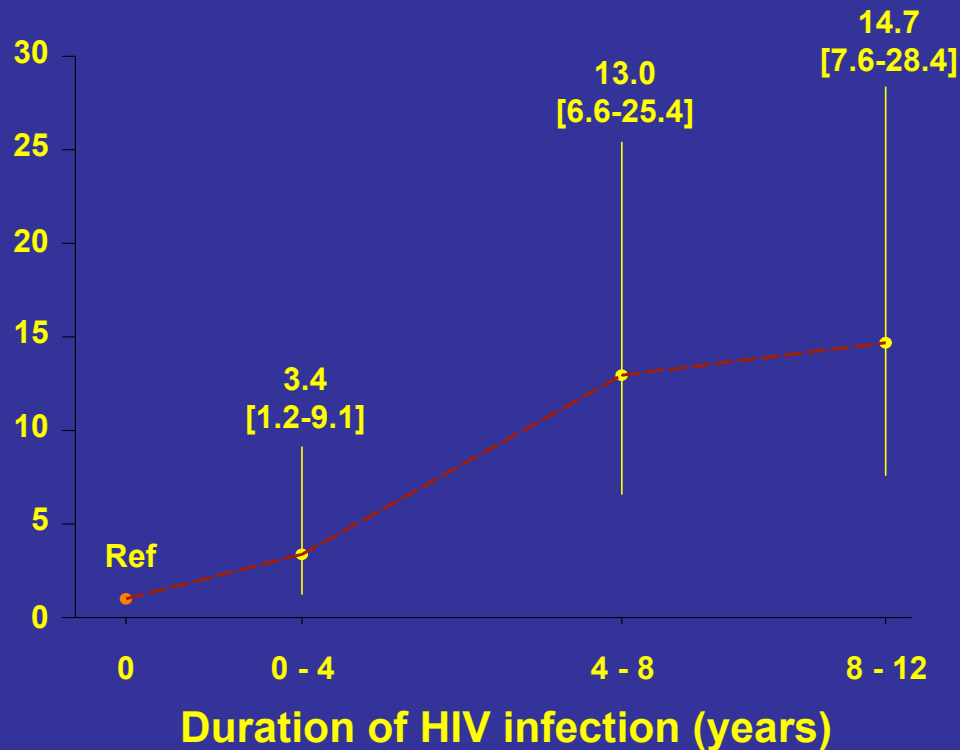
Aging-related needs

Current HAART Era: Age Issues

- HIV as a “chronic disease”
 - Chronic disease care and prevention principles
 - Co-morbidities / toxicities (malignancies, metabolic perturbations, cardiovascular, kidney, liver, neurologic) - which have an age component
 - Geriatric conditions:
 - Polypharmacy
 - Falls
 - Frailty
 - Disability
 - Mortality

Duration of HIV Infection is Associated with Presence of Frailty-related phenotype (FRP) - Pre-HAART

Odds ratio [95% CI] to manifest the FRP*



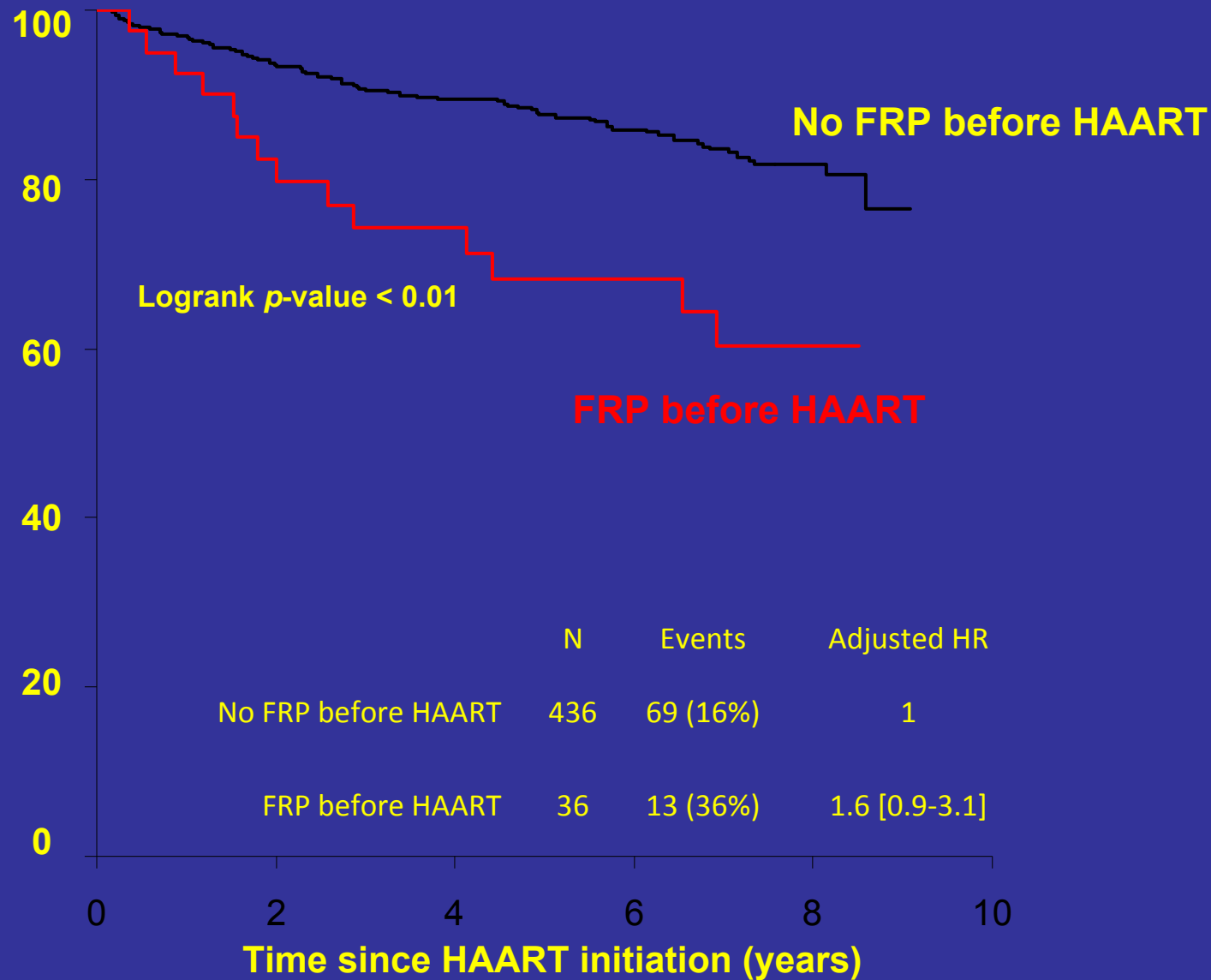
Same FRP prevalence between a 55-year old man infected < 4 years and a >65-year old uninfected man

*Logistic regression models (GEE)

Desquilbet L, et al, *J Gerontol A Biol Sci Med Sci* 2007; 62:1279-86.

Prognostic Effect of FRP on HAART Response – Results

% alive without clinical AIDS



Age-specific and health needs-specific public health and health care approaches matter

Integration of public health and medicine
– in both prevention and care

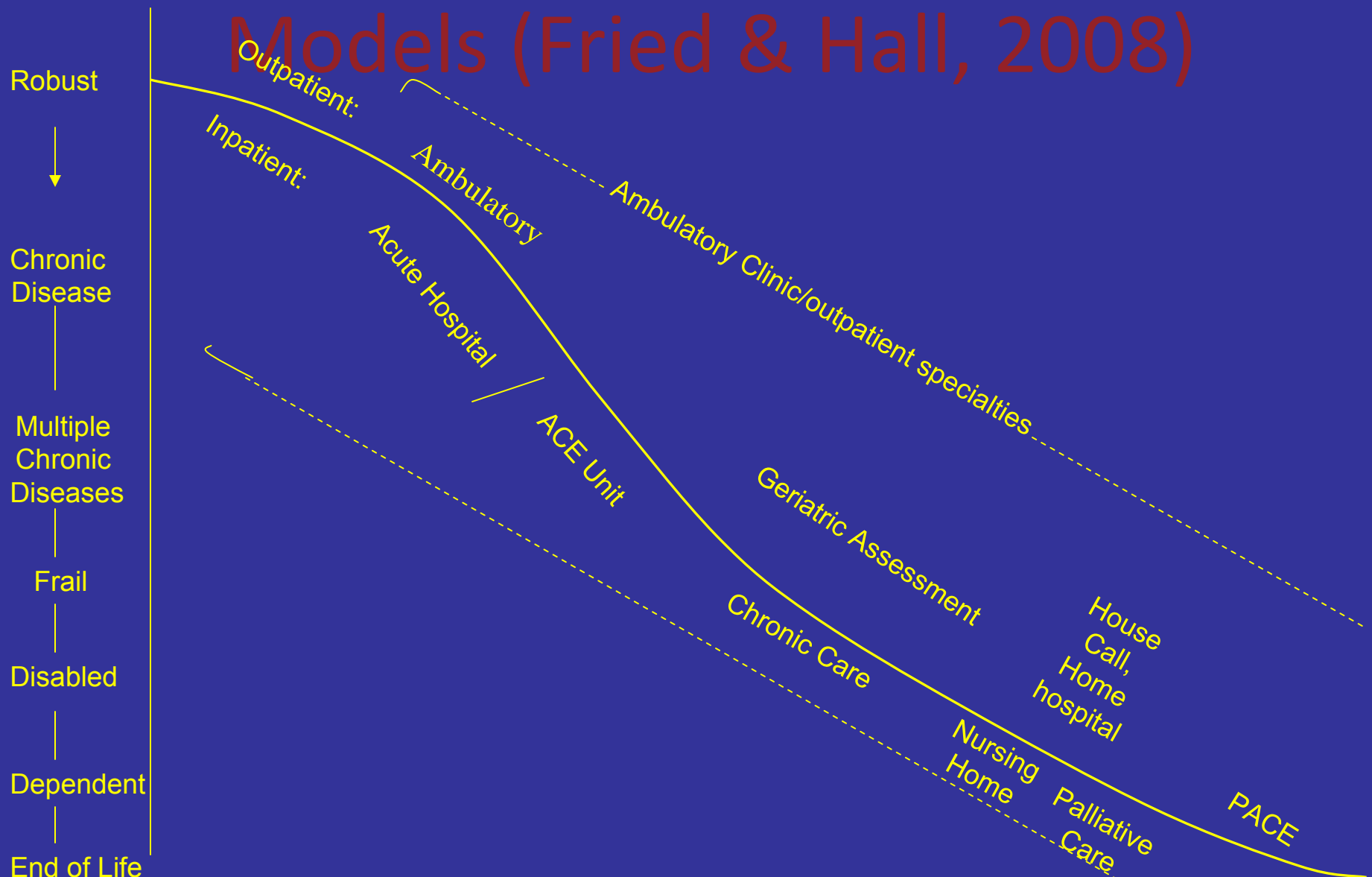
4. An older population will require new approaches to both public health and health care delivery

- Transformed Systems: care and prevention
- Contents: Geriatrically knowledgable
- Distributed locations – for health promotion & care
- Integration of public health and medical care
- Workforce: geriatrically knowledgable
- Environment: built and physical
- Opportunity: for giving back, productive and social engagement

Challenge: The Health Care System Not Ready for Chronic Care Needs of Older Adults

- Acute event-based care in a chronic disease world
- Effective geriatric care not implemented or reimbursed
 - Community-based coordinated continuum of care needed
- Prevention needs to be inserted into medical care and linked to community-based approaches
 - *Institute of Medicine. Retooling for an Aging America: Building the Health Care Workforce, 2008*

Continuum of Geriatric Care Models (Fried & Hall, 2008)



There are now evidence-based interventions for geriatric syndromes that *improve* outcomes

- Medications management
- Delirium
- Falls
- Incontinence
- Weakness, frailty, mobility
- Function, disability
- Transitions in care

These interventions are linked to positive outcomes

Sarcopenia modifiable

- In “frail”, disabled nursing home patients, resistance exercise increased:
 - muscle mass by 180%
 - strength by 100%

Fiatarone et al, 1993

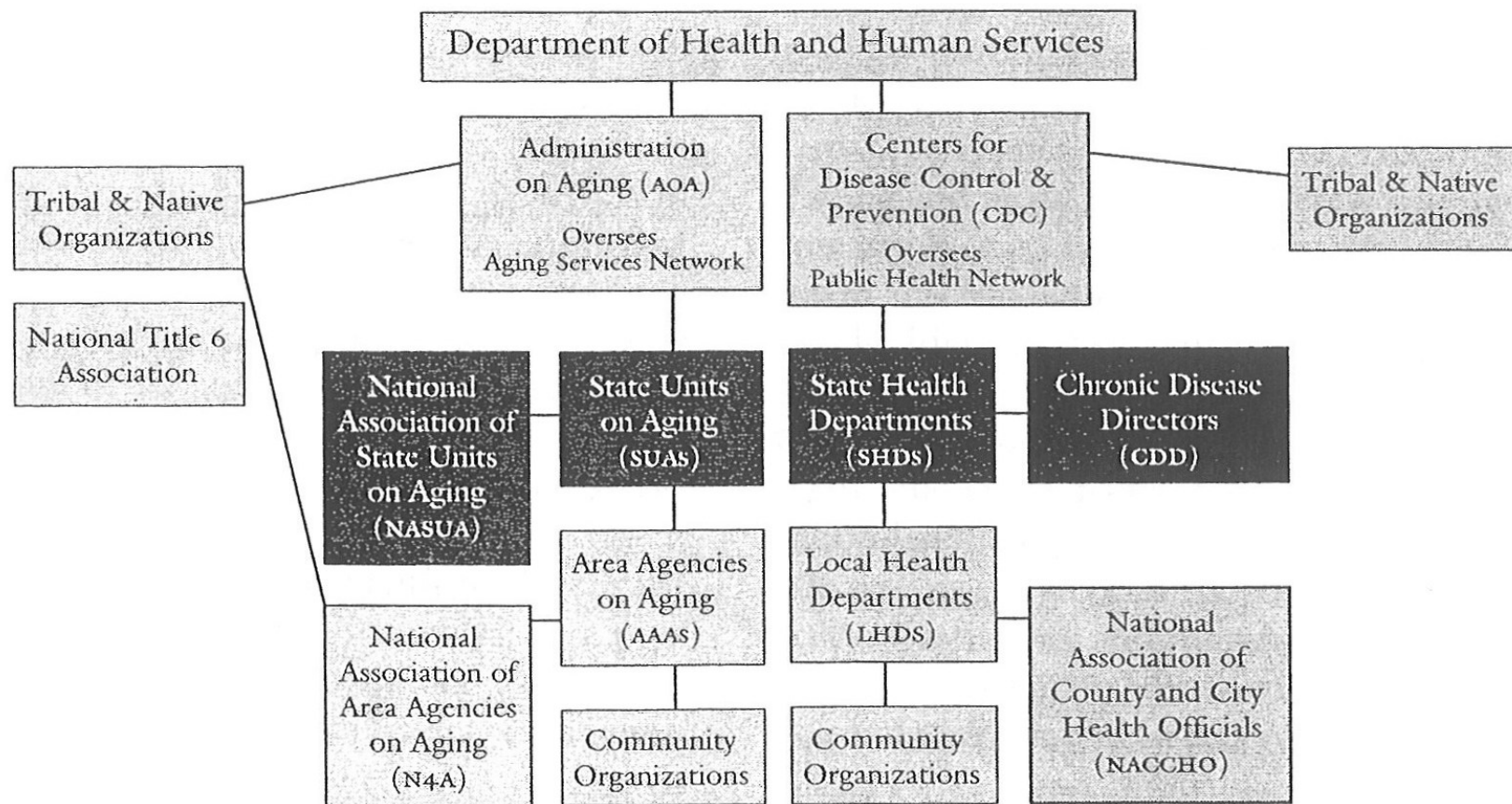
Building Blocks for Effective Health Promotion for an Aging Population beyond Disease Focus

Access to Care	Meaningfully engaged	Financial Resources	Environmental Stressors
Polypharmacy	Disability	Health Disparities	
Frailty	Multimorbidity	Falls	Depression/Loneliness
Primary, Secondary Prevention; Disease Management			

Key Policy Challenge: Who is on first?

- Public health and aging services networks are siloed

Public Health and Aging Services Networks



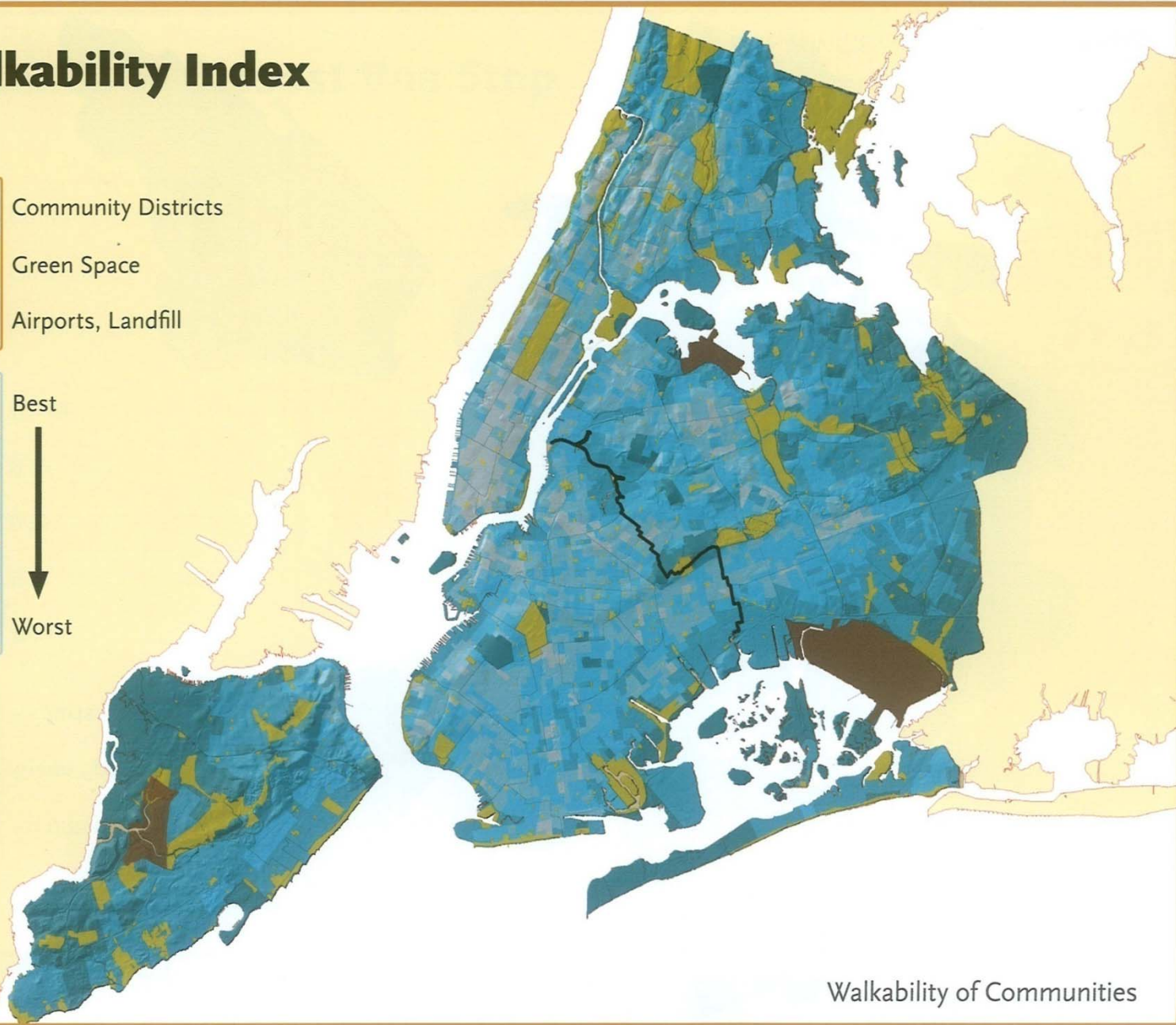
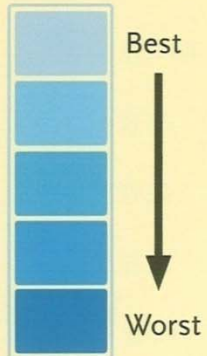
Designing New York to support
successful and healthy aging

Age-Friendly City: Key Domains (WHO)

- Housing
- Transportation
- Outdoor spaces and buildings
- Social participation
- Civic engagement and employment
- Respect, social inclusion v. ageism, fear
- Communication re: services
- Community support and health services

Accomplishing positive health behaviors for an aging population will require novel, multilevel, community-based approaches

Walkability Index



Walkability of Communities

Maps prepared by the Built Environment and Health Project, Columbia University (www.beh.columbia.edu) in collaboration with The New York Academy of Medicine (www.nyam.org).
Data sources include: US Census Bureau, Census of Population and Housing, 2000 and the NYC Department of City Planning.

Increasing Physical Activity in Older Adults: public health and health care working together

- Physician prescription
- Communities that permit/support being active
- Exercise programs
- Community-based approaches to enhance usual physical activity

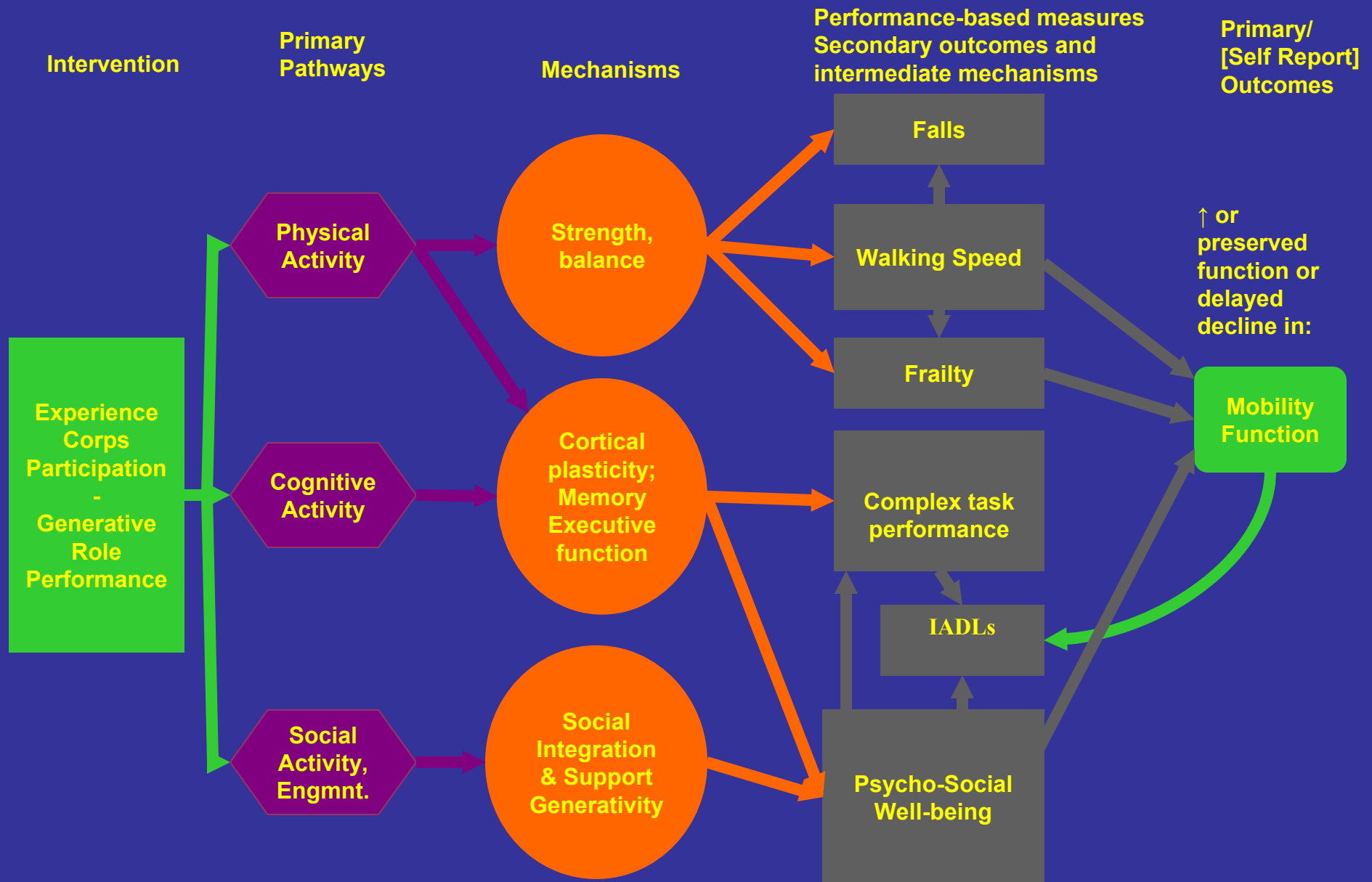
Community-based approaches to
health promotion:

Generativity as a vehicle

Meaningful roles to make a
difference:

Experience Corps as an example

Causal Pathway: Experience Corps



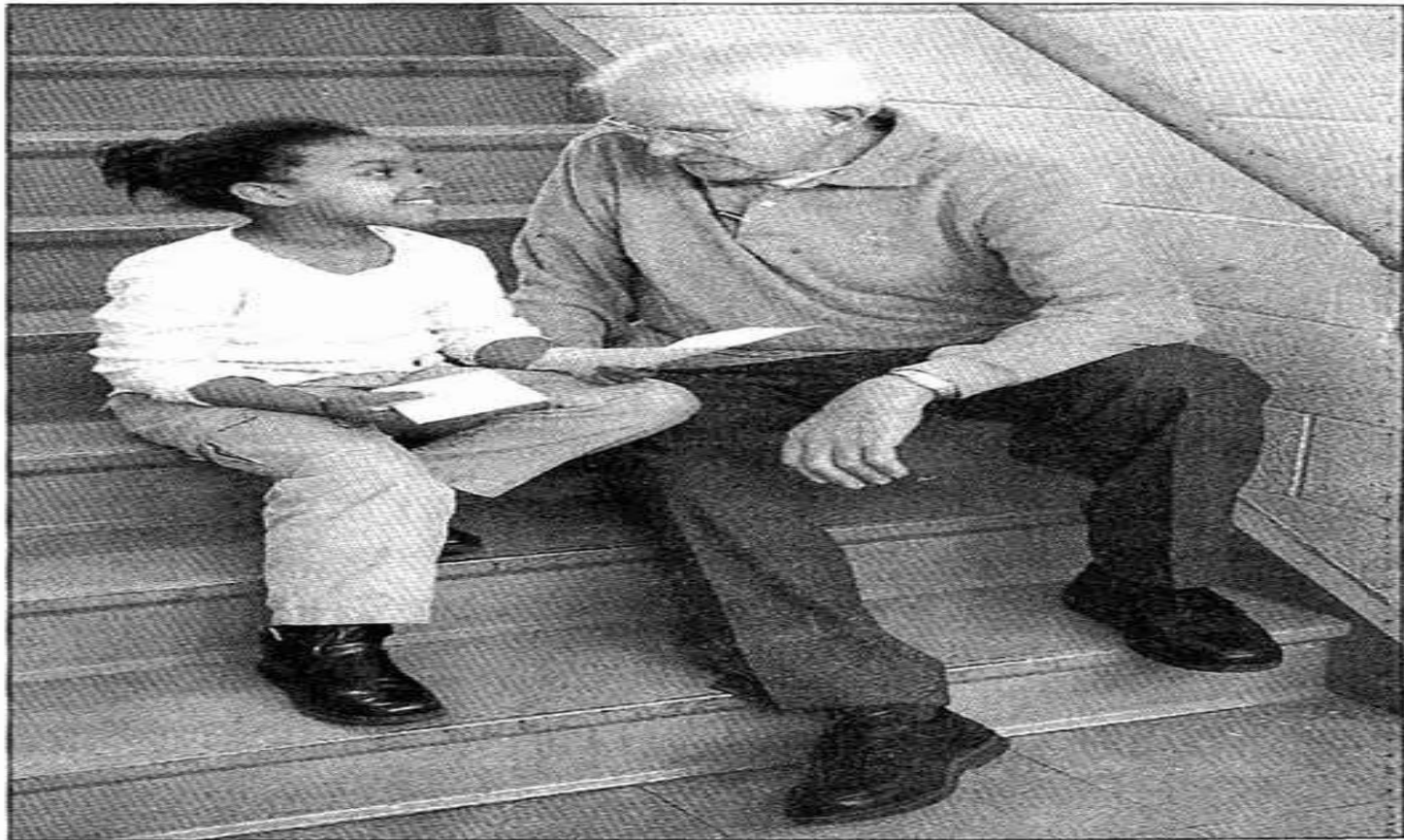


Preliminary evidence: EC Baltimore Pilot RCT, Physical Activity

- For those previously with low activity:
 - Increased physical activity within range obtained by exercise trials (≤ 700 kcal/wk)
 - sustained dose; high retention
 - Increase sustained over 3 years, relative to comparison group
- Participation by high risk subsets, including African American older adults

(Tan et al, 2006, 2009)





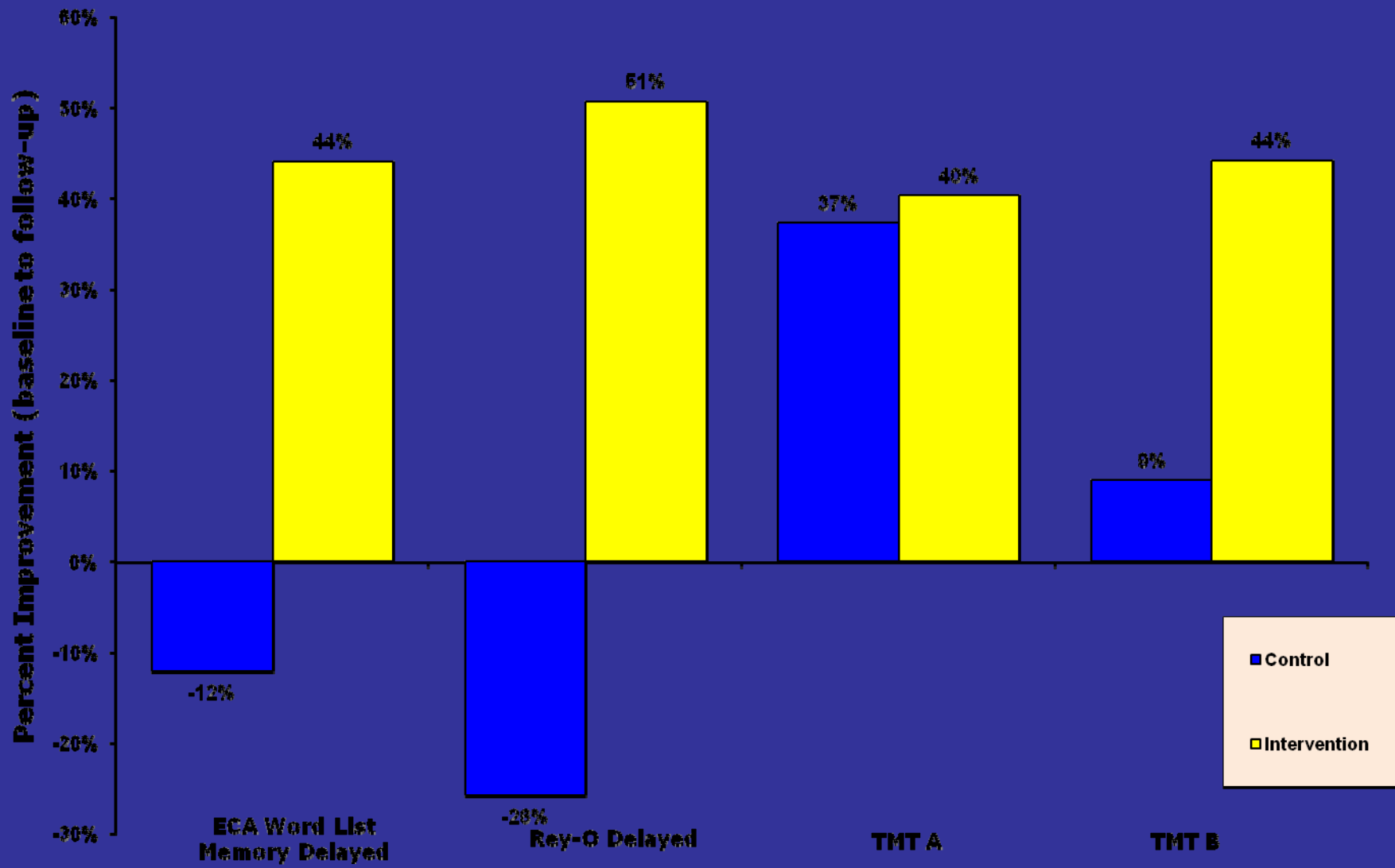
**“IT SEEMED A GOOD
OPPORTUNITY TO
HELP OUT.”**

DICK FRYER, 73, HAS A PRIVATE CHAT WITH A SECOND-GRADER OUTSIDE THE CLASSROOM. A RETIRED CIVIL ENGINEER FROM PERRY HALL, FRYER LEARNED ABOUT THE EXPERIENCE CORPS THROUGH AN AARP MAILING.

Change in Executive Function in EC volunteers v. controls

- Low normal executive function at baseline
- 6 month follow-up

(Carlson et al 2009)

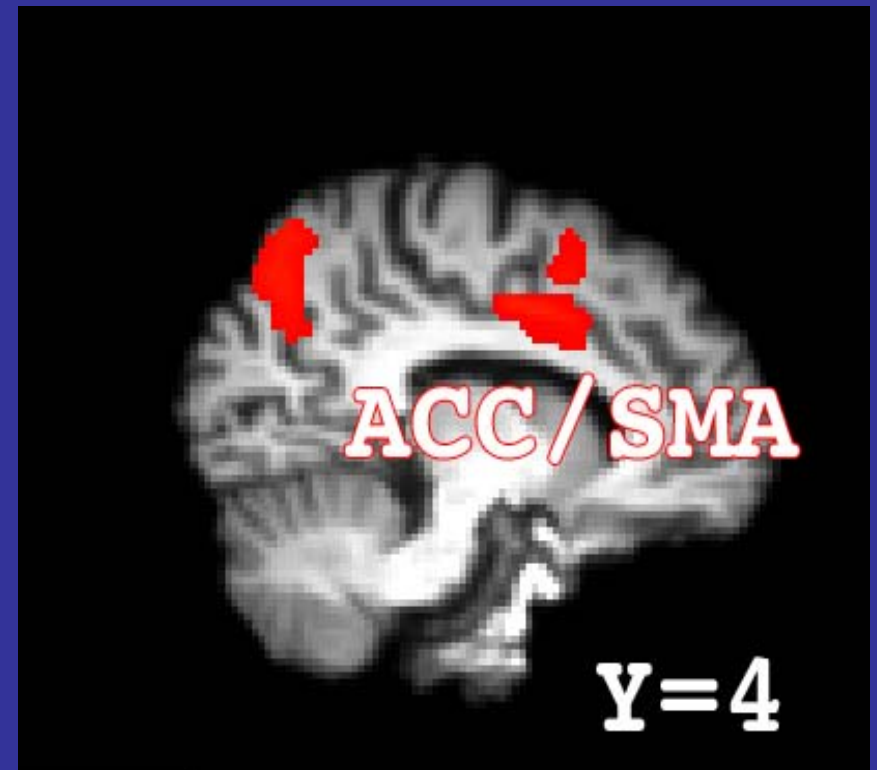
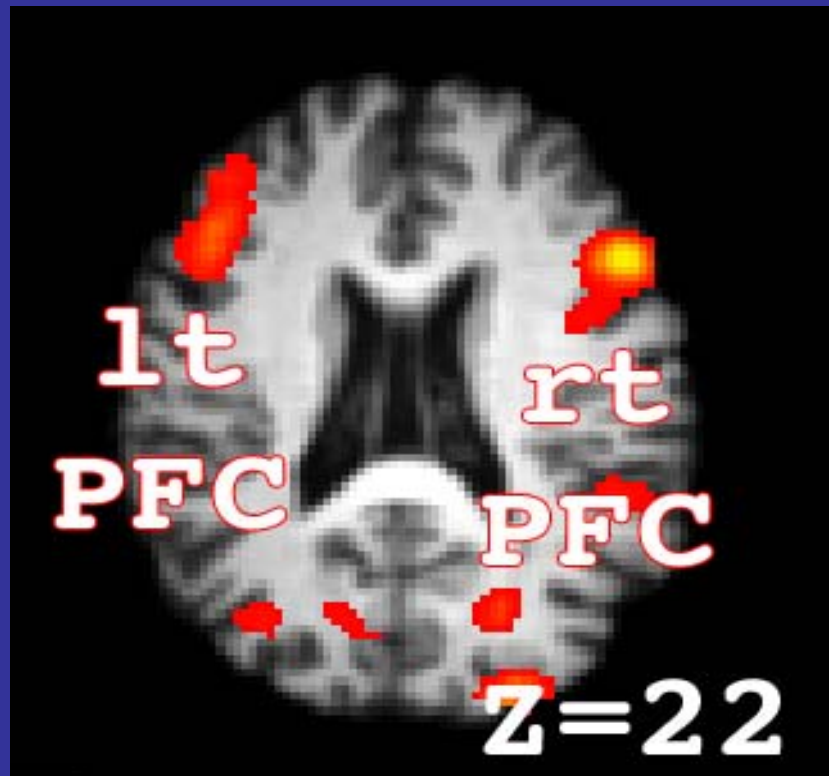


Evidence for neurocognitive plasticity in at risk older adults: EC pilot study

- EC (8) v. matched Controls (9) at risk for cognitive impairment:
 - African-American women, mean age 68, HS education, low income, mean MMSE 25-26
- Baseline and 6-month follow up
- Intervention-specific short-term gains:
 - Executive Function
 - Activity of prefrontal cortical regions (fMRI)

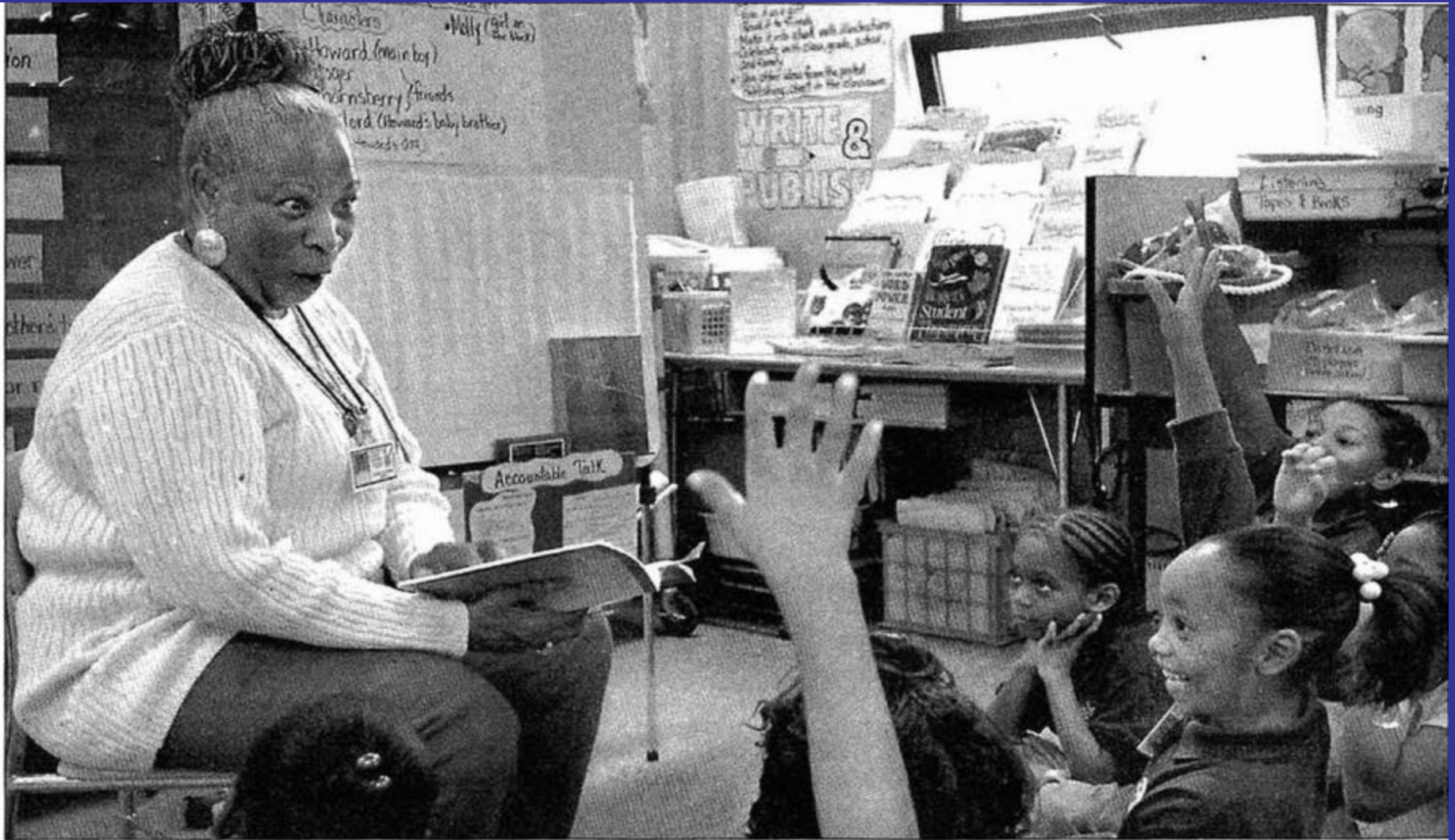
Carlson et al JGMS, 2009

EC participants > Controls on test of executive function following 6 month exposure



PFC= prefrontal cortex; ACC= anterior cingulate cortex

Carlson, Erickson, Kramer, ...Fried, JGMS 2009



"YOU ARE A FRIEND IN THE CLASSROOM."

AUDREY WEEMS, 70, READING A STORY TO STUDENTS IN A THIRD-GRADE CLASS AT WAVERLY. A MOTHER OF EIGHT, SHE WORKED AT THE SOCIAL SECURITY ADMINISTRATION FOR 35 YEARS, RETIRING IN 2002. WEEMS LEARNED ABOUT THE BALTIMORE EXPERIENCE CORPS PROGRAM THROUGH HER CHURCH.

Role of Public Health in Goals for an Aging World

- Population goals: “compression of morbidity”
 - Individuals: Health behaviors; life course perspective for health promotion
 - Health system redesign: hospital, clinic, home & community; integration
- Design for an aging society which all ages will benefit from
 - Families, communities and cities

Healthy, successful aging:
a critical societal goal







PUBLIC HEALTH AND COMMUNITY HEALTH: OPTIMIZING HEALTH IN AGING		FUNCTIONAL STATUS SPECTRUM: OLDER ADULTS					
					Dependent		End-of-Life/ Hospice
		Community			Home-Based or Institutionalized Care		
		30%	30%	20%	10-15%		
TOPICS/ISSUES		High Functioning/ Robust	Usual Functioning - at risk	Low Functioning	Assisted Living	Longterm Care	
A	MOBILITY / PHYSICAL ACTIVITY						
B	CIVIC ENGAGEMENT						
C	MENTAL HEALTH						
D	ISSUES OF ISOLATED OLDER ADULTS						
E	*EMERGENCY PREPAREDNESS/ SAFETY/ MONITORING						
F	ENSURING RECEIPT OF NEEDED CLINICAL CARE						
G	CHRONIC DISEASES PREVENTION & MANAGEMENT						

FRP in the 3 years before HAART initiation independently predicts subsequent clinical AIDS or death (N= 596 men in MACS) - adjusting for other predictors (multivariate Cox models)

Exposures at HAART initiation	AIDS-free at HAART (N=472)		AIDS at HAART (N=124)	
	aHR ¹	P-value	aHR ²	P-value
Education \geq college	1.01	0.96	0.87	0.70
Ethnicity = White non Hispanic (vs others)	1.32	0.45	0.64	0.38
Age (per 10 years increase)	1.43	0.03	1.31	0.32
Nadir CD4+ T-cell count (per 100 cell/mm ³ increase) ³	0.85	0.05	0.94	0.61
Maximum plasma viral load (per 1 log ₁₀ copies/ml increase) ³	2.08	< 0.01	1.31	0.38
<i>Proportion of FRP + visits before HAART (for a 25% increase)³</i>	1.35	0.04	1.63	<0.01

FRP, frailty-related phenotype; aHR, hazard ratios adjusted for variables listed in the table; ¹ adjusted hazard ratios for AIDS/death; ² adjusted hazard ratios for death only; ³ within the 3 years before HAART

Colin A Depp and Dilip V Jeste

Definitions and Predictors of Successful Aging

A Comprehensive Review of Larger Quantitative Studies

Components for the Definition of Healthy Aging

- Disability Physical Functioning
- Cognitive Functioning
- Life satisfaction/ Well-Being
- Social/Productive Engagement
- Illness/Disease
- Longevity
- Self-Rated Health
- Personality
- Environment/Finances
- Self-rated successful aging

Physical disability in community-dwelling persons 70 and older, U.S.: due to chronic diseases

- 10% incidence/year

- Difficulty: Men Women

- Walking 30% 33%

- IADLs 13% 7%

- ADLs 22% 17%

NCHS, 1999

Preliminary Evidence of Intermediate Effects: EC vs. Controls

- Behavioral Risk Factors:
 - Physical activity
 - Cognitive activity
 - Social supports
- Intermediate effects:
 - Cognition (executive); brain activation
 - Strength, performance, energy, falls
 - Depressive symptoms

Why is Executive Function Important?

- Ability to plan, initiate, and carry out a course of action, shift flexibly and modify goals
- Integral to performing many independent activities of daily living (e.g., Grigsby et al., 1998; Carlson et al., 1999)
- Age-related changes in executive function may precede changes in memory (Carlson et al.)

Large-scale RCT of Experience Corps being conducted in Baltimore, MD

- Funded by NIA BSR, plus AmeriCorps, Weinberg Foundation, MacArthur Foundation
- Initiated Fall 2006
- Randomizing 600 people 60 years and older to EC or low-activity volunteering control;
- 20+ public elementary schools to EC; matched controls
- Outcomes for older adults: physical disability, mobility, frailty, fall risk, memory and intermediate neurocognitive changes
- Outcomes for children: standardized scores;

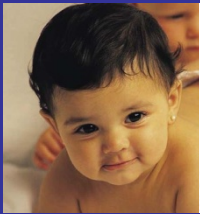
Implications of Experience Corps Design

- Generative potential attracts and fulfilled generativity retains older adults
- Lifestyle activity, in a 15-hour/week dose, shows increased physical, cognitive and social activity and is neurocognitively protective
- High longterm retention: sustained dose of prevention
- Designed for a win-win
 - Social model for health promotion
 - Harnessing social capital of aging society

New York: growing older

- 2005, New York City
 - 65 and older: 12% of New York City; 17% of some neighborhoods
 - 43% with some disability
 - 18.1% poor (U.S. 9%)
 - 12.1% poor and disabled (U.S. 5.5%)

Life course approach to healthy aging is essential



Chronic Disease & Susceptibility Over the Life Course

- Cumulative disease processes develop chronically and progressively over the life course
- Initiators, Promoters, Modifiers and Mediators are active
- In many trajectories, exposure and intervention at any point can make a difference; however, there may be points of particular opportunity for prevention
- At different ages, additional age-specific issues emerge and may require unique interventions

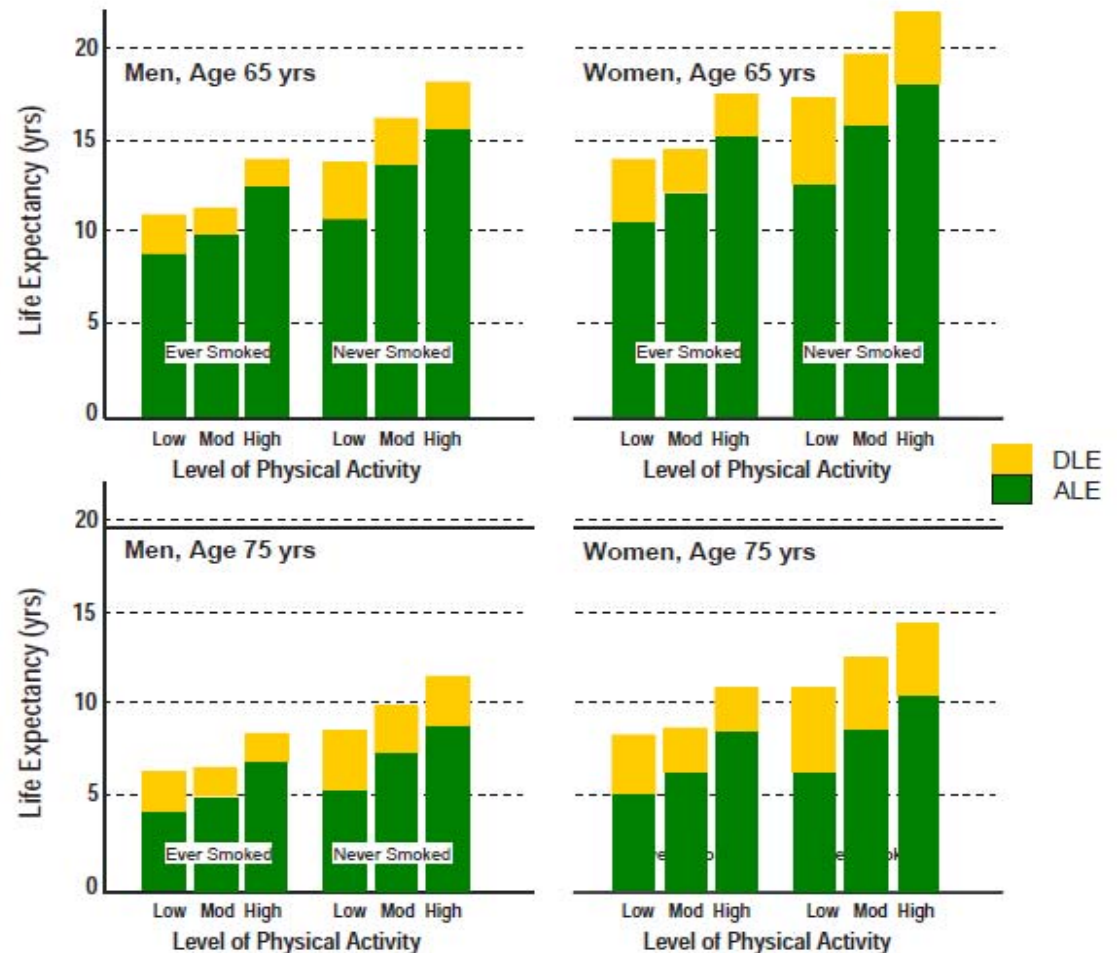
"... the ills that flesh is heir to... it must be that many a taint grows deeply, mysteriously grained in their being ..."
-Virgil, *The Aeneid*, 1st Century BCE



Smoking, Physical Activity, and Active Life Expectancy

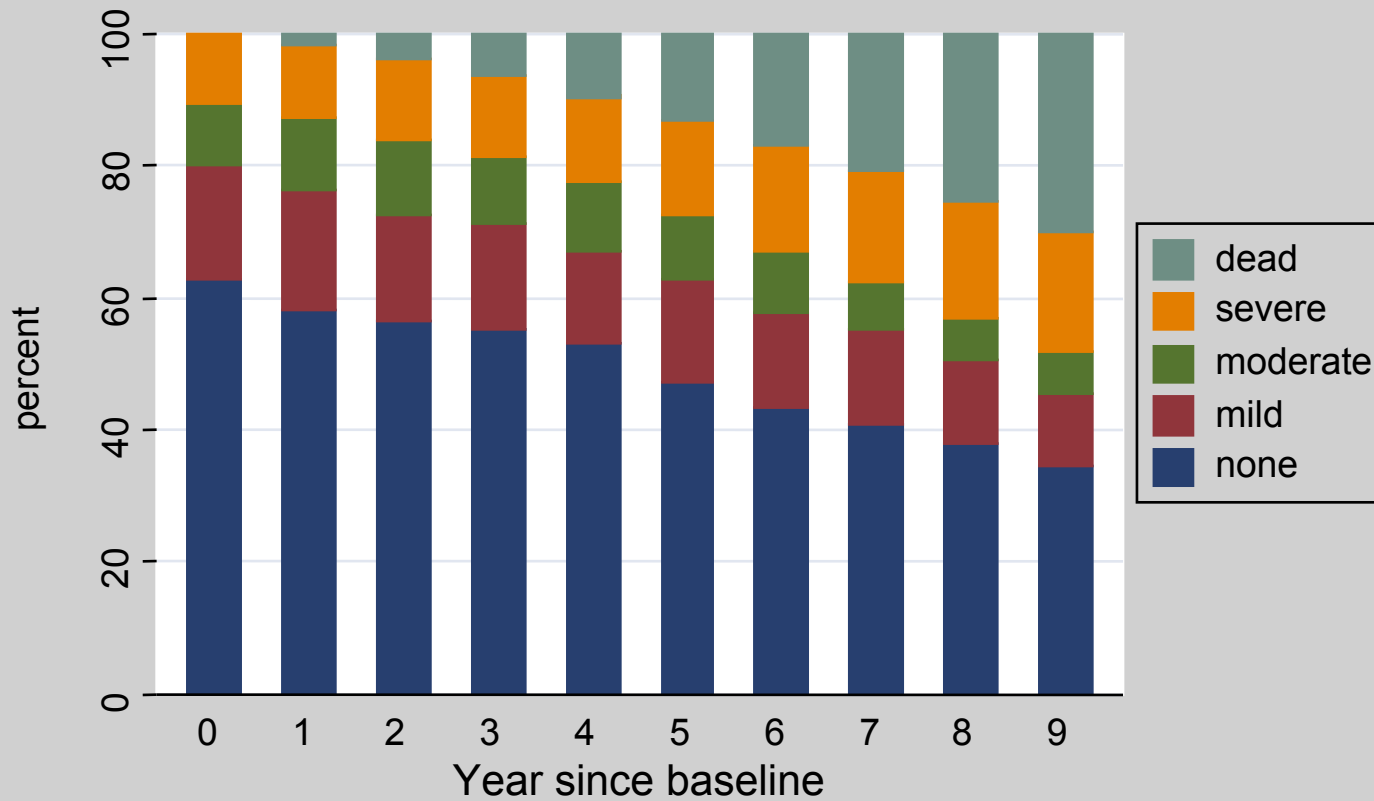
Luigi Ferrucci,^{1,2} Grant Izmirlian,² Suzanne Leveille,² Caroline L. Phillips,² Maria-Chiara Corti,³ Dwight B. Brock,² and Jack M. Guralnik²

1. Higher physical activity positively impact ALE more than DLE
2. Smoking has the highest impact on longevity
3. However, people who never smoked and were sedentary have the same longevity but shorter ALE than those who smoked and were physically active
4. These findings are consistent in men and women and according to age group
5. Women live longer but have only slightly longer ALE than men.



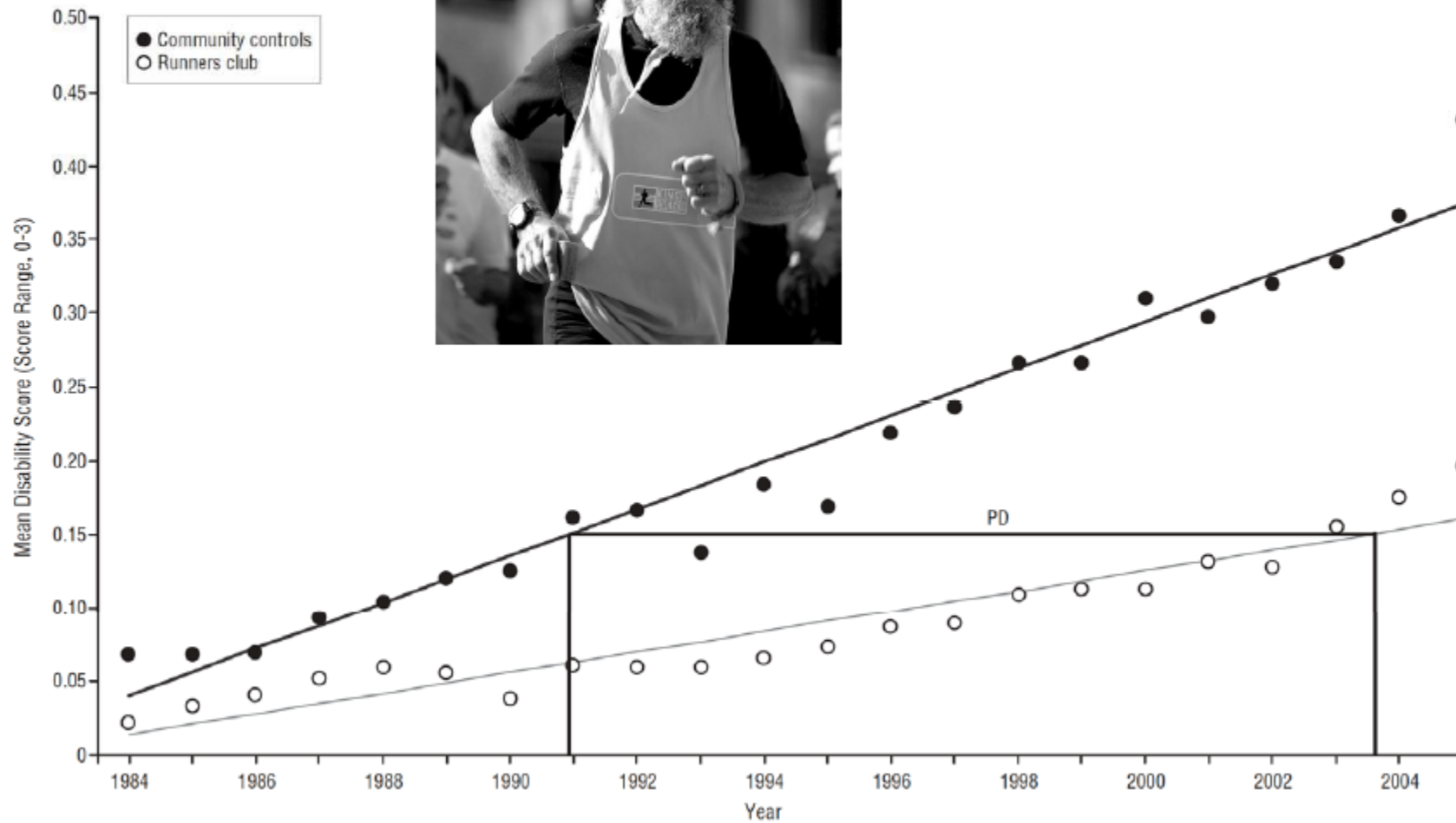
Trends in the overall CHS cohort

Difficulty in physical function: high energy tasks
In full CHS cohort

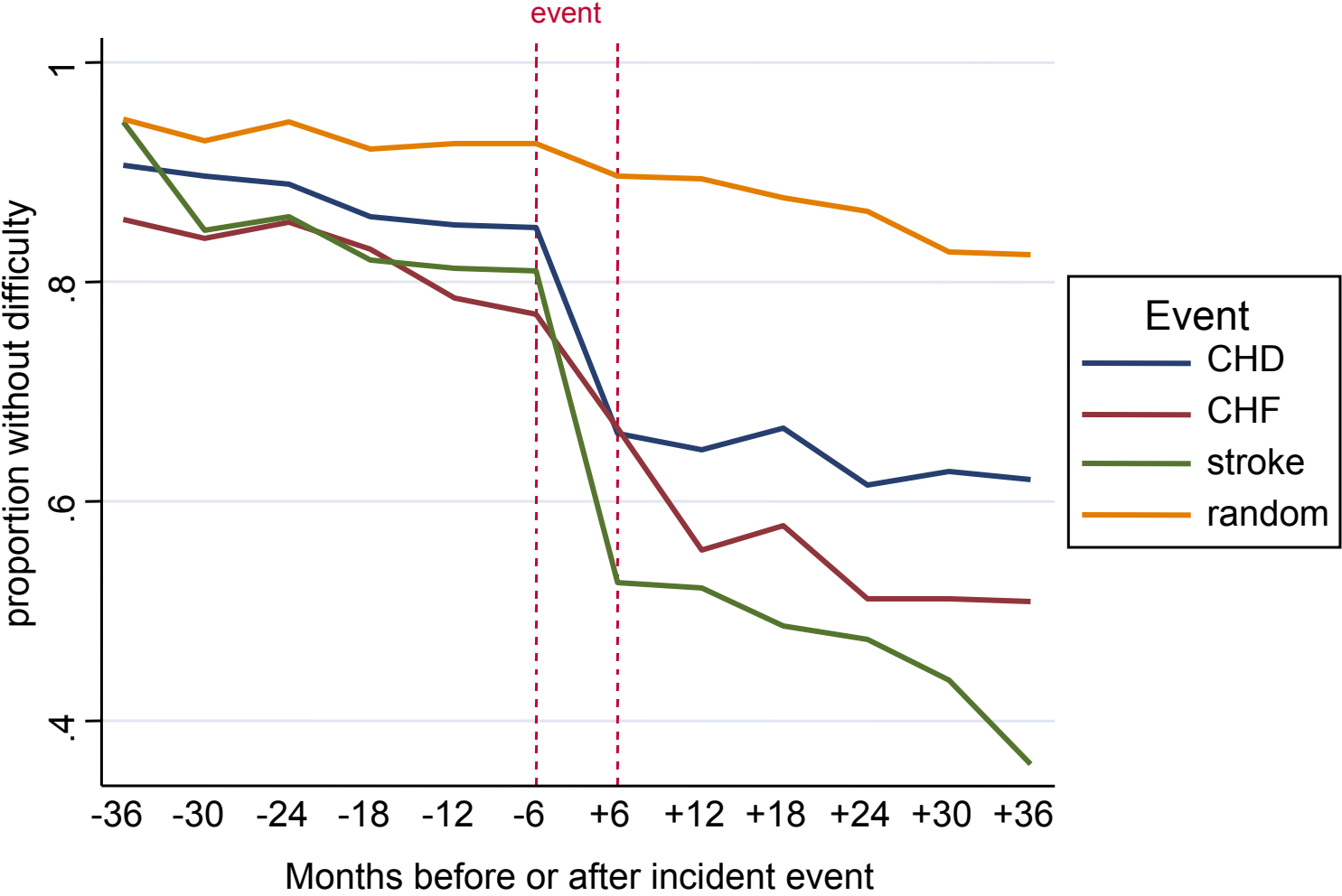


Reduced Disability and Mortality Among Aging Runners: A 21-Year Longitudinal Study

Eliza F. Chakravarty, MD, MS; Helen B. Hubert, PhD; Vijaya B. Lingala, PhD; James F. Fries, MD



Self-reported physical function: medium energy tasks



Compression of Morbidity:

Major public health goal for an aging society

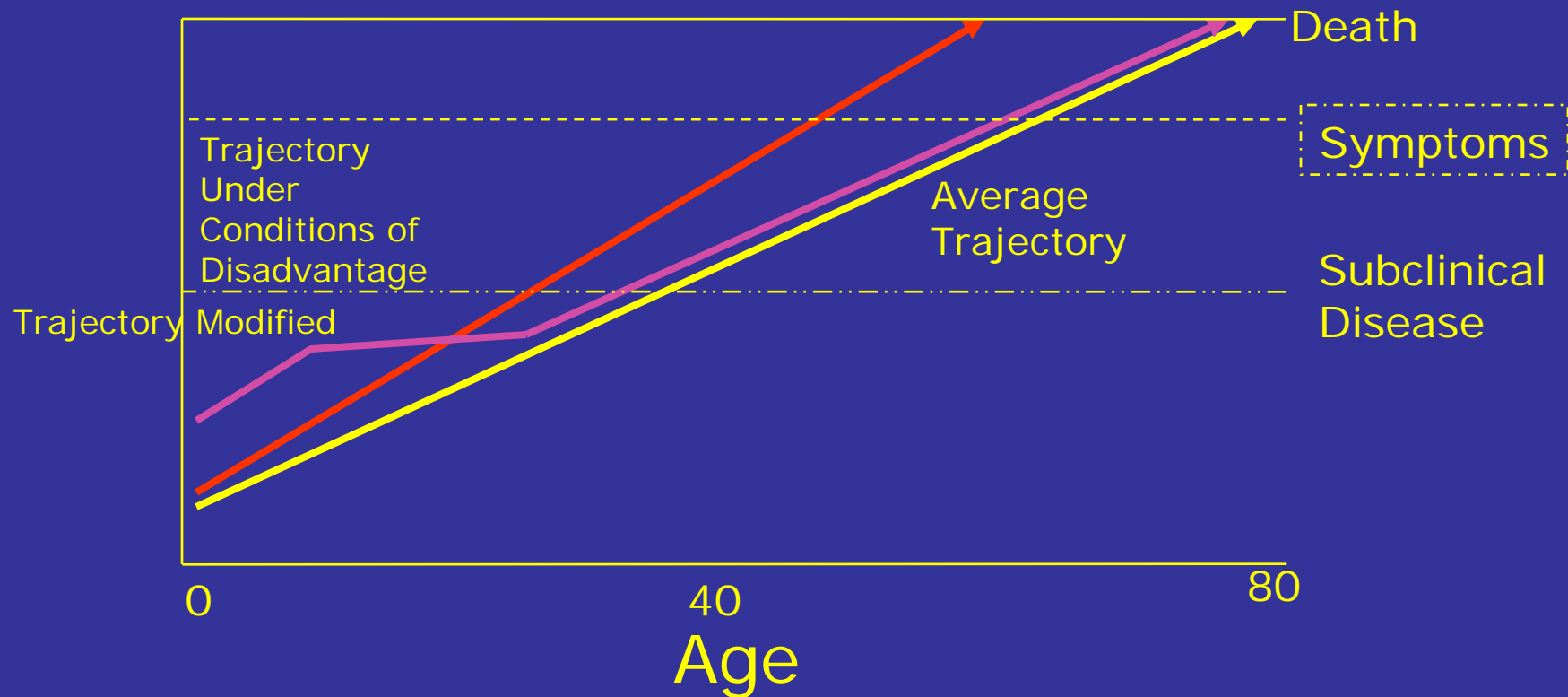
- Push back development of disease and disability to latest points in human life span
- Add "life to years"

Life Course Across the Lecture



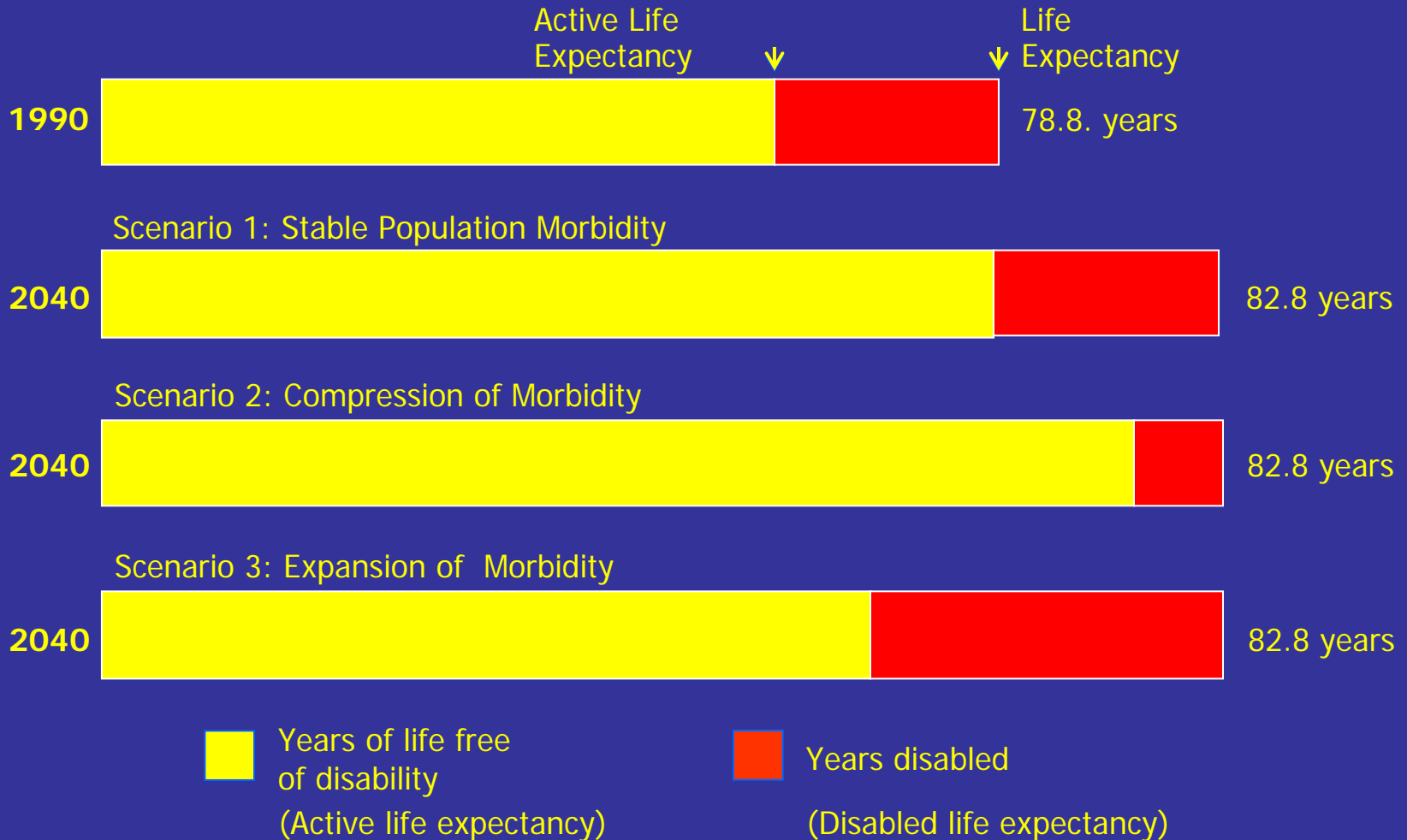
- Frederica Perera
 - *in utero* environmental exposures and effects in childhood and beyond
- Ezra Susser
 - latent and transgenerational effects
- Bruce Link
 - disparities and disease risk
- Linda Fried

Chronic Diseases Emerge Over the Life Course



Health Biography of Older Age

Burden of Illness/Disability from 1990 to 2040



Compression of Morbidity: Major public health goal for an aging society

- Push back development of disease and disability to latest points in human life span
- Add “life to years”

Potential for Prevention and Compression of Morbidity

Impact of Effective Prevention: Change in U.S. death rates, 1950-1997

	Age Group	% change
Heart Disease	65-74	-58%
	75-84	-54%
	85+	-31%
Stroke	65-74	-76%
	75-84	-65%
	85+	-46% <i>Bureau of the Census</i>

Approaches Demonstrated: Prevention or Treatment of Chronic Conditions

	Med Tx/Prev	Exercise
Arthritis	+	+
High Blood Pressure	+	+
Osteoporosis	+	+
Diabetes	+	+
Heart Attack	+	+
Angina	+	+
Cancer	+	
Lung Disease	+	
Stroke	+	+

(moderate/vigorous) and 5-year mortality,

Men and Women 65-101 years of age,

Kcals/week:	Death rate/1000 person-years	RR	p
≤ 67.5	52.1	1.00	
67.5-472.5	32.8	0.78	(*)
472.5-980	28.7	0.81	(*)
980-1890	22.9	0.72	*
>1890	15.5	0.56	*

Fried. *JAMA*, 1998

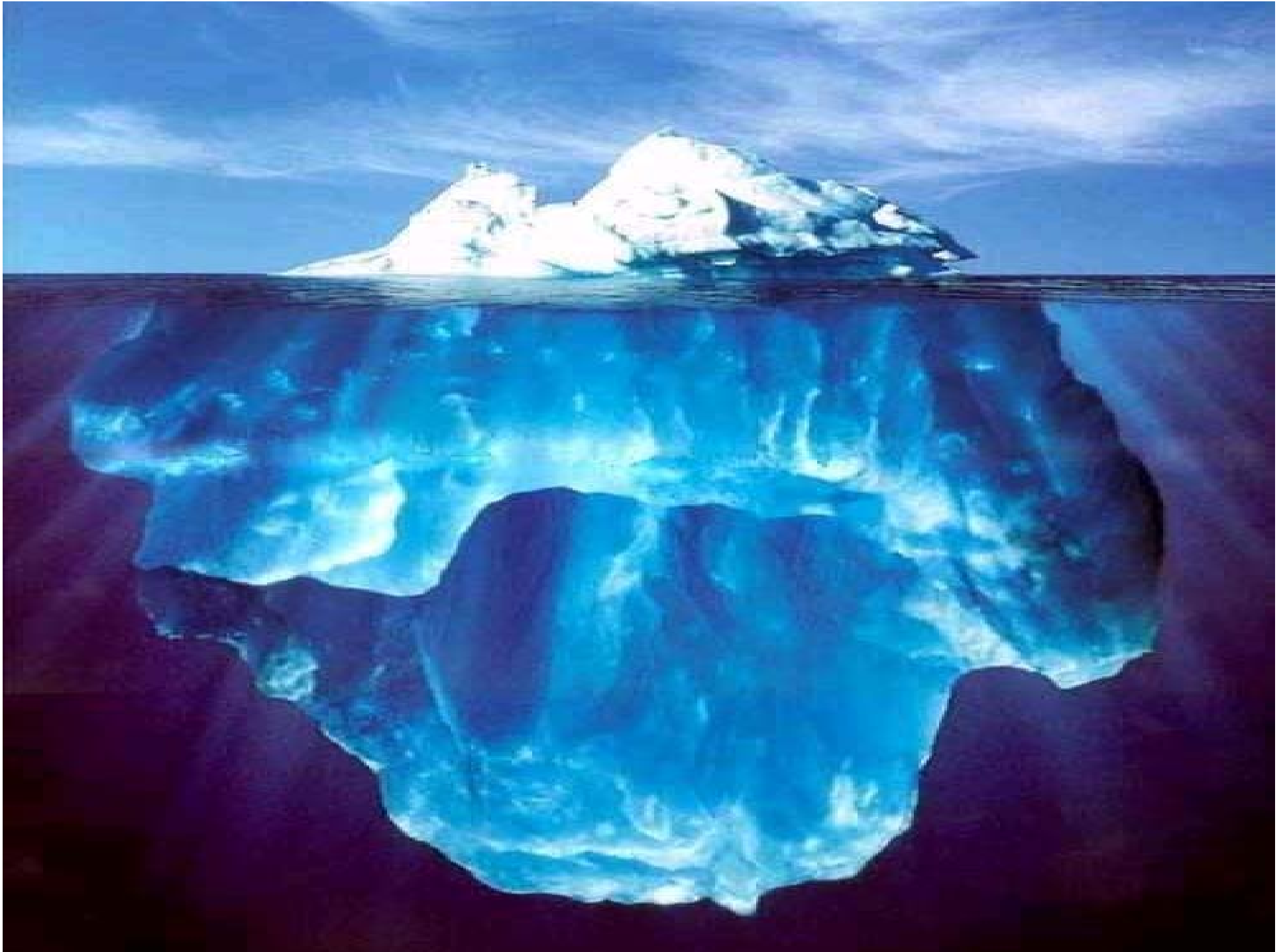
Erikson's 8 Stages of Psycho-Social Development

Trust Vs Mistrust	Autonomy Vs Shame & Doubt	Initiative Vs Guilt	Industry Vs Inferiority	Identity Vs Role Confusion	Intimacy Vs Isolation	-----Generativity----- Ego ----- Integrity Vs Despair	Stagnation
Stage 1	Stage 2	Stage 3	Stage 4	Stage 5	Stage 6	Stage 7	Stage 8
Infancy 0 to 18 Months	Toddlerhood 18 months to 3 years	Early Childhood 3 to 5 years	Middle Childhood Elementary 5 to 12 Years	Adolescence 12 to 18 years	Young Adult 20s +	Middle Adulthood 25 to 65 years	Older Adulthood 60 years+

Life course



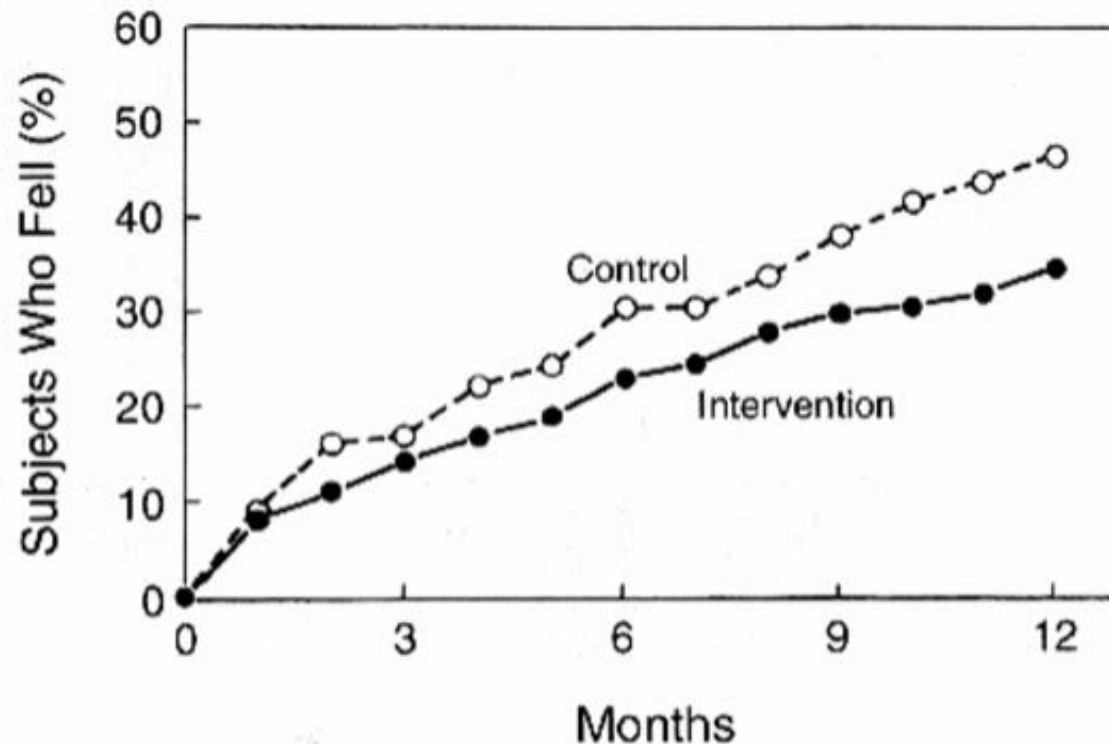
- **Goal: Healthy aging**
- **Opportunities for optimizing health and wellbeing?**
- **When to intervene?**



Times of particular risk, of development and of malleability

- Perinatal
- Infancy
- Childhood
- Adolescence
- Young adulthood
- Middle age
- Young-old
- Old-old

Falls prevention through multirisk factor intervention leads to 30% decline in falls, 1 Year of Follow up



Intervention	153	130	113	103	95
Control	148	123	102	89	76
Relative risk	—	0.86	0.77	0.79	0.75

Tinetti M et al. *N Engl J Med* 1994

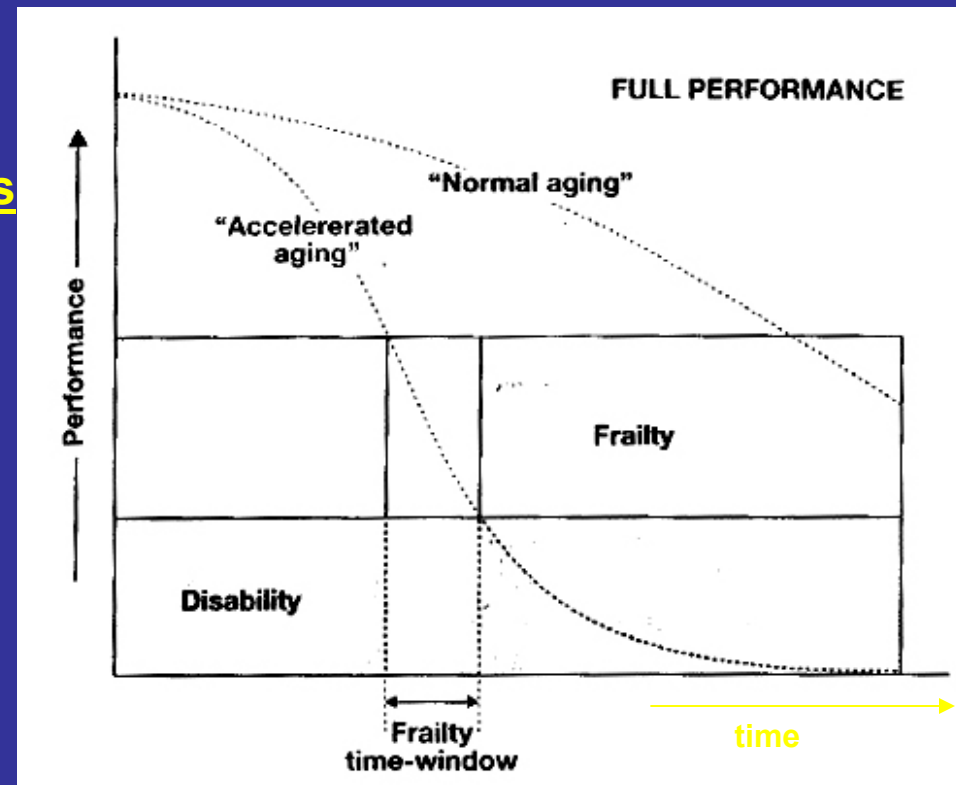
Frailty: A Brief Overview

Definition

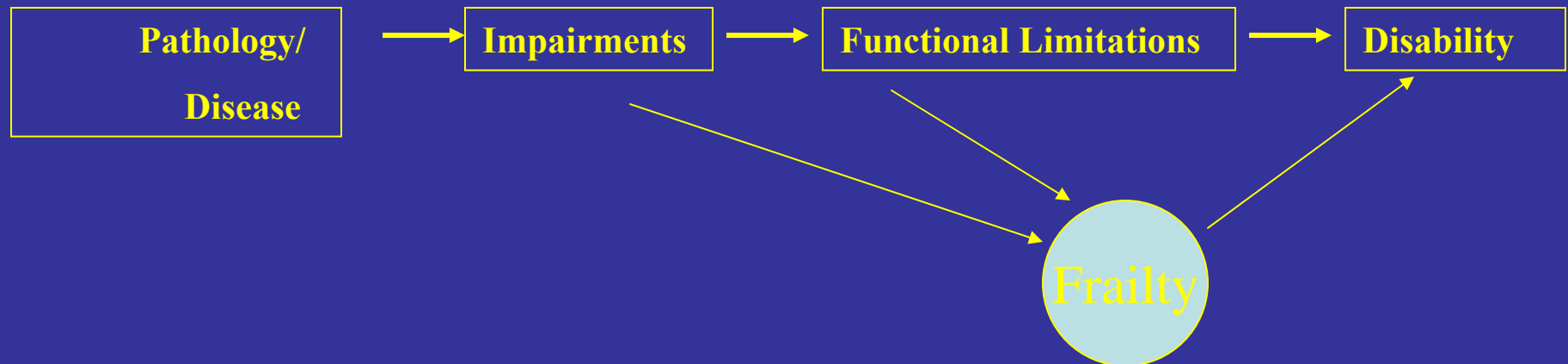
“A central definition of frailty in geriatric medicine is that it is a clinical state of vulnerability to stressors, [...] resulting from aging-associated declines in resiliency and physiologic reserves and a progressive decline in the ability to maintain a stable homeostasis.” [1]

Frailty is a predictor of poor outcomes

- Falls
- Hospitalization
- Institutionalization
- Disability
- Mortality [2]



WHO/IOM Pathway to Disability



WHAT older adults *DO* affects their health...

- Activity and engagement
 - Physical
 - Social
 - Networks, support
 - Engagement;
 - Regular structured activities
 - Cognitive

A Social Model of Health Promotion for an Aging Society: Experience Corps

- High intensity volunteering for older adults
- High impact roles in public elementary schools improving outcomes for children
- Critical mass of older adults:
 - Shift outcomes for schools
 - Force for social benefit
 - Social networks and friendships
- Health promotion program embedded

» Fried et al, 2004

Aging issues in HIV/AIDS

- Frailty
- Multimorbidity
- Polypharmacy
- Disability

HIV and Aging

- **Similarities between HIV and aging at the biological level**
 - T-lymphopenia, decreased cellular immunity
 - Replicative senescence of T-lymphocytes
 - ↑ pro-inflammatory markers (IL-6, TNF- α , IFN- γ)
- **Similarities between HIV and aging at the clinical level**
 - Sarcopenia, weight loss, wasting
 - Cognitive disorders, dementia
 - Rheumatologic disorders, decrease in bone mineral density
 - Frailty-like clinical presentation presaging death

Definition of a frailty-related phenotype (FRP)

- The FRP definition was based on the frailty phenotype of Fried et al.
- Components of the frailty phenotype:

→	→	→	→
---	---	---	---

▪ FRP in the MACS
- Physical shrinking (unintentional weight loss) → - available
- Weakness (grip strength) → - not available *
- Exhaustion (self-reported) → - available
- Slowness (time to walk 15 feet) → - approximated (SF-36) *
- Low physical activity level (a weighted score of kilocalories/week) → - approximated (SF-36)

Exhaustion: During the past 4 weeks, as a result of your physical health, have you had difficulty performing your work or other activities (for example, it took extra effort)?

Slowness: Does your health now limit you in walking several blocks?

Low physical activity: Does your health now limit you in vigorous activities, such as running, lifting heavy objects, participating in strenuous sports?

* Grip strength and time to walk have recently been incorporated in the MACS

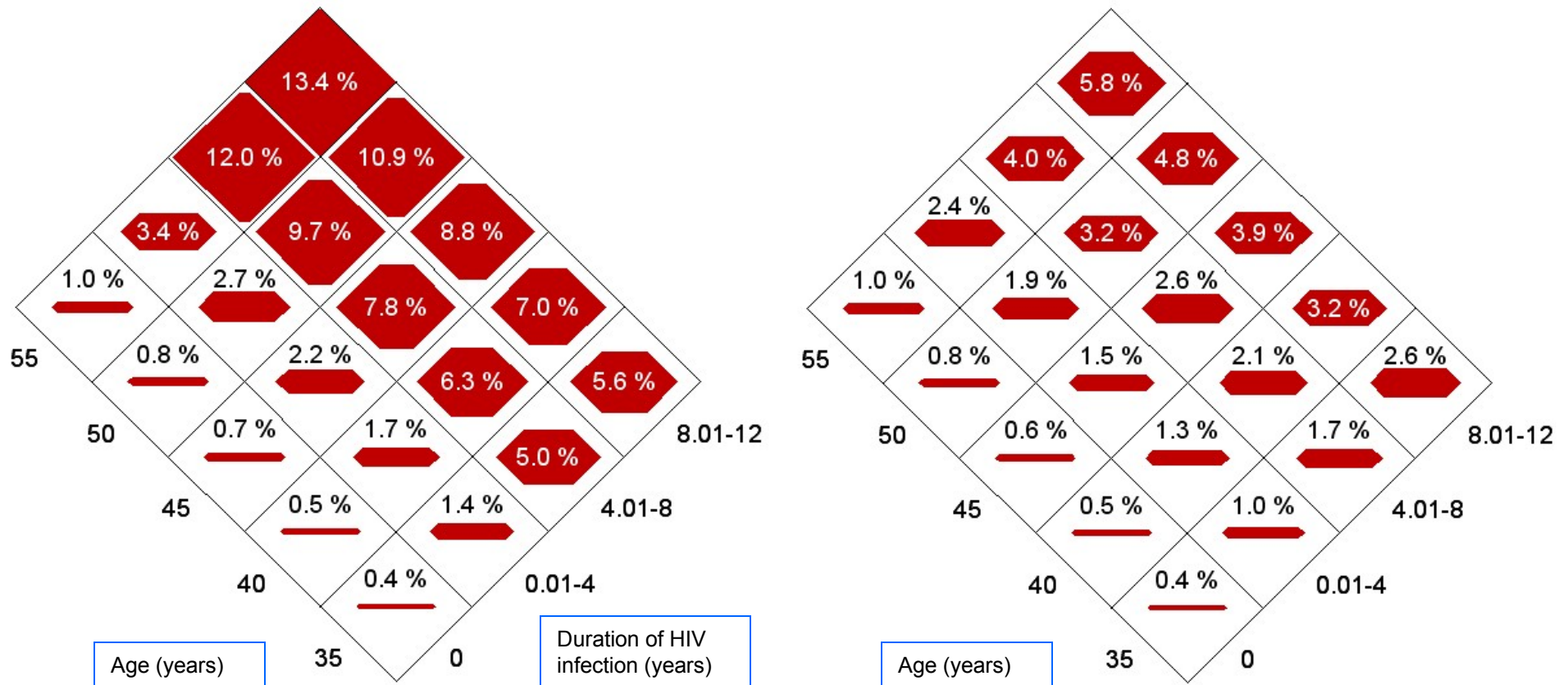


Figure 2. Adjusted prevalence of having a frailty-related phenotype according to age and presence and duration of HIV infection, for fixed values of ethnicity (white non-Hispanic) and educational level (college), among HIV-seronegative and -seroconverted men in the Multicenter AIDS Cohort Study (MACS) between April 1994 and January 1, 1996 when (a) all person-visits were included and (b) excluding person-visits occurring later than 6 months before the first AIDS-defining illness. The ratio of the areas (and middle vertical length) of any two shaded polygons equals the ratio of the two corresponding prevalences written in their centers. For example, the area corresponding to a prevalence of 1.7% is equal in a and b, and is the equivalent of half the area corresponding to a prevalence of 3.4%.

Summary

- HIV-1 infection was associated with a >10-year earlier occurrence of a phenotype related to frailty (FRP) [1]
- Non-linear association between CD4 cell count and FRP
 - Risk of FRP increased with decreasing CD4 cell count, especially when CD4 cell count < $\sim 400/\text{mm}^3$ [2]
- After adjusting for ages and CD4 cell count, FRP prevalence decreased after the introduction of HAART, but has not further diminished with the establishment of HAART [2]
- Older age, lower educational level, and clinical AIDS were independently associated with FRP among HIV+ men [1]
- Proportion of visits with FRP prior to HAART initiation independently predicted the subsequent risk of AIDS or death, even after HIV suppression
- True Frailty Phenotype is under investigation.

[1] Desquilbet L et al, J. Gerontol_A Biol. Sci. Med. Sci. 62:1279-1286, 2007.

[2] Desquilbet L et al, J. Acquir. Immune Def. Syndr. 50:299-306, 2009 .

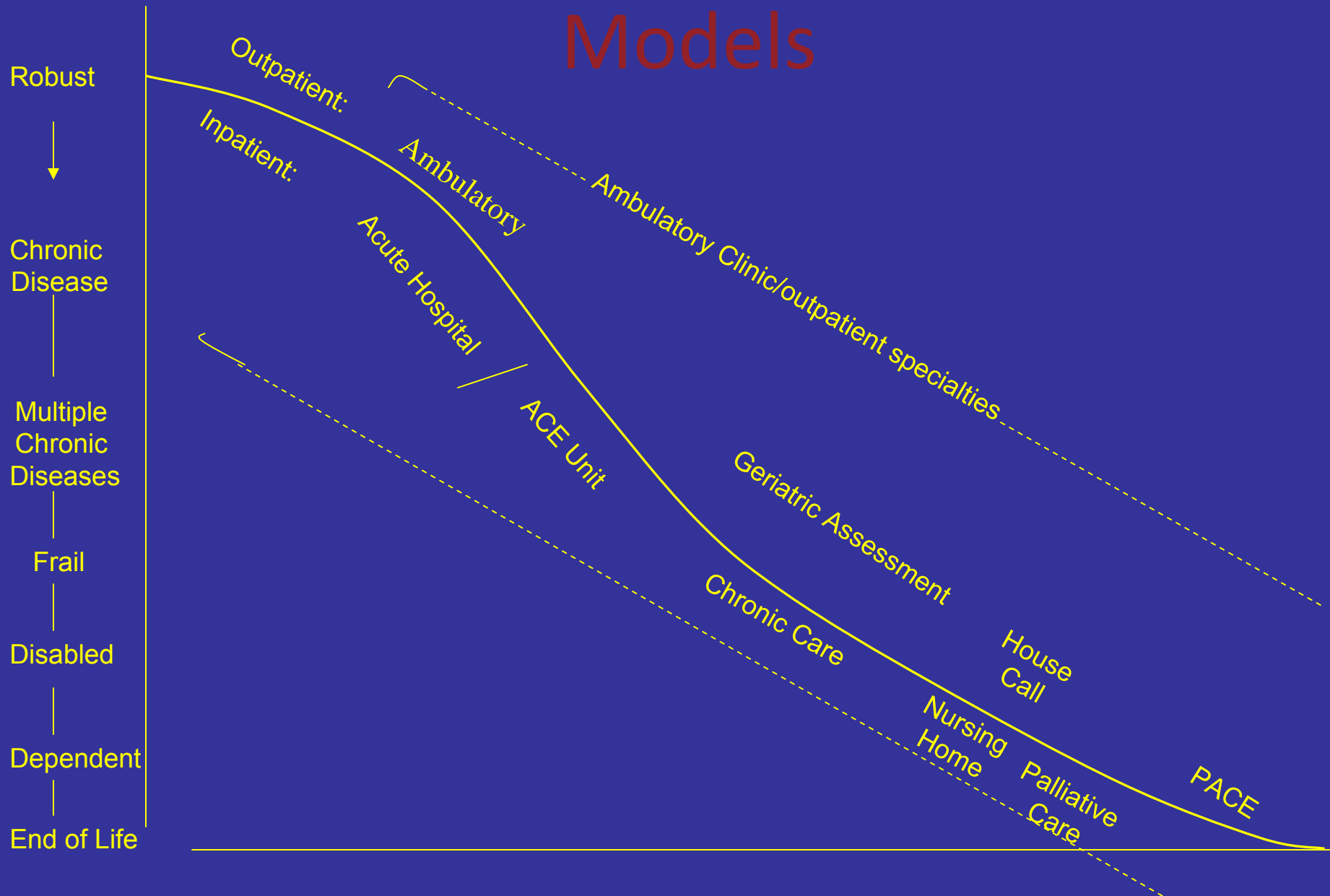
Completing the epidemiologic transition: preventing chronic disease - associated with aging

Transitioning to a chronic disease world

Chronic, progressive
Multifactorial etiology
Multilevel solutions

Both public health and medical care

Continuum of Geriatric Care Models



Challenge: The Public Health System Not Ready for Aging

- Evidence on primary, secondary and tertiary prevention for older adults not implemented through public health
- 20% of older adults in community isolated, without access to care; ineffective links to clinical care provision
- Responsibility for public and community health for an aging population not allocated

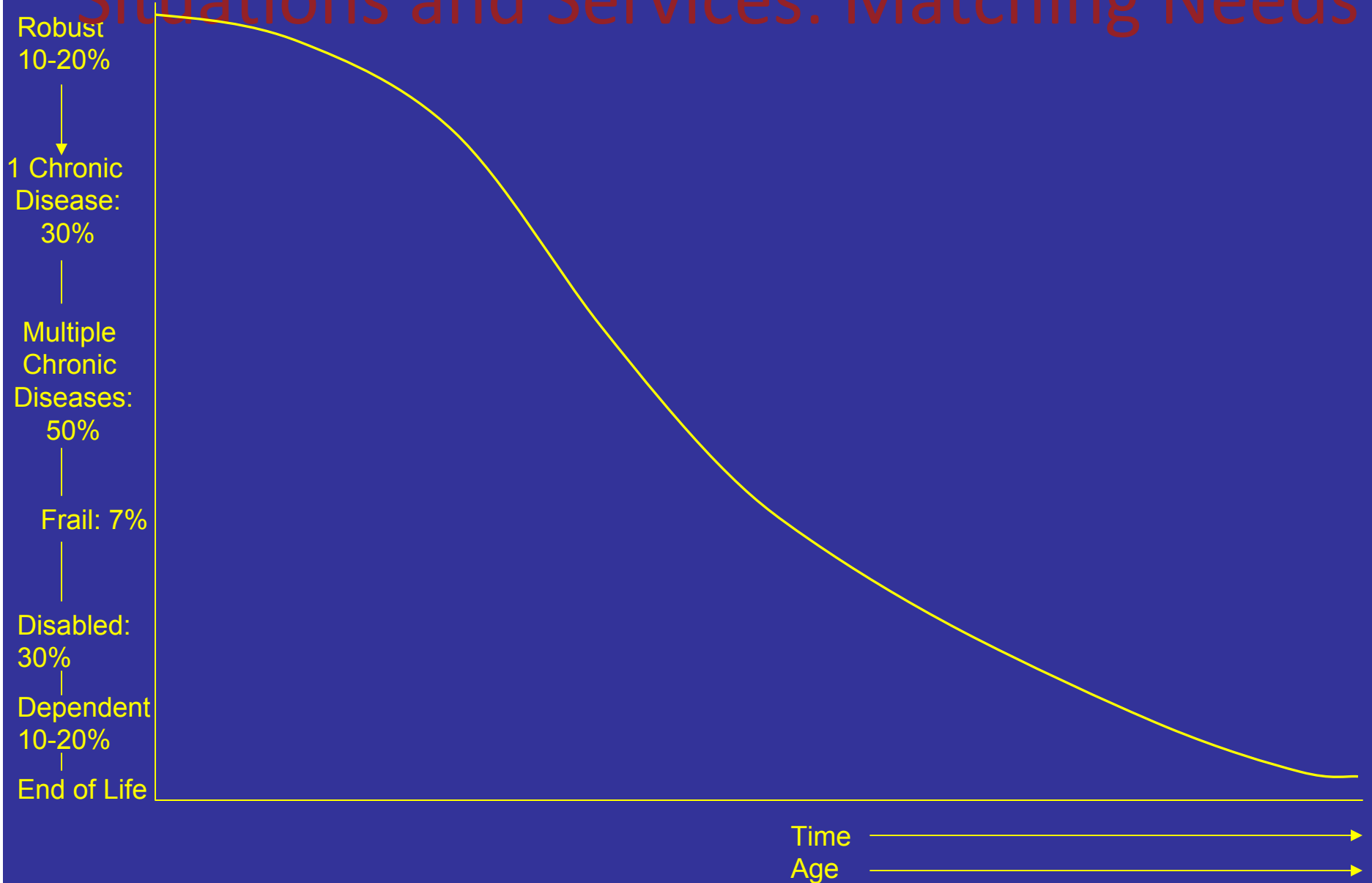
Challenge: Public Health could lead by designing an integrated health system – that invests in prevention

Public health and medical care in one coordinated system

Building Blocks for Effective Health Promotion for an Aging Population beyond Disease Focus

Access to Care	Widowhood	Financial Resources	Environmental Stressors
Polypharmacy	Disability	Health Disparities	
Frailty	Multimorbidity	Falls	Depression/ Loneliness
Primary, Secondary Prevention; Disease Management			

Community-based Continuum of Living Situations and Services: Matching Needs



Age-Friendly City: Key Domains

- Housing
- Transportation
- **Outdoor spaces** and buildings
- Social participation
- **Civic engagement** and employment
- Respect and social inclusion; address ageism, fear
- Communication and information re: services
- Community support and health services

WHAT we DO Matters with aging: Improving Health Outcomes

- Physical Activity
- Cognitive Activity
- Social engagement:
 - Social networks and support; loneliness
 - Meaningful, productive roles
 - Leaving a legacy

Population based studies. Lack of activity and engagement adversely affect health outcomes for older adults.

- Social isolation
- Loneliness
- Non-stimulating environments
- Sedentariness
- Little sense of purpose

Few meaningful roles for older adults

- At same time, society needs the contributions of older adults:
 - Only increasing natural resource
 - Best educated cohort of older adults in history of world
 - Time, experience, wisdom
 - Wanting to leave the world a better place

Ho: “generativity” is key to successful aging

- Leaving a legacy; Leaving the world better for future generations (Erikson)
 - Productive, meaningful engagement

What if we could create new,
meaningful, generative ways for
people to stay engaged after
retirement - and design them to
promote health?

A win-win: target roles to societal unmet
needs

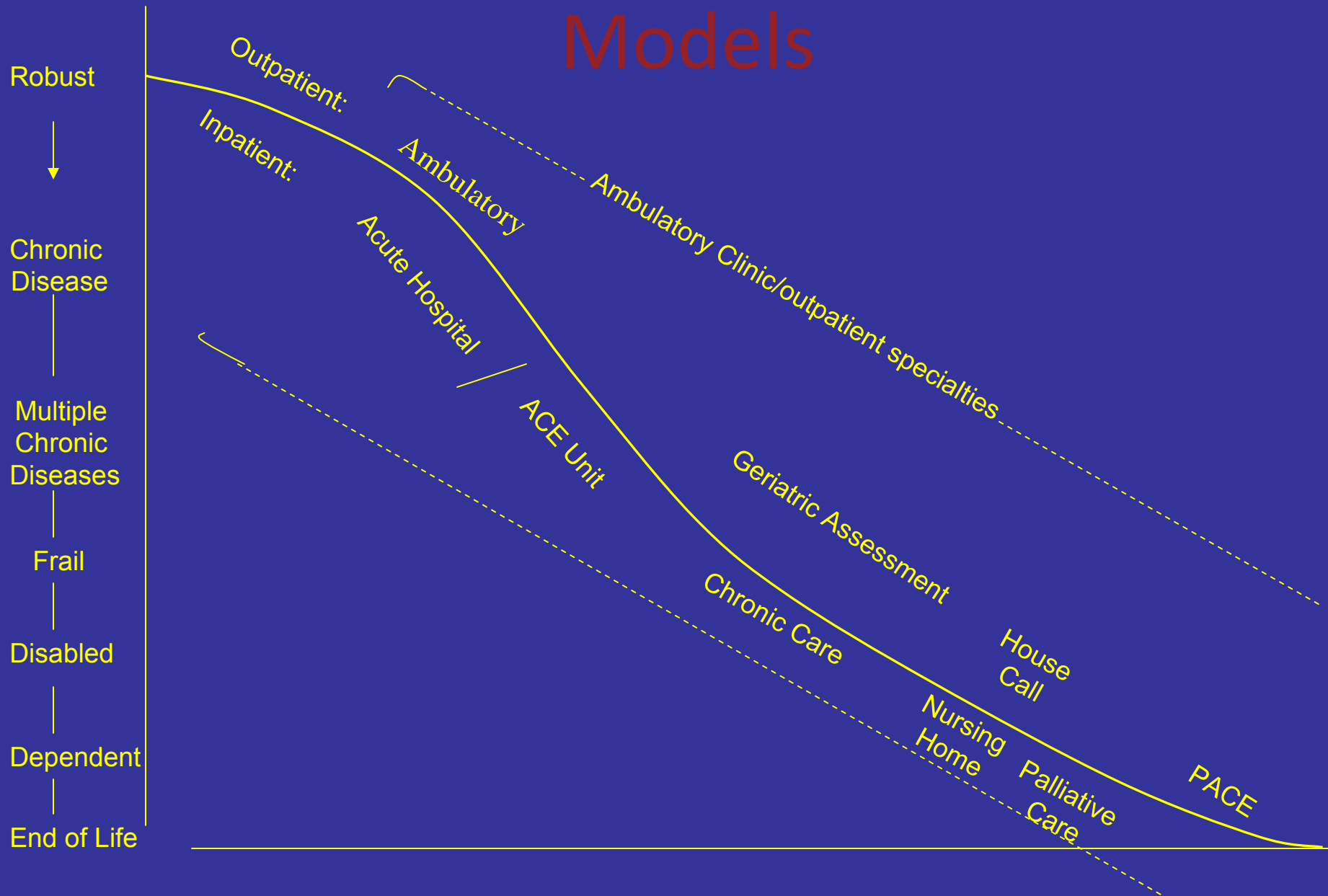
Public health for an aging society will benefit all of us

- Systems for health designed to prevent and ameliorate chronic conditions
- Healthy older adults; successful aging
- Healthy communities
- Healthy cities
- Institutions that support continued engagement and productivity with aging
- Strengthening win-wins across generations
- Who we are as a society

Challenge: The Health Care System Not Ready for Chronic Care Needs of Older Adults

- Acute event-based care in a chronic disease world
- Effective geriatric care not implemented or reimbursed
 - Community-based coordinated continuum of care needed
- Prevention needs to be inserted into medical care and linked to community-based approaches
 - *Institute of Medicine. Retooling for an Aging America: Building the Health Care Workforce, 2008*

Continuum of Geriatric Care Models



Challenge: The Public Health System Not Ready for Aging

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Primary, Secondary Prevention; Disease Management			





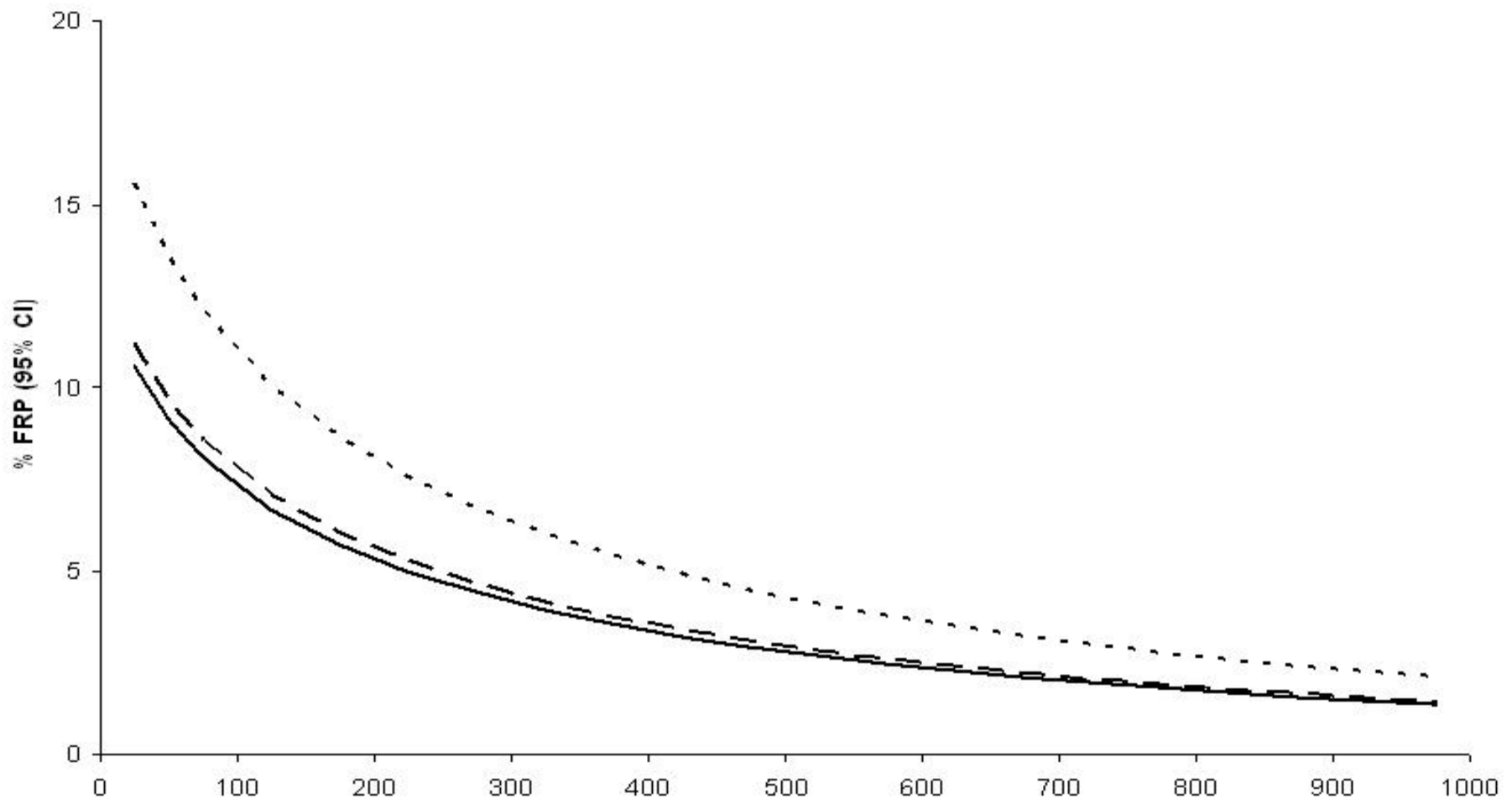


FIGURE 1. Estimated prevalences of a FRP as a function of CD4 T-cell count in the MACS for fixed values of age (45 years), fixed percentage for ethnicity (80% white non-hispanic), education (52% \geq college), and prevalence of AIDS (20%), in the pre-HAART era (1994-1995; dotted line), introduction of HAART era (1996-1999; dashed line), and established HAART era (2000-2005; plain line). The curves for the 2 HAART eras do not differ significantly from each other, but both are significantly different from that of the pre-HAART era (see text for odds ratios and P values comparing the 3 different eras).

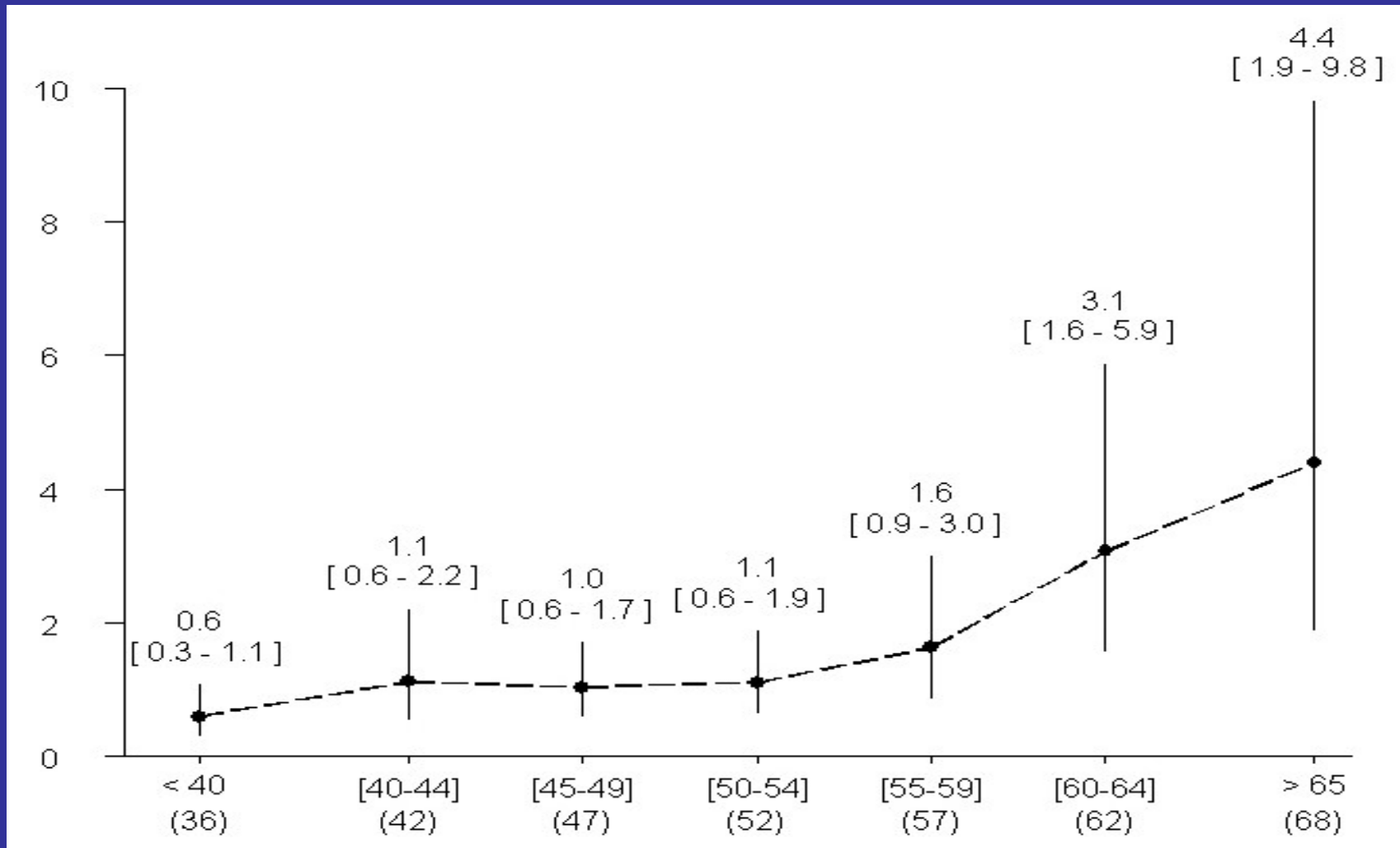


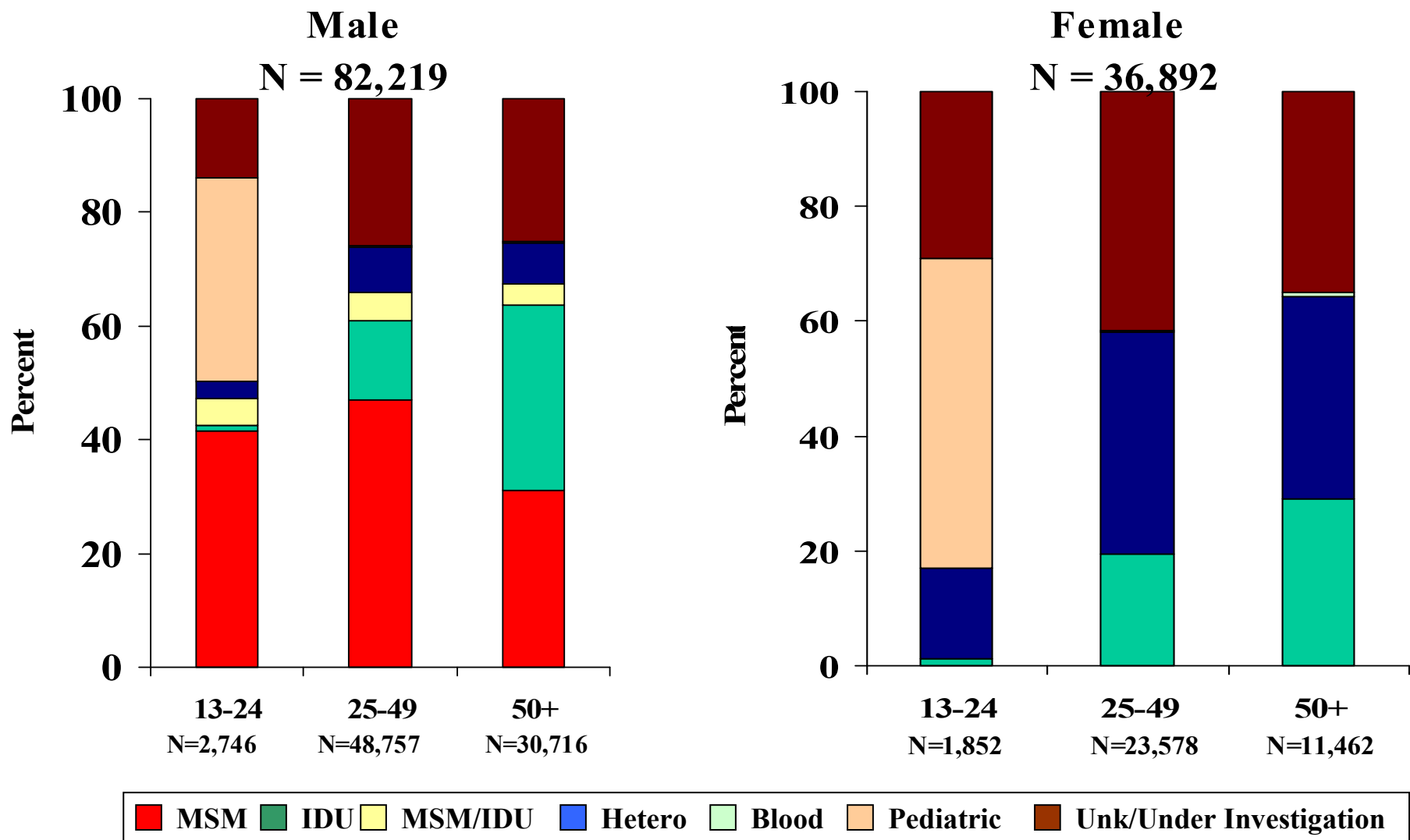
Figure 1. Estimated prevalence of a frailty-related phenotype (FRP) as a function of age among 1977 HIV-seronegative individuals from the Multicenter AIDS Cohort Study (MACS) cohort (April 1994–November 2004) (95% confidence interval [CI])

Area Profile
New York State

2007

New York State Department of Health
Bureau of HIV/AIDS Epidemiology

Persons* Ages 13 and Older, Living with HIV/AIDS by Gender, Risk and Current Age New York State



* Includes prisoners.

Persons Living with HIV/AIDS and Cumulative AIDS Cases*

New York State

Region of Residence at Time of Diagnosis	Living with HIV (non AIDS)	Living with AIDS	Cumulative AIDS Cases
New York City	35,640	57,029	144,887
Rest of State	10,400	16,860	35,787
New York State	46,040	73,889	180,674

*Includes prisoners.

Estimated numbers of persons living with HIV/AIDS, by year and selected characteristics, 2004–2007—34 states and 5 U.S. dependent areas with confidential name-based HIV infection reporting (Numbers derived from Table 9)

Age	2004	2005	2006	2007
50–54	53,903	60,030	67,082	74,582
55-59	28,077	33,023	38,186	43,985
60-64	13,363	15,309	17,705	20,962
≥65	<u>10,512</u>	<u>12,361</u>	<u>14,363</u>	<u>16,982</u>
	105855	120,723	137,336	156,511

Note. These numbers do not represent reported case counts. Rather, these numbers are point estimates, which result from adjustments of reported case counts. The reported case counts have been adjusted for reporting delays and missing risk-factor information, but not for incomplete reporting.

Centers for Disease Control and Prevention. *HIV/AIDS Surveillance Report, 2007*. Vol. 19. Atlanta: U.S. Department of Health and Human Services, Centers for Disease Control and Prevention; 2009:[inclusive page numbers]. <http://www.cdc.gov/hiv/topics/surveillance/resources/reports/>.

Estimated numbers of persons living with AIDS, by year and selected characteristics, 2003-2007 - United States (50) and the District of Columbia (numbers derived from Table 12)

Age	2003	2004	2005	2006	2007
50-54	46,661	53,125	59,045	66,003	72,991
55-59	23,976	28,149	33,279	38,626	44,298
60-64	11,224	13,232	15,265	17,878	21,196
≥65	<u>8,842</u>	<u>10,450</u>	<u>12,232</u>	<u>14,386</u>	<u>17,005</u>
	90,703	104,956	119,821	136,893	155,490

Note. These numbers do not represent reported case counts. Rather, these numbers are point estimates, which result from adjustments of reported case counts. The reported case counts have been adjusted for reporting delays and missing risk-factor information, but not for incomplete reporting.

Centers for Disease Control and Prevention. *HIV/AIDS Surveillance Report, 2007*. Vol. 19. Atlanta: U.S. Department of Health and Human Services, Centers for Disease Control and Prevention; 2009:[inclusive page numbers]. <http://www.cdc.gov/hiv/topics/surveillance/resources/reports/>.

Frailty and HIV in the Multicenter AIDS Cohort Study (MACS)

Joseph B. Margolick, L. Desquilbet, L. P. Fried, L. P. Jacobson

Studies of a Frailty-Related Phenotype in the MACS

- ▶ 4954 MSM followed semiannually since 1984
- ▶ The FRP was present if ≥ 3 of the above 4 components were answered “yes” (#1 and #2) or “yes, limited a lot” (#3 and #4)
- ▶ Covariates: Age, Education, Ethnicity, CD4 cell count, HIV RNA
- ▶ Study population
 - MACS individuals enrolled before 1996
 - Seroconverter and seroprevalent men
 - ≥ 1 measurement of CD4 cell count between visit 21 and visit 41
- ▶ Visits:
 - All HIV+ visits between visit 21 and visit 41
- ▶ Final study population: N = 1045 (N person-visits = 12,916)
 - 98 men had no measurement of CD4 count

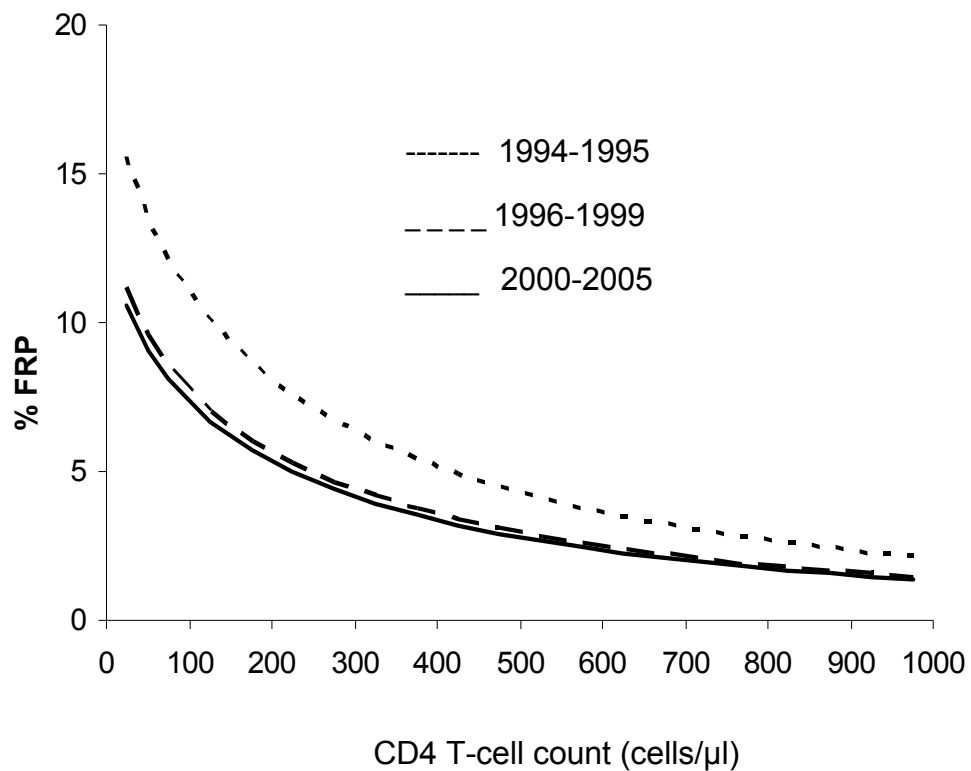
Number of person-visits (1994-2004)

N_{total} = 12,916

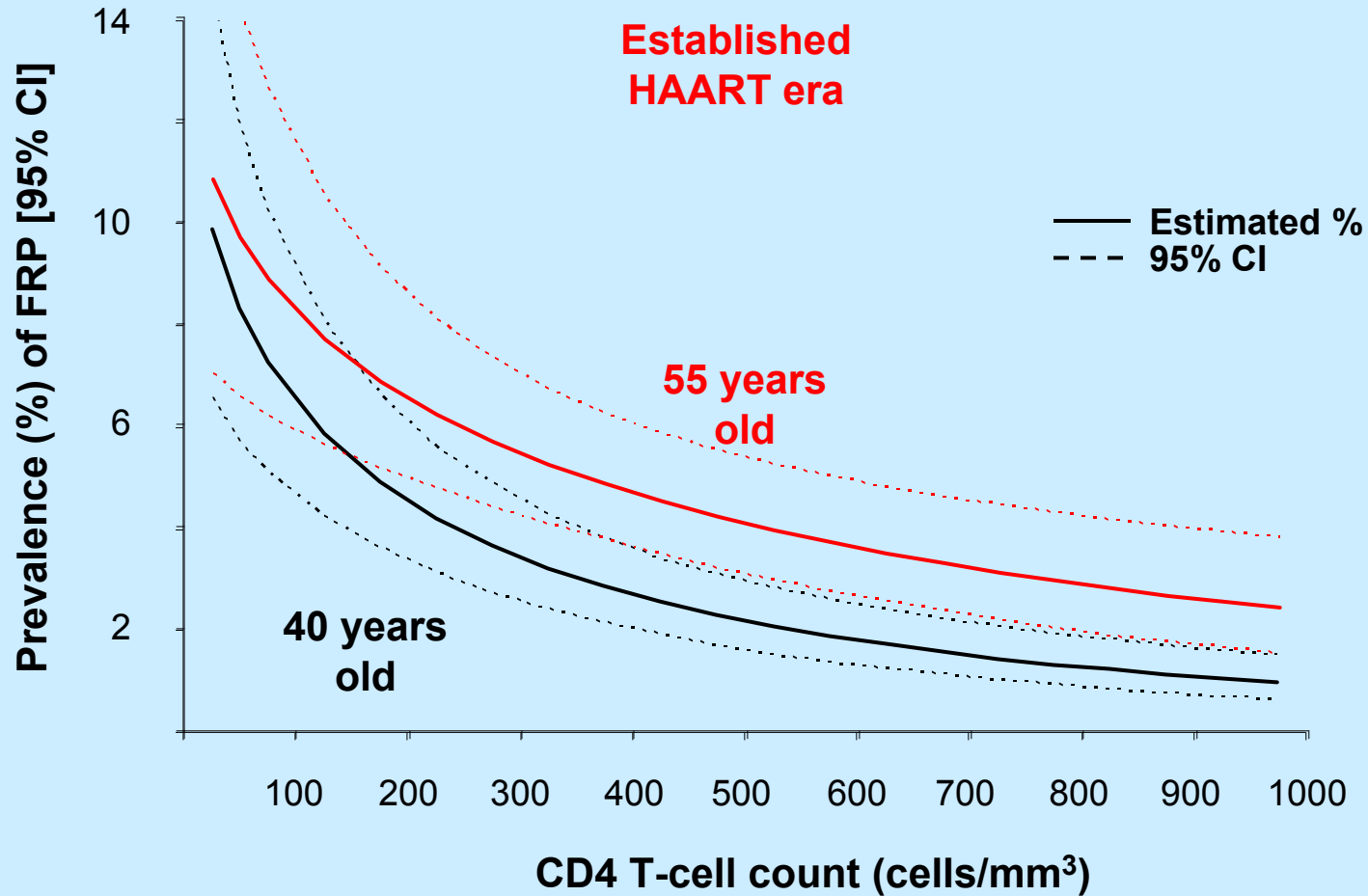
Variables	< 1996 (N=2,660)	1996-1999 (N=5,104)	2000-2004 (N=5,152)
Visits with FRP % (N)	7.6% (184)	5.3% (247)	4.4 (198)
Current age*	41 (37-46)	44 (39-48)	48 (44-52)
Current CD4 cell count*	320 (133-500)	415 (262-609)	489 (324-696)
Current CD4:CD8 ratio*	0.31 (0.15-0.51)	0.42 (0.26-0.63)	0.51 (0.33-0.75)

* Median (IQR)

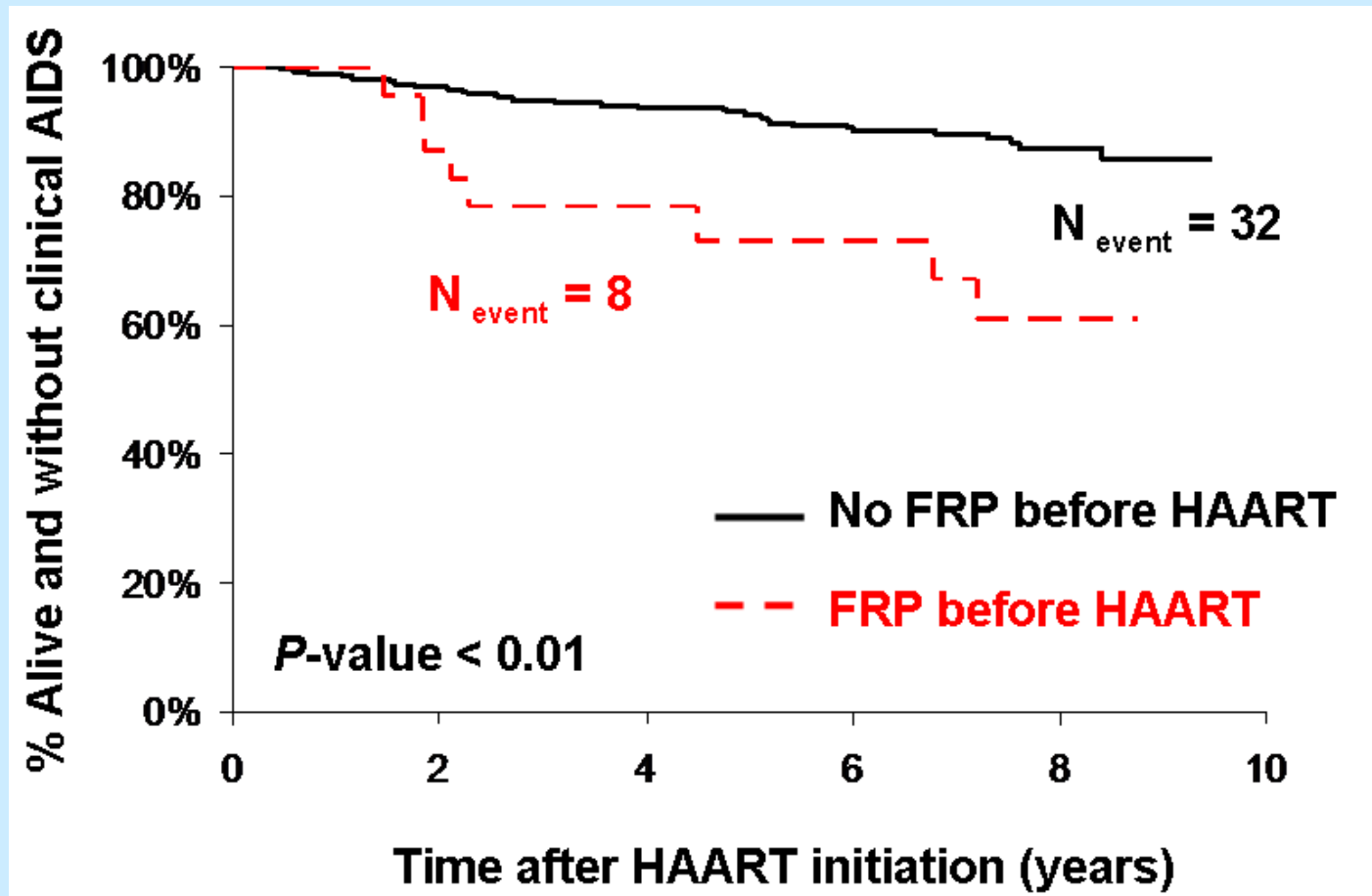
Relationship between CD4 T-cell count and Prevalence of Frailty-Related Phenotype, by Calendar Period



Effect of Age Decreases for Low CD4 T-Cell Counts



Time to AIDS or Death After Starting HAART, by Presence or Absence of a Frailty-Related Phenotype



Acknowledgements

▶ **Multicenter AIDS Cohort Study (MACS)**

- J. Phair (Chicago)
- R. Detels, B. Jamieson (Los Angeles)
- L. Jacobson (Baltimore)
- C. Rinaldo, M. Holloway (Pittsburgh)

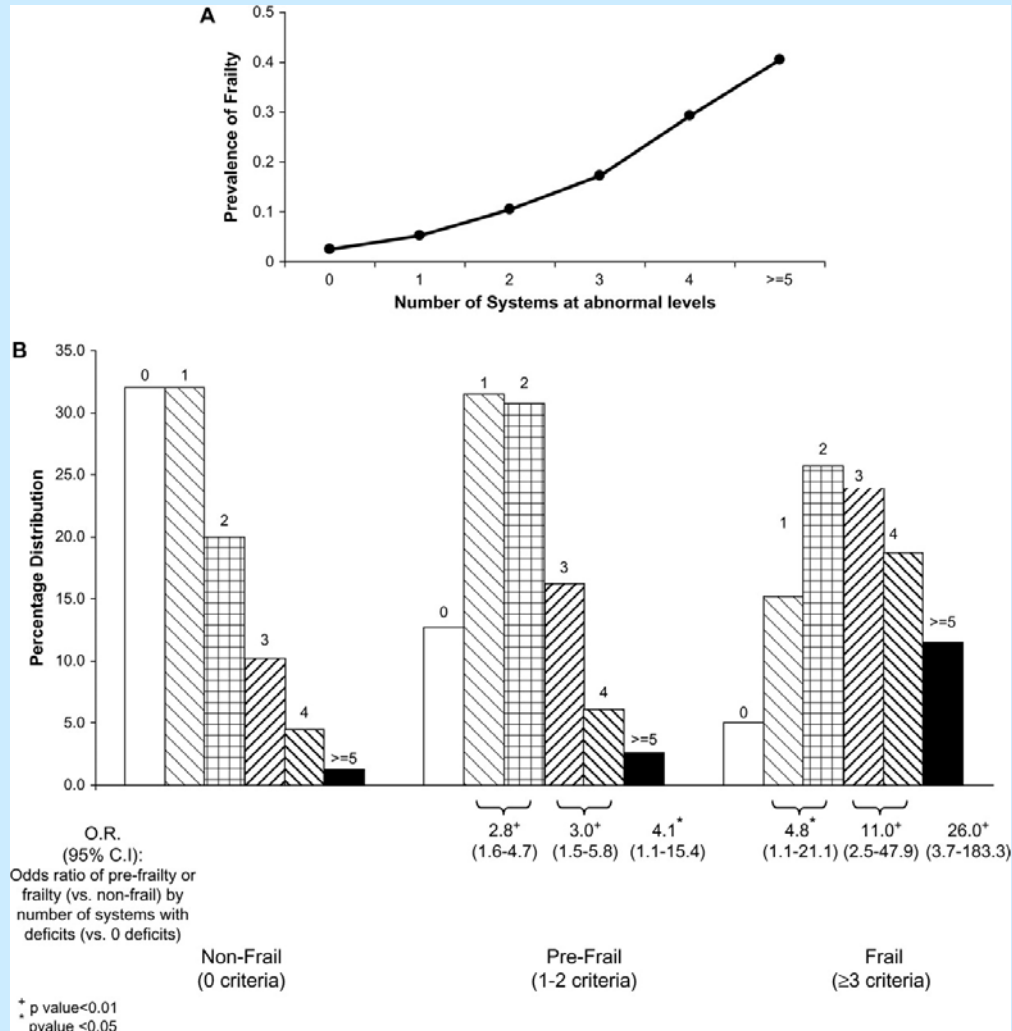
▶ **Columbia Mailman School of Public Health**

- L. Fried

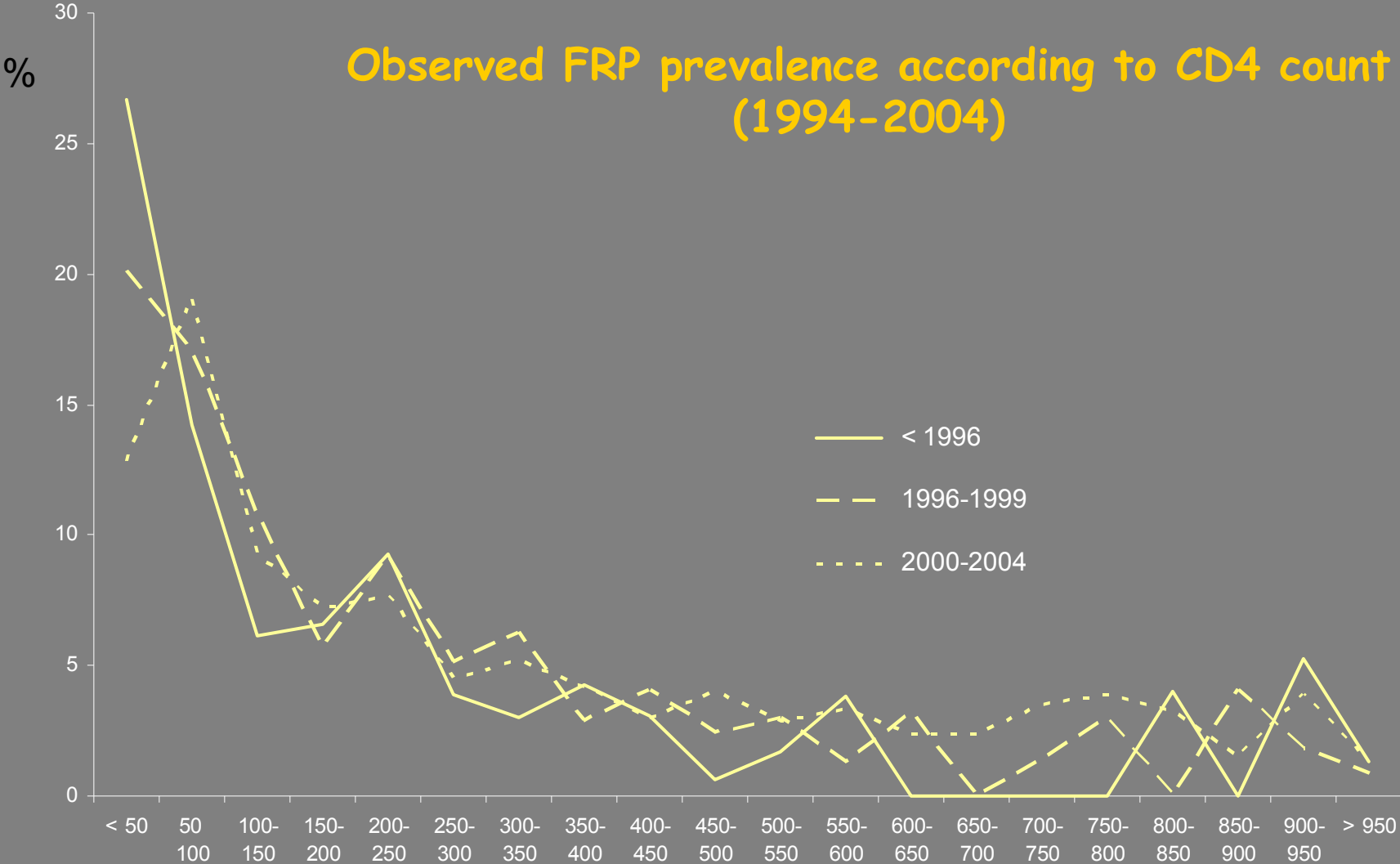
▶ **Parisian National Veterinary School**

- L. Desquilbet

Association of number of physiological systems at abnormal levels with being frail, women aged 70-79 years (p < .01 for qualitative trend)



Observed FRP prevalence according to CD4 count (1994-2004)



		CD4 Count Ranges																				Total
		< 50	50-100	100-150	150-200	200-250	250-300	300-350	350-400	400-450	450-500	500-550	550-600	600-650	650-700	700-750	750-800	800-850	850-900	900-950	> 950	
< 1996	FRP	86	22	8	10	14	7	5	7	5	1	2	4	0	0	0	0	1	0	1	1	174
	Total	322	155	131	152	151	179	165	165	163	156	116	104	71	69	38	41	25	24	19	76	2322
1996-1999	FRP	34	26	22	14	27	18	25	10	14	8	9	3	7	0	2	4	0	4	1	2	230
	Total	169	152	204	246	294	351	398	350	345	330	297	227	219	205	145	132	102	99	55	221	4541
2000-2004	FRP	9	13	9	13	17	12	18	13	10	13	8	9	6	5	6	6	4	2	4	5	182
	Total	70	68	97	180	222	266	348	310	336	325	280	271	252	208	172	155	124	131	103	389	4307

Effect of FRP status at HAART Initiation on Outcomes

- **Among AIDS-free men**

Outcomes n (%)	FRP prior to HAART ¹		Total (n=511)
	No (n=475)	Yes (n=36)	
No outcome	394 (83)	22 (61)	416 (81)
AIDS/death	81 (17)	14 (39)	95 (19)
Among AIDS/death			
AIDS	54 (67)	8 (57)	62 (65)
Death with no previous AIDS	27 (33)	6 (43)	33 (35)

FRP, frailty related phenotype

¹ within a 3-year period prior to HAART

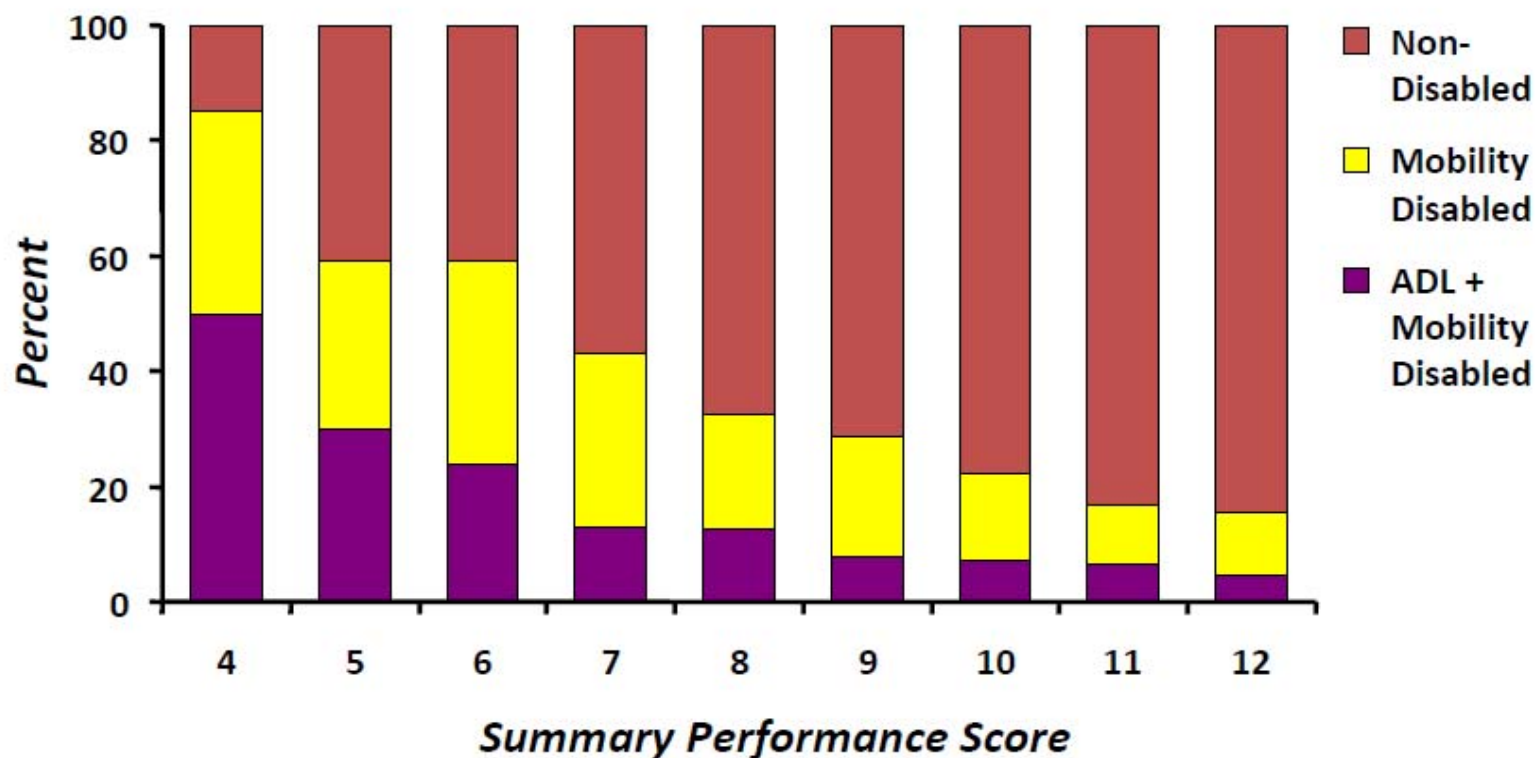
- **Among AIDS-diagnosed men**

Outcome n (%)	FRP prior to HAART ¹		Total (n=141)
	No (n=91)	Yes (n=50)	
No outcome	71 (78)	29 (58)	100 (71)
Death	20 (22)	21 (42)	41 (29)

FRP, frailty related phenotype

¹ within a 3-year period prior to HAART

Disability Status at Four Years According to Baseline Summary Performance Score Among Those Non-Disabled at Baseline (Iowa – EPESE)



ADL = activity of daily living

Guralnik JM, et al. *N Engl J Med.* 1995;332:556-561.

Current HAART Era: Age Issues

- ▶ **Immunologic mechanisms leading to worse HIV disease**
 - Decreased T-cell replacement
 - Decreased cellular response to HAART
 - Decreased function on a per lymphocyte basis
 - Dysregulation leading to impaired responses
 - Exacerbation of normal age-related immune deterioration
 - Immune activation (remains present on HAART)
- ▶ **Non-immunologic mechanisms leading to worse prognosis**
 - Frailty- may have immunologic basis even without HIV
 - Age-related morbidities
 - Interactions with HAART
- ▶ **Possibility of immune-modulatory therapy for HIV**
- ▶ **Vaccine responses**
 - HIV
 - Others

(Margolick, JB, unpublished)