

# Invecchiamento

## La Rete Omeostatica in Sciopero

**Luigi Ferrucci, MD, PhD**

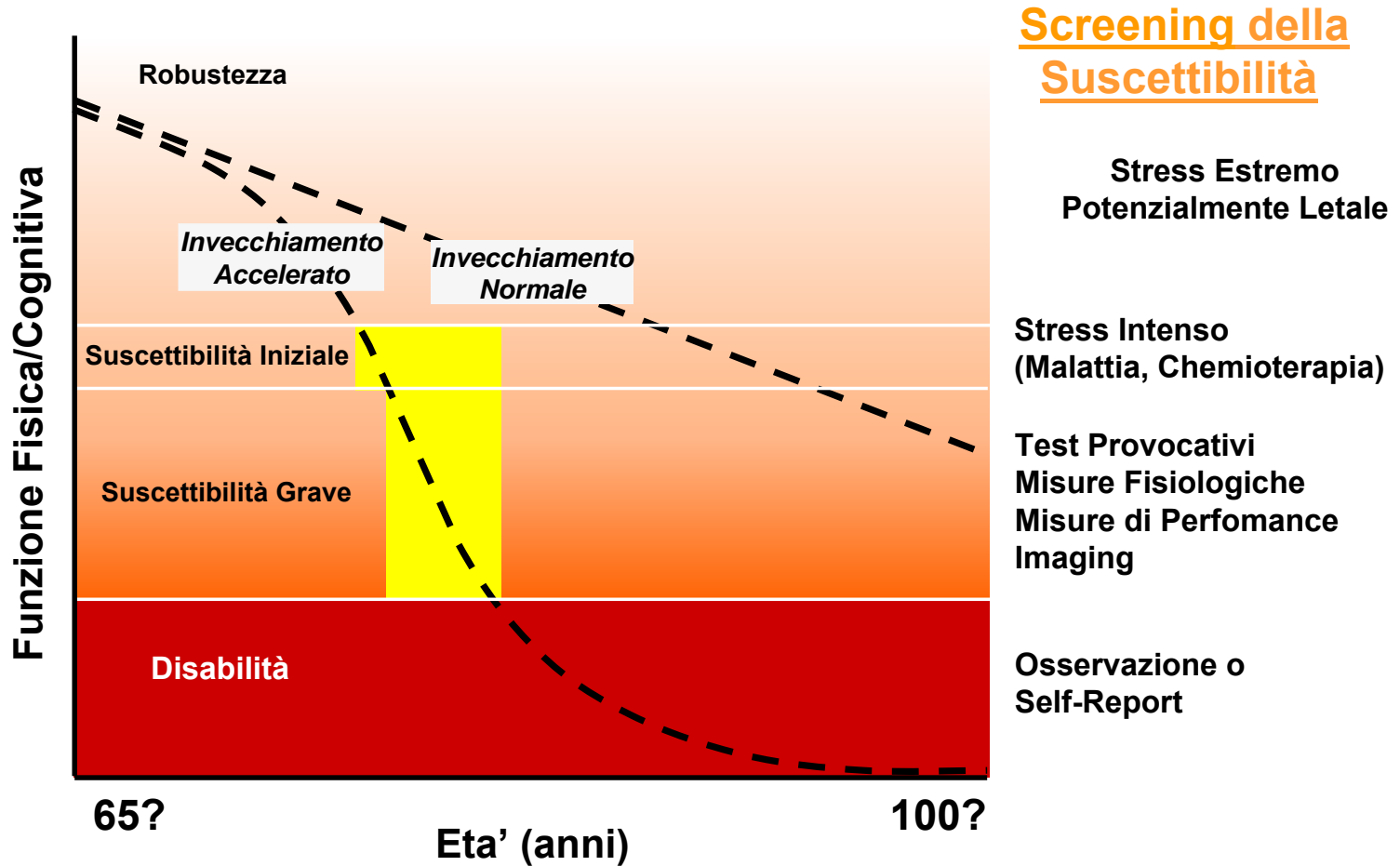
Clinical Research Branch  
National Institute on Aging  
Baltimore, MD



Pieter Bruegel: La torre di Babele, 1563

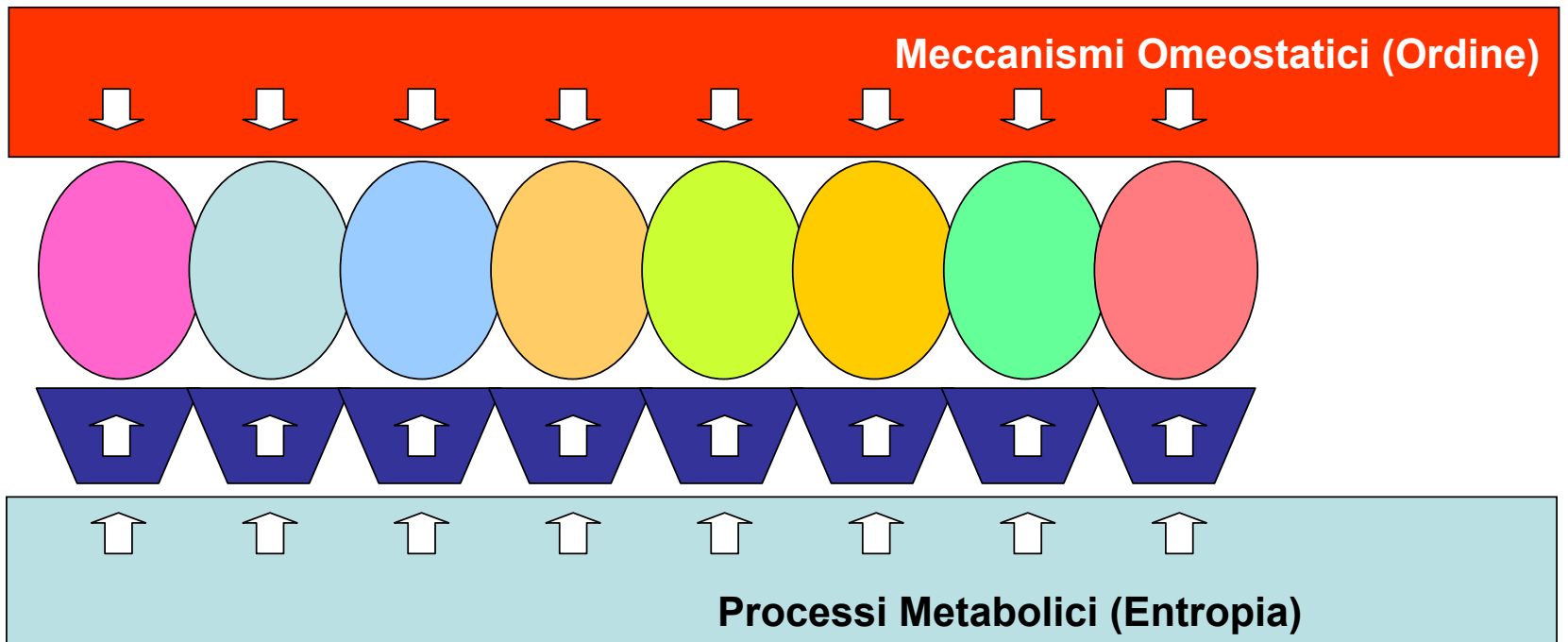
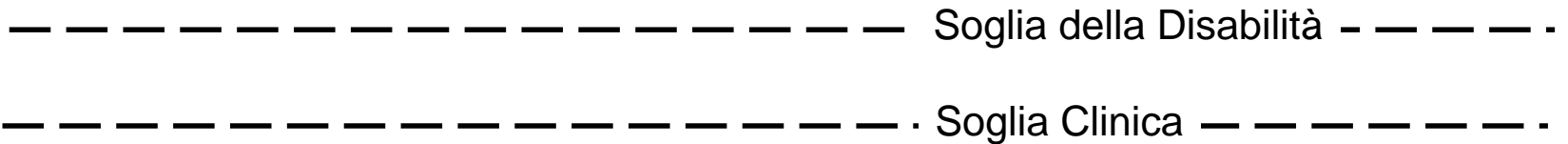
# Paradigma dell'Invecchiamento

## “La Velocità e le Soglie del Declino”



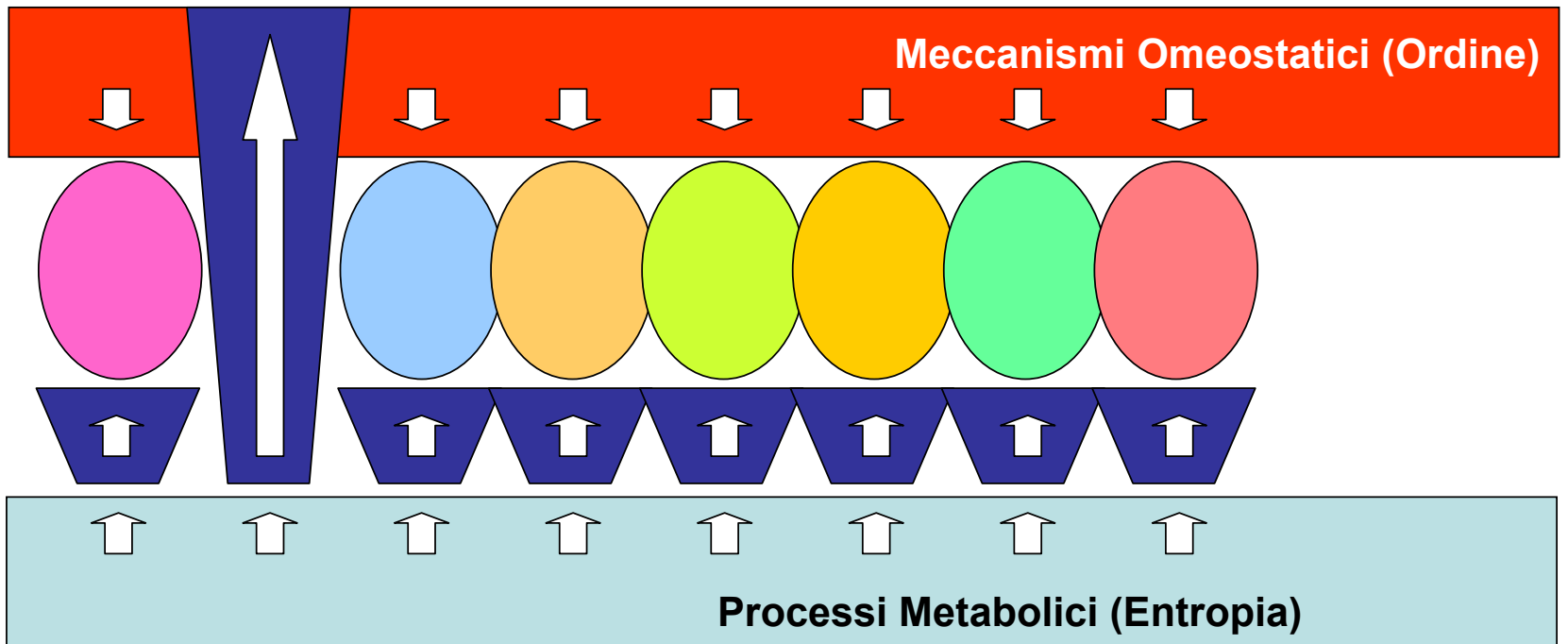
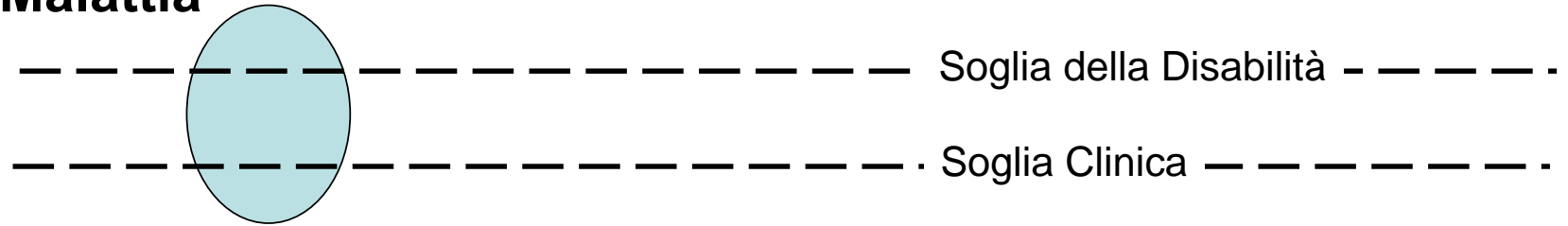
# Il Modello Omeostatico

## Perfezione Omeostatica



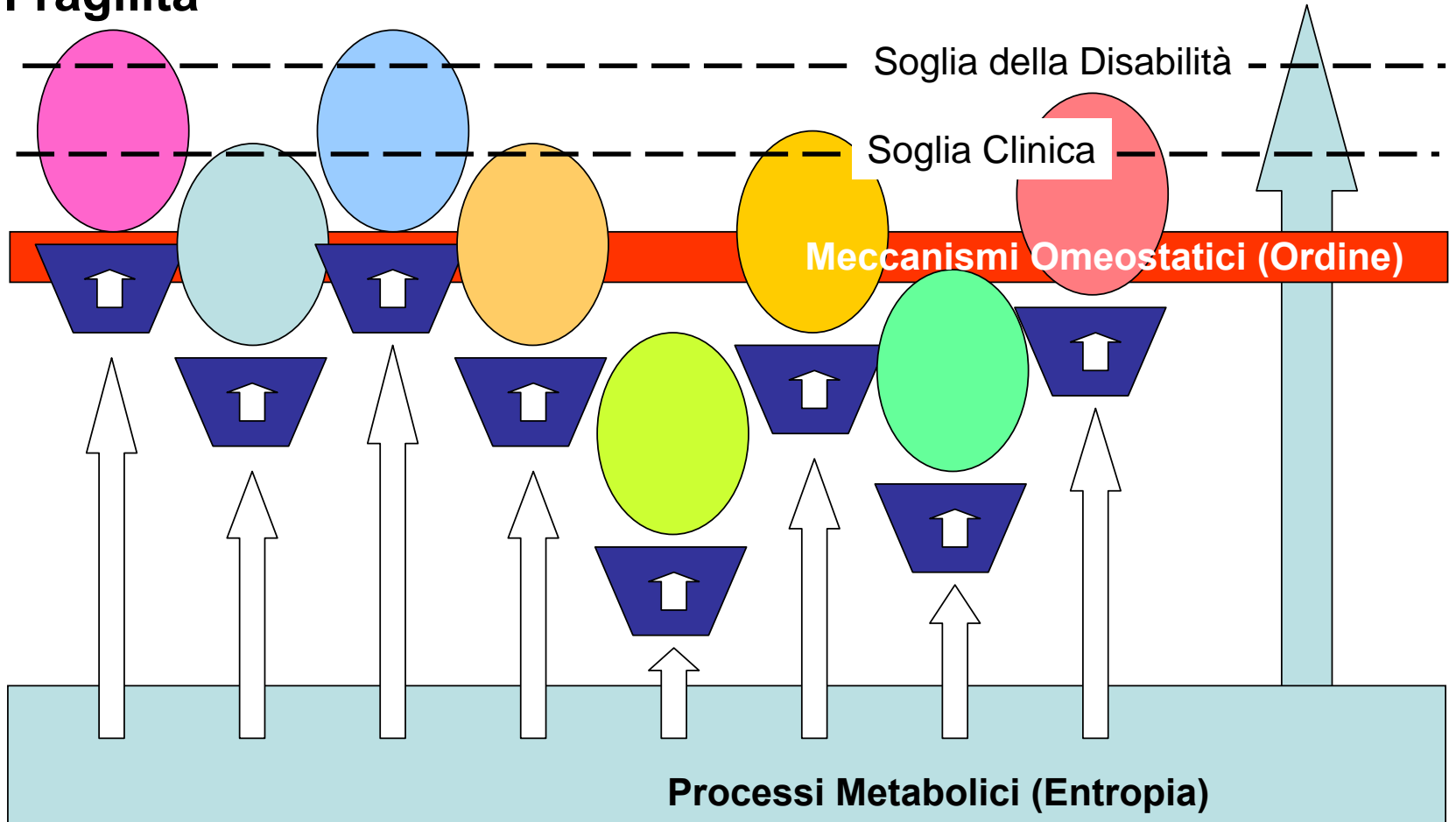
# Il Modello Omeostatico

**Malattia**

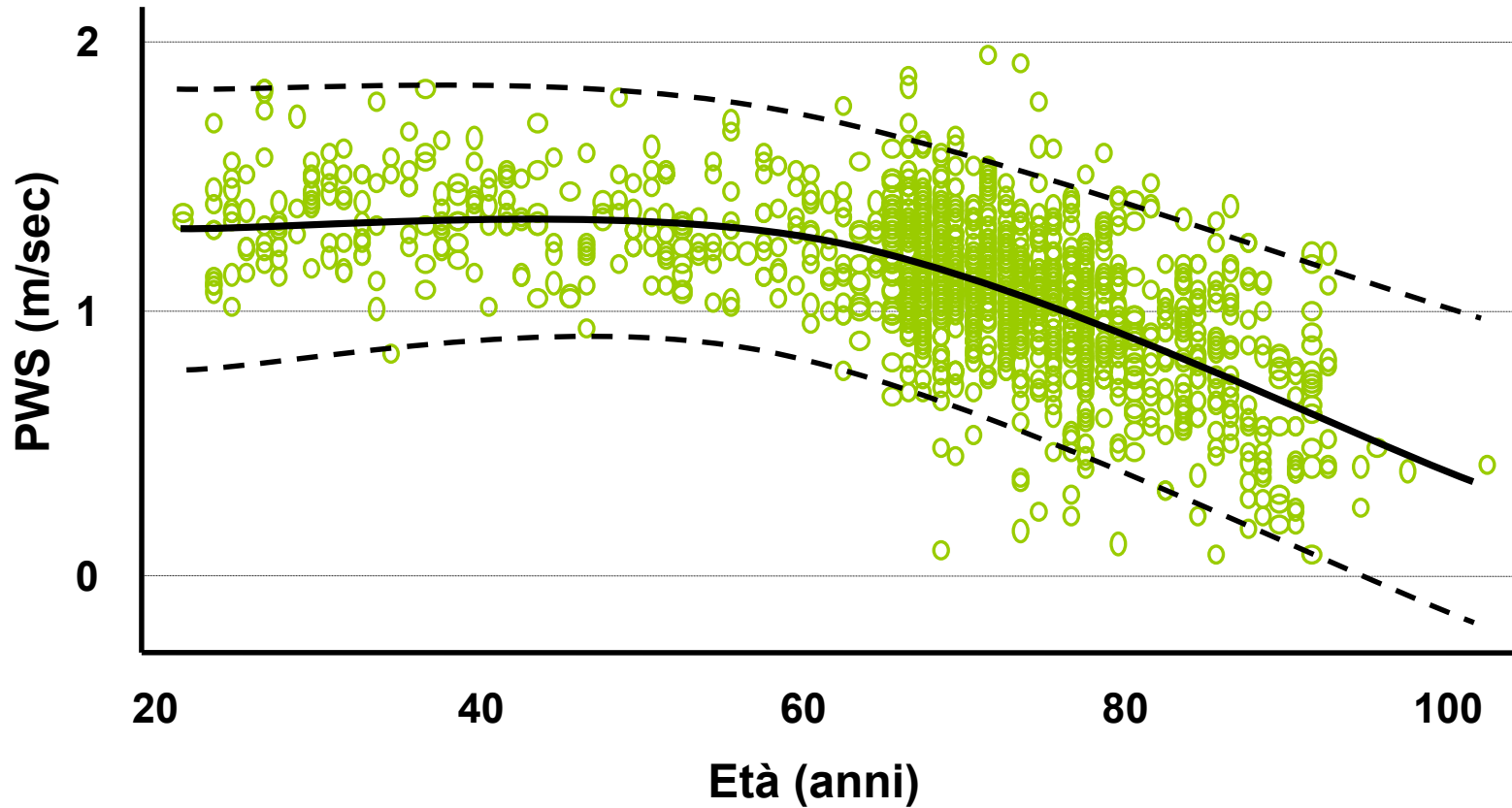


# Il Modello Omeostatico

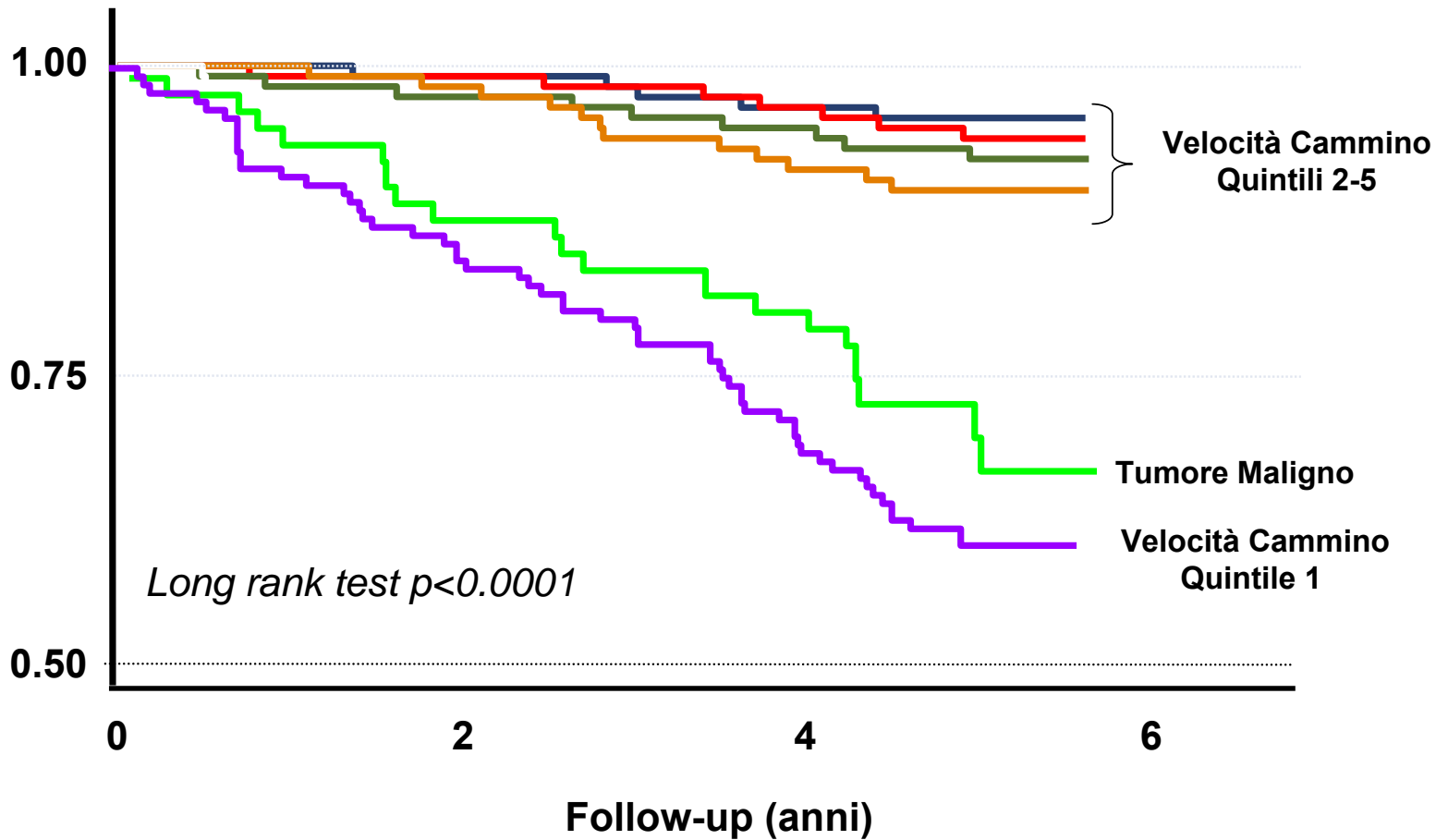
**Fragilità**



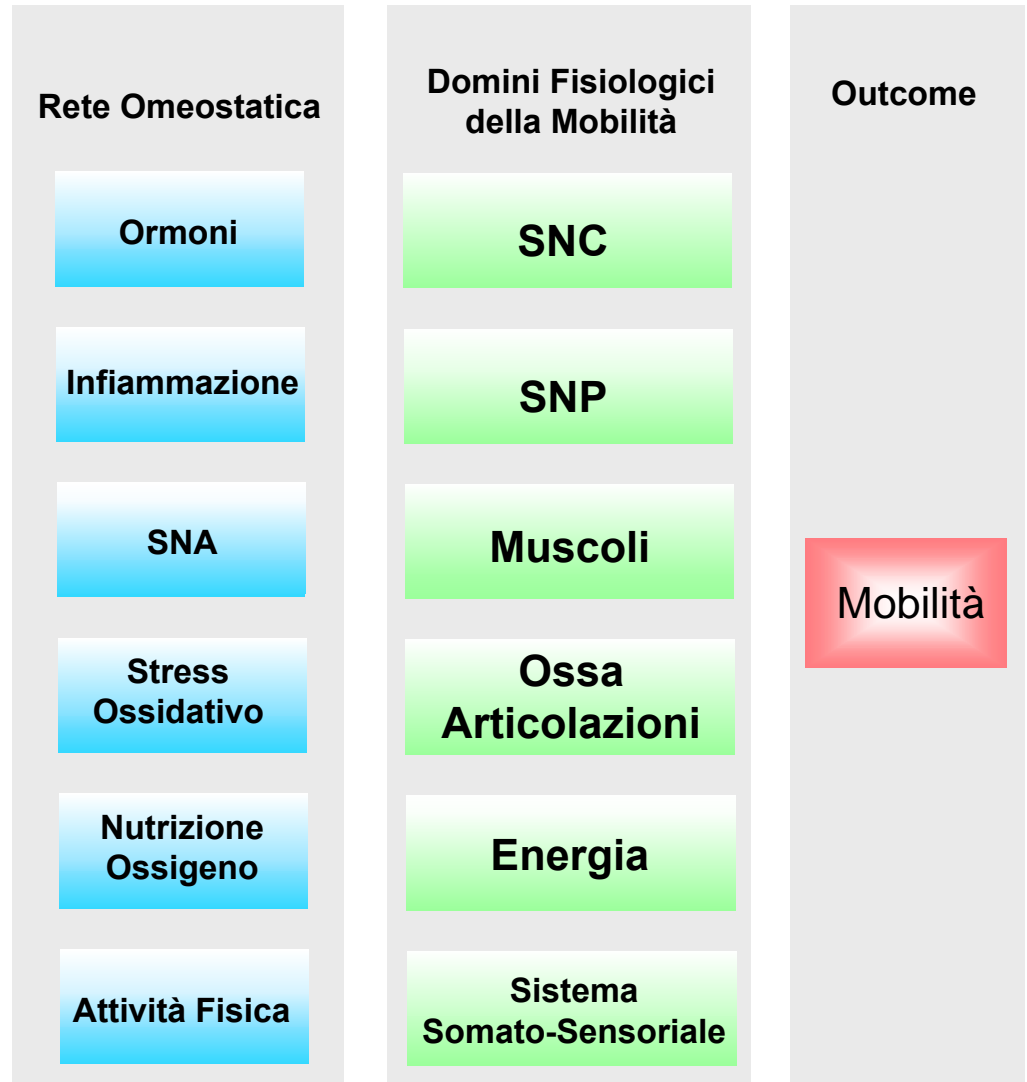
# Velocità del Cammino ed Età



# La Velocità del Cammino Predice la Mortalità



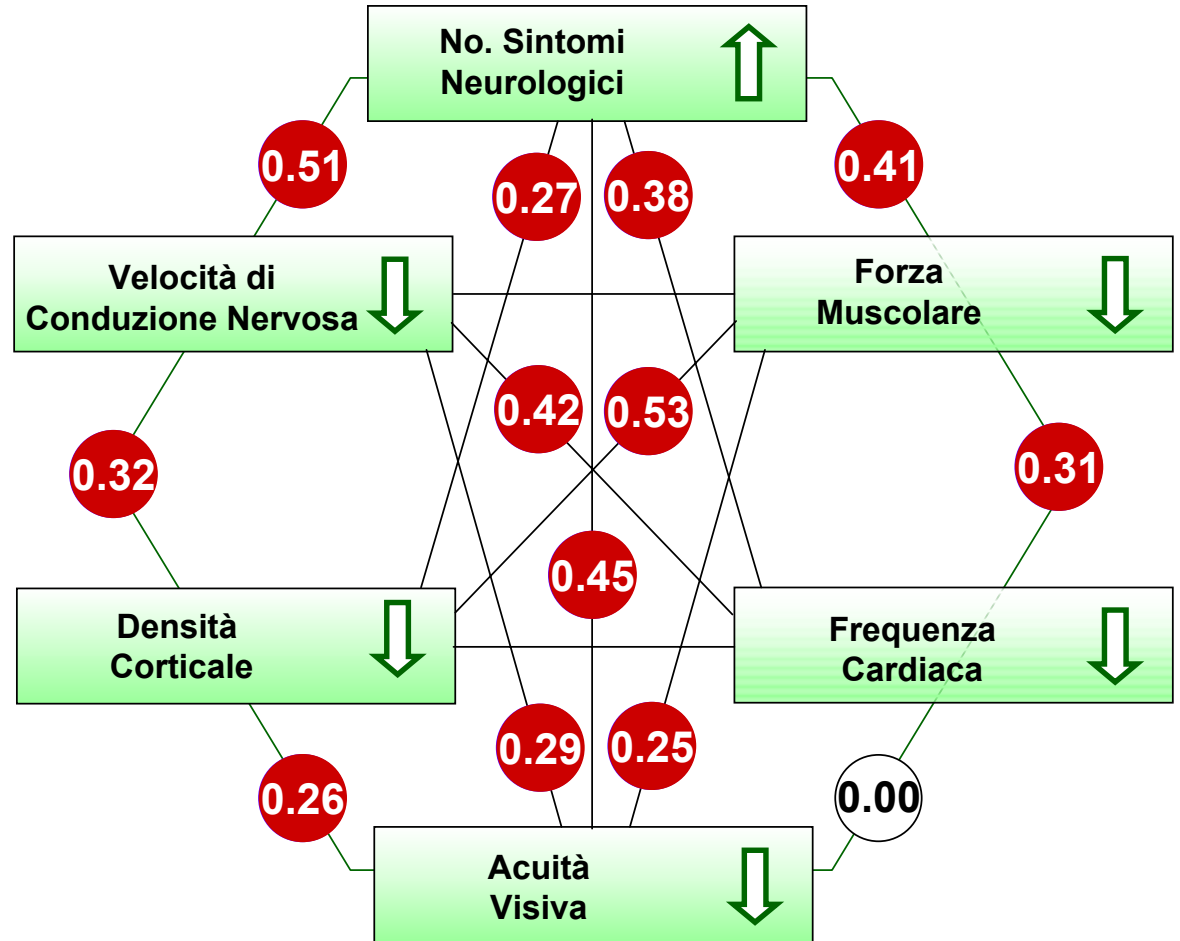
# La Rete Omeostatica





# Correlazione tra le Velocità di Variazione di Parametri Fisiologici Importanti per il Cammino

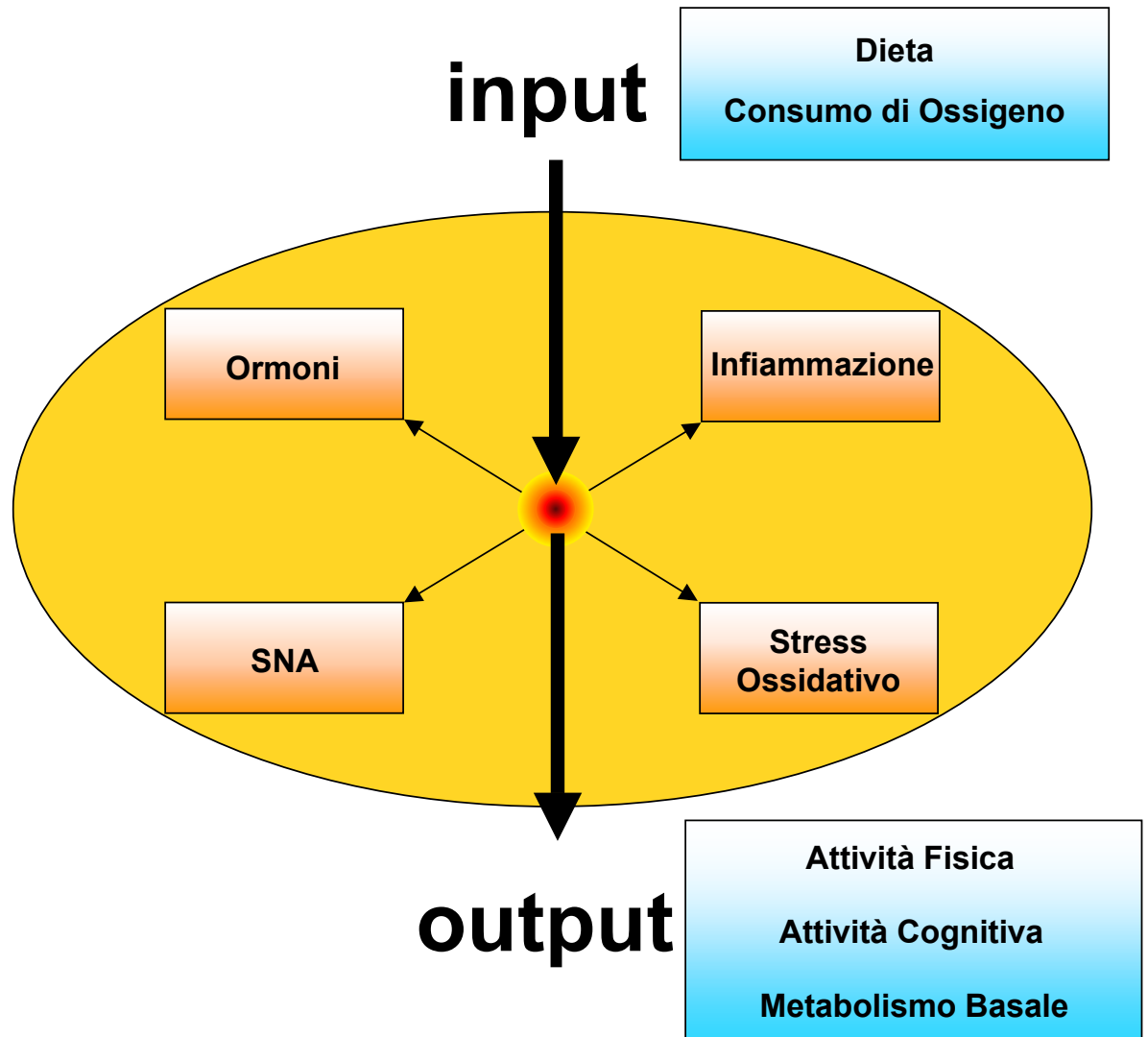
(InCHIANTI, n=1055)



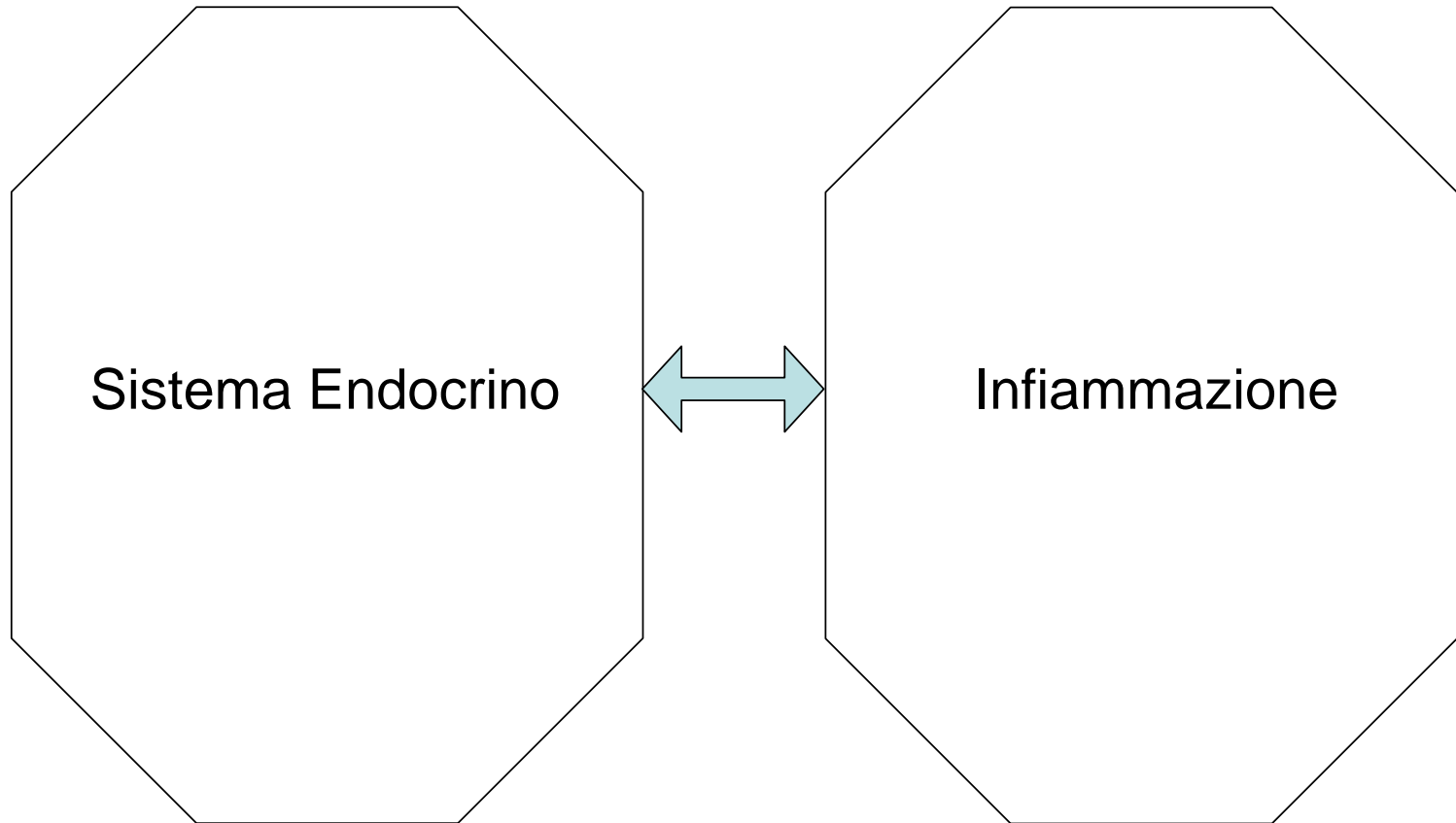
**Media Cruda**  $r=0.39$  ( $p<.001$ )

**Media Aggiustata per Età**  $r=0.13$  ( $p<.001$ )

# Rete Omeostatica



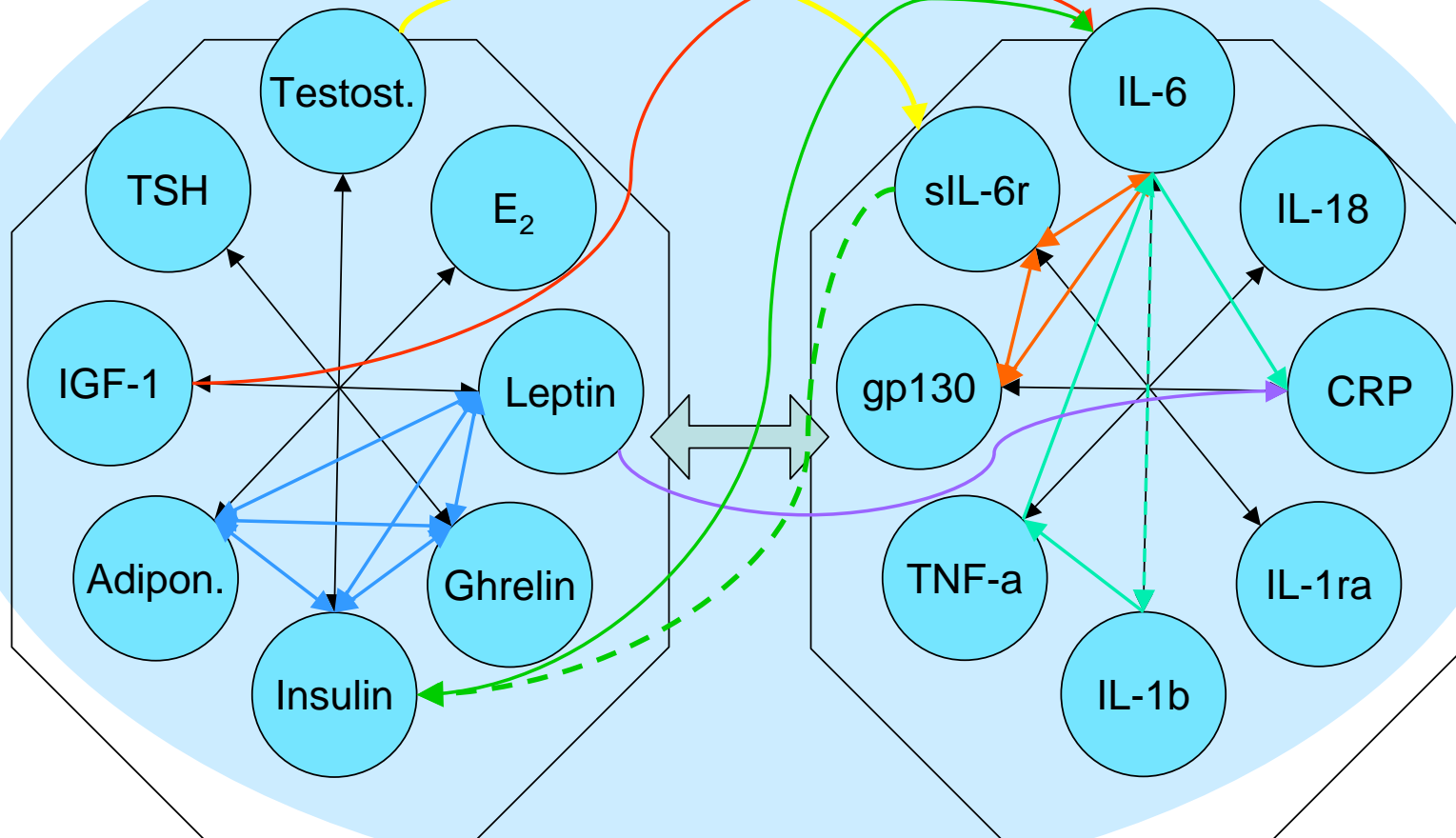
## Interazione tra Domini Diversi della Rete Omeostatica



# Rete Omeostatica

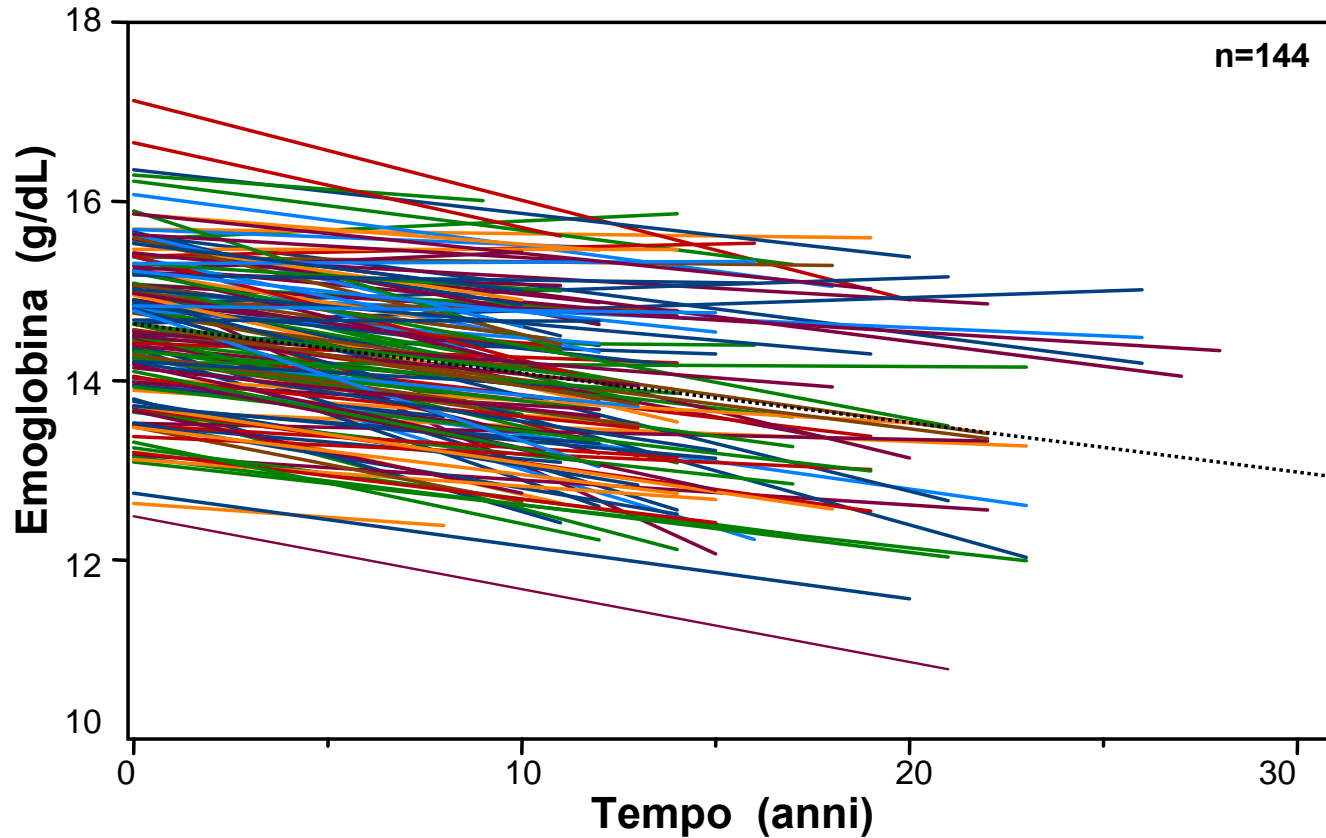
Sistema Endocrino

Inflammazione



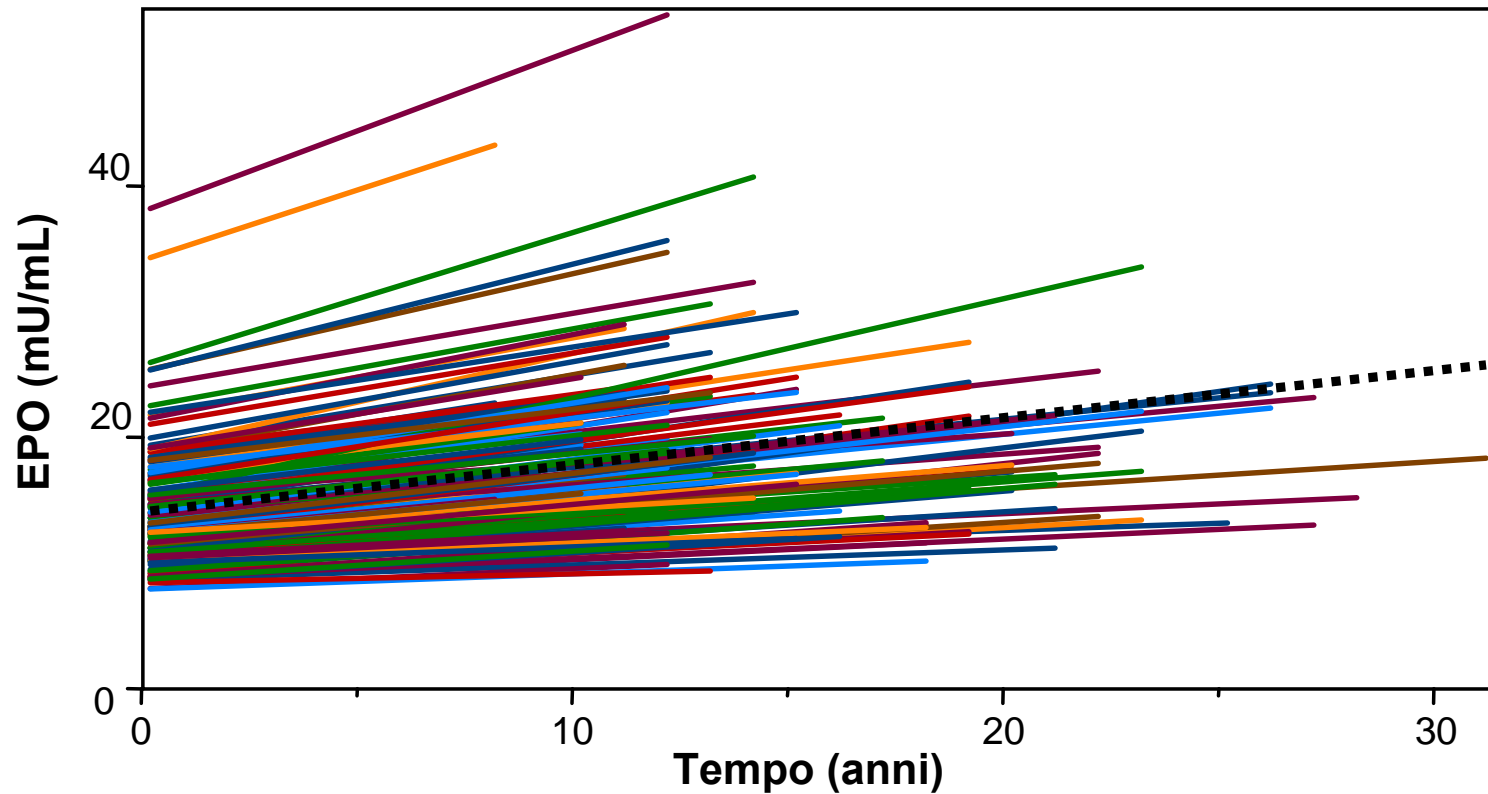
# Anemia nel Vecchio

Traiettorie Longitudinali Hb in 144 Partecipanti (BLSA)



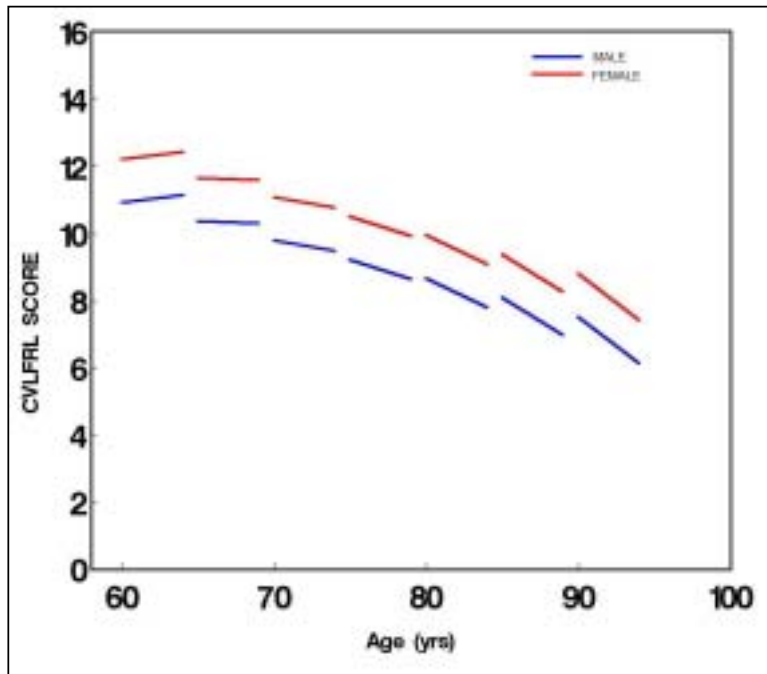
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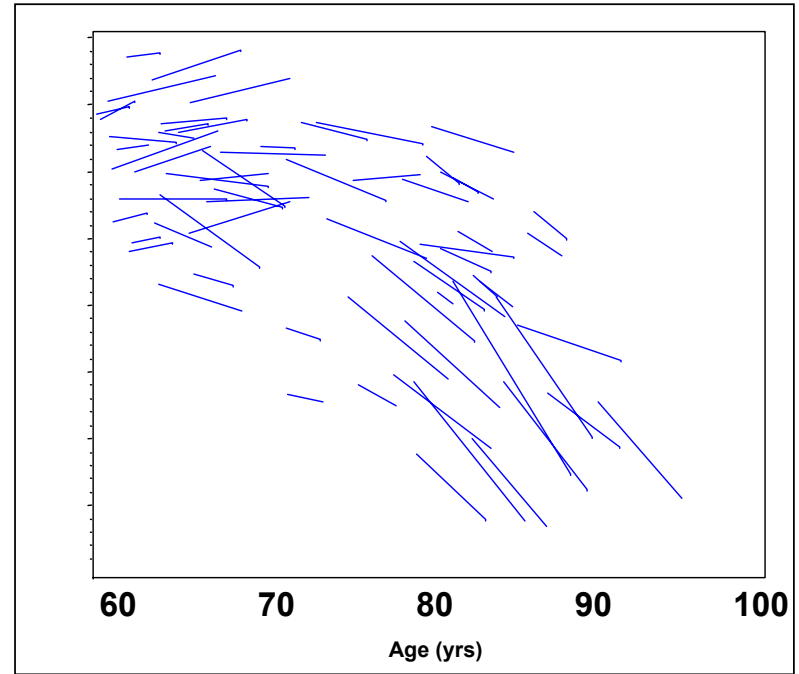


# California Verbal Learning Test (CVLT): Age Differences and Age Changes in Delayed Free Recall

Adapted from Lamar et al., Neurology, 2003



621 Men  
532 Women

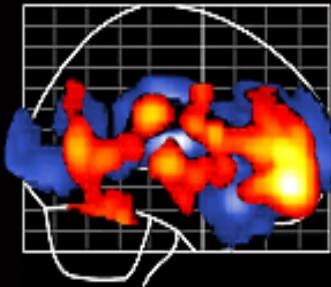


Randomly selected individual  
trajectories Age  $\geq$  60, mean = 72.08  
85 Men, 52 Women

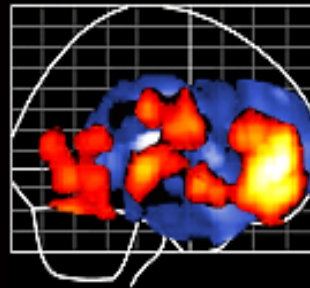
# Longitudinal Changes in rCBF from Year 1 to 9

Beason-Held, et. al. Neurobiol Aging 2008 [epub 2006]

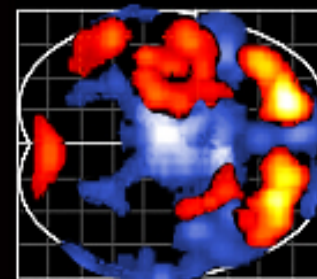
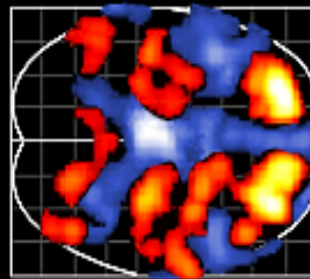
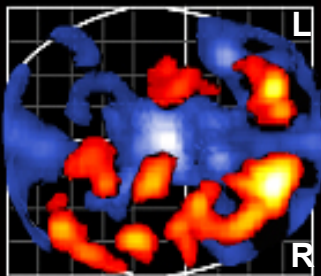
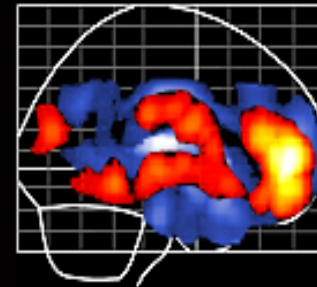
**Rest**



**Verbal**



**Figural**

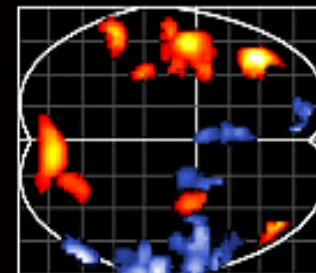
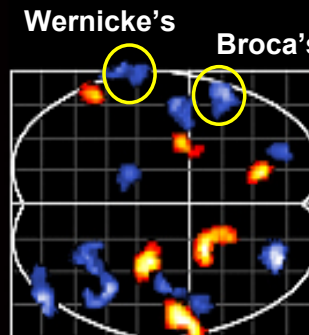
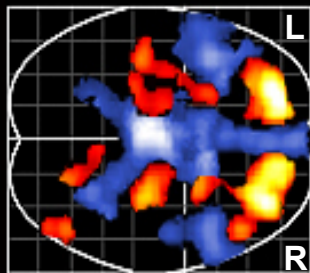
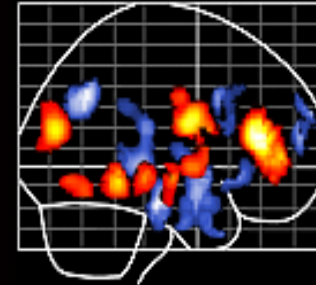
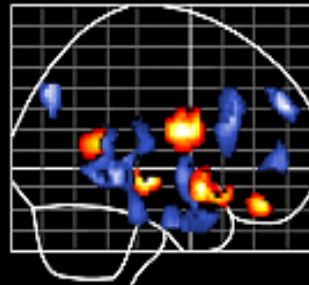
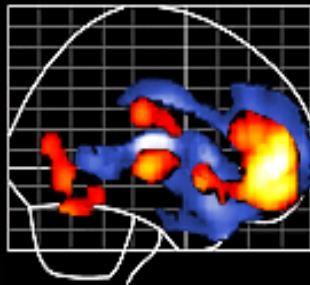


■ Increased Blood Flow  
■ Decreased Blood Flow

# Common and Modality-Specific rCBF Changes from Year 1 to 9

Common Changes Verbal Specific

Figural Specific



- Increased Blood Flow
- Decreased Blood Flow

# Longitudinal $\Delta$ rCBF: High vs. Low [ $^{11}\text{C}$ ]PIB Retention Group

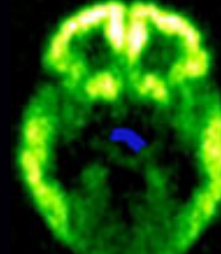
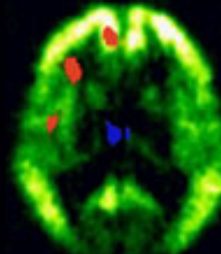
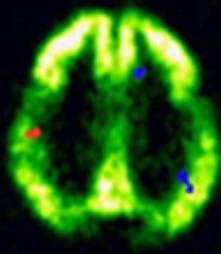
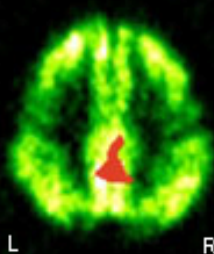
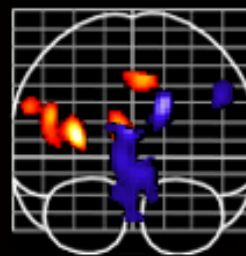
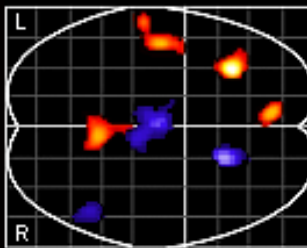
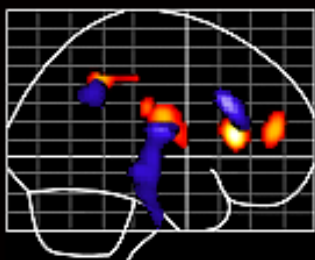
Sojkova et al. J Nucl Med, 2008;49:1465-71

## Greater decreases in $\Delta$ rCBF

- Anterior/mid cingulate gyrus (R)
- Supramarginal gyrus (R)
- Thalamus (L)
- Midbrain (B)

## Greater Increases in $\Delta$ rCBF

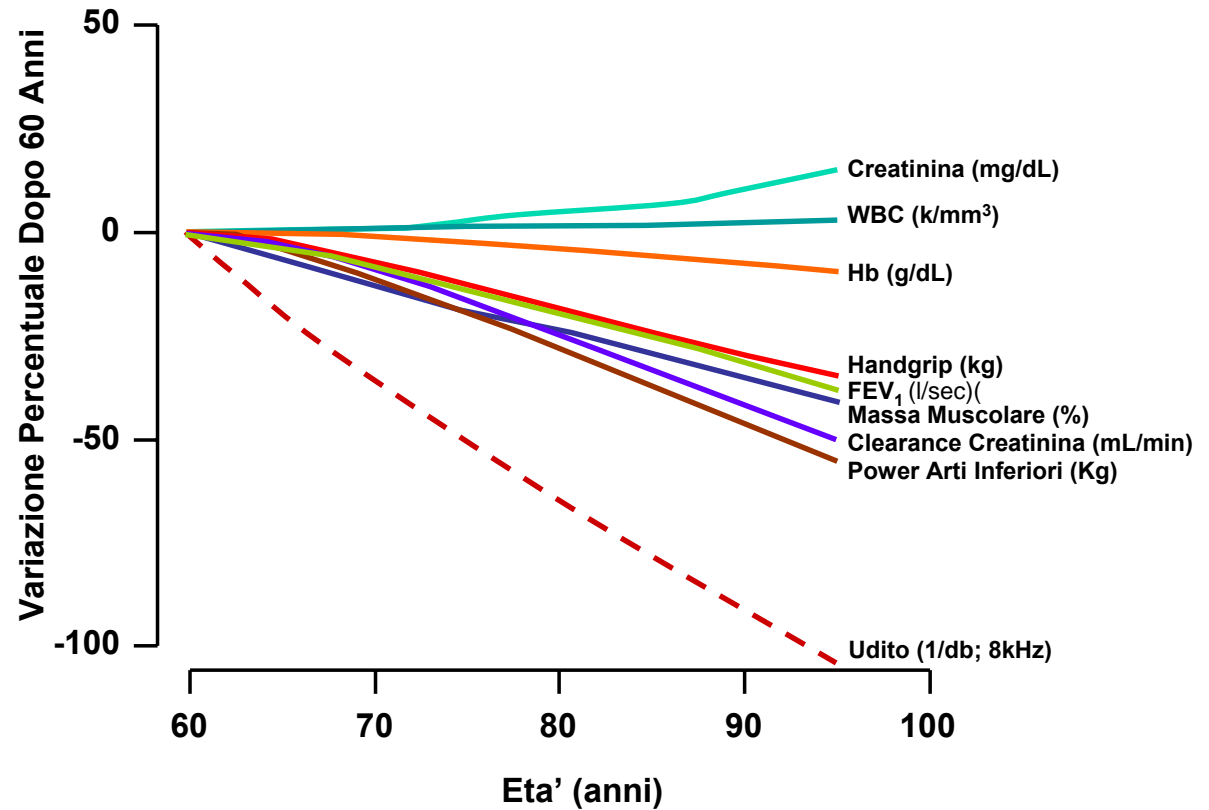
- Precuneus (R)
- Medial and inferior frontal gyrus (L)
- Postcentral gyrus (L)
- Inferior parietal lobule (L)





## Variazioni Percentuali con l'Invecchiamento di Parametri Fisiologici Multipli (60+)

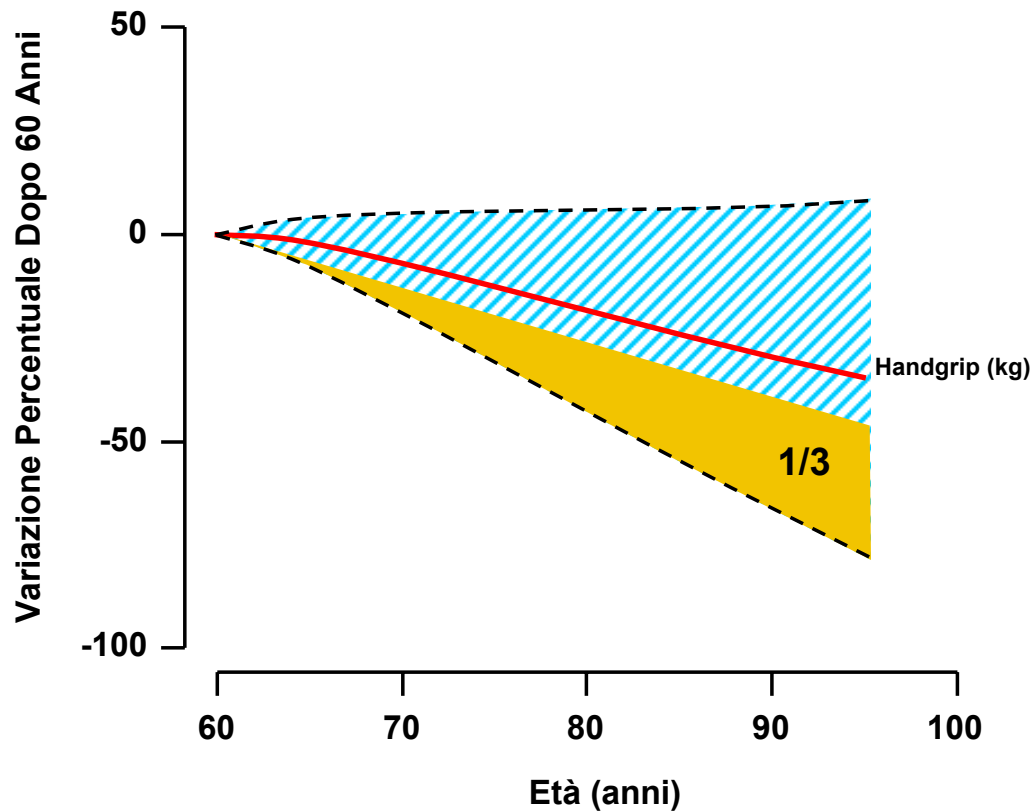
(BLSA, n=184, Valutazioni=2757)





# Variazione Percentuale Handgrip con l'Invecchiamento (60+)

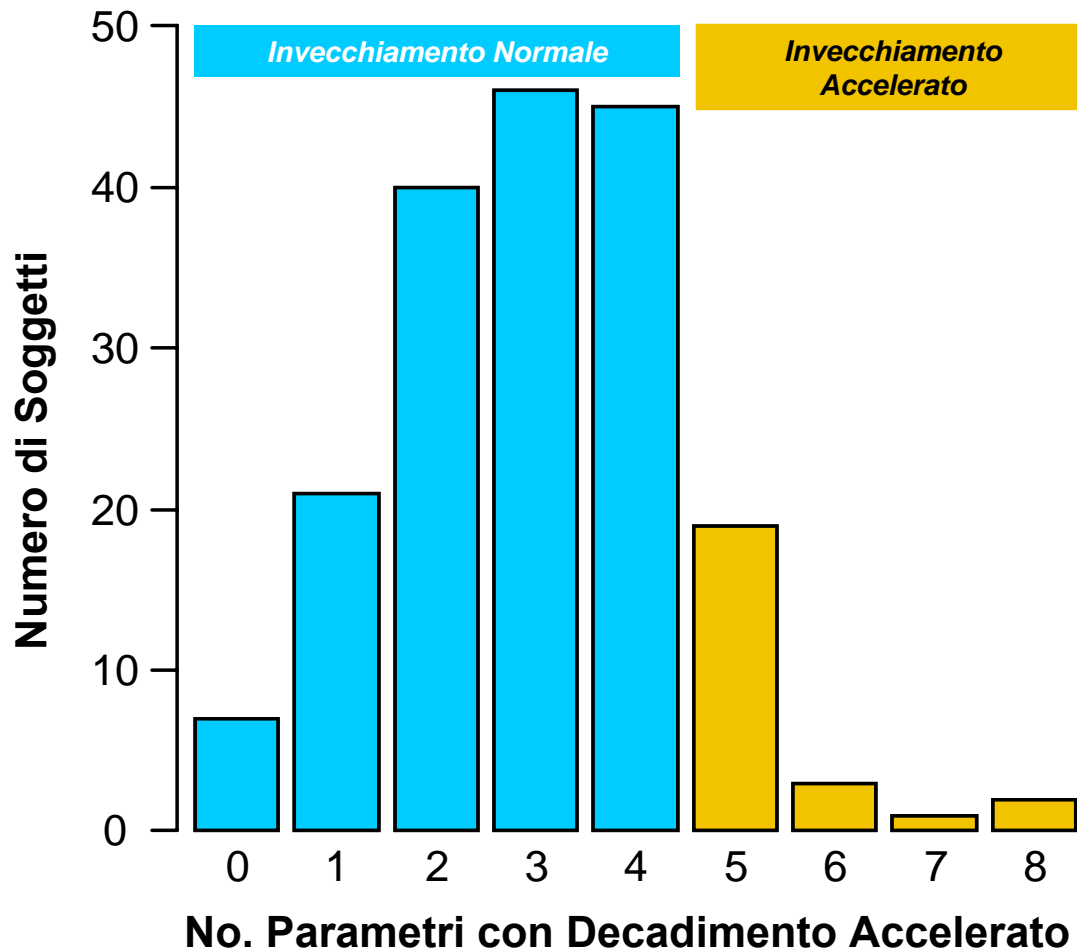
(Schematic example)





## No. Partecipanti in Funzione del Numero di Parametri Fisiologici con Decadimento Accelerato

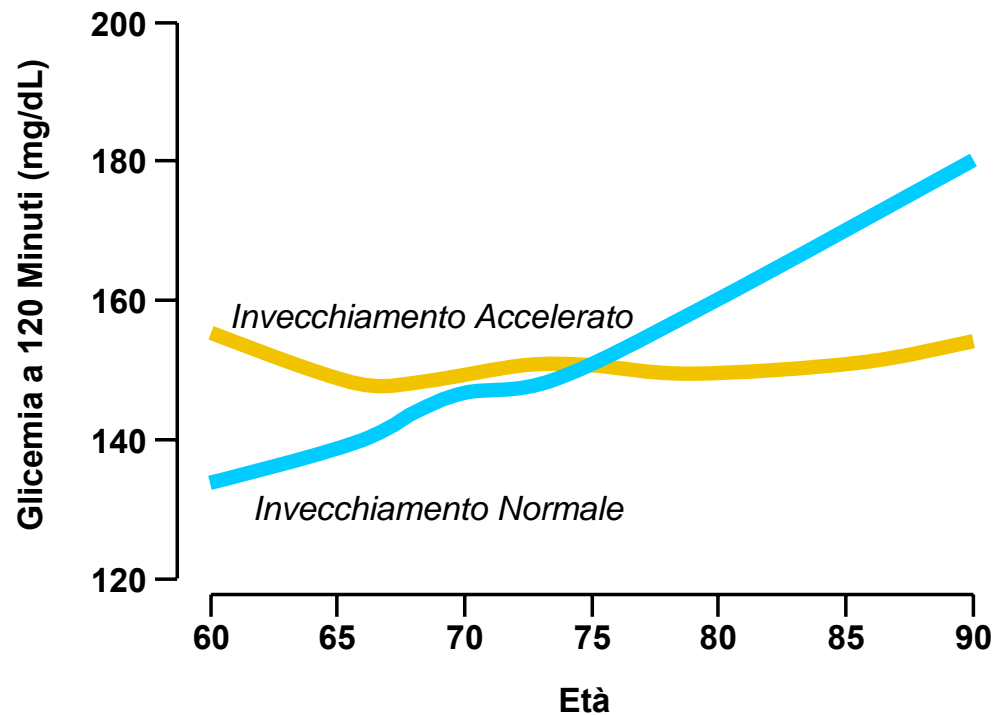
(BLSA, n=184, Visits=2757)





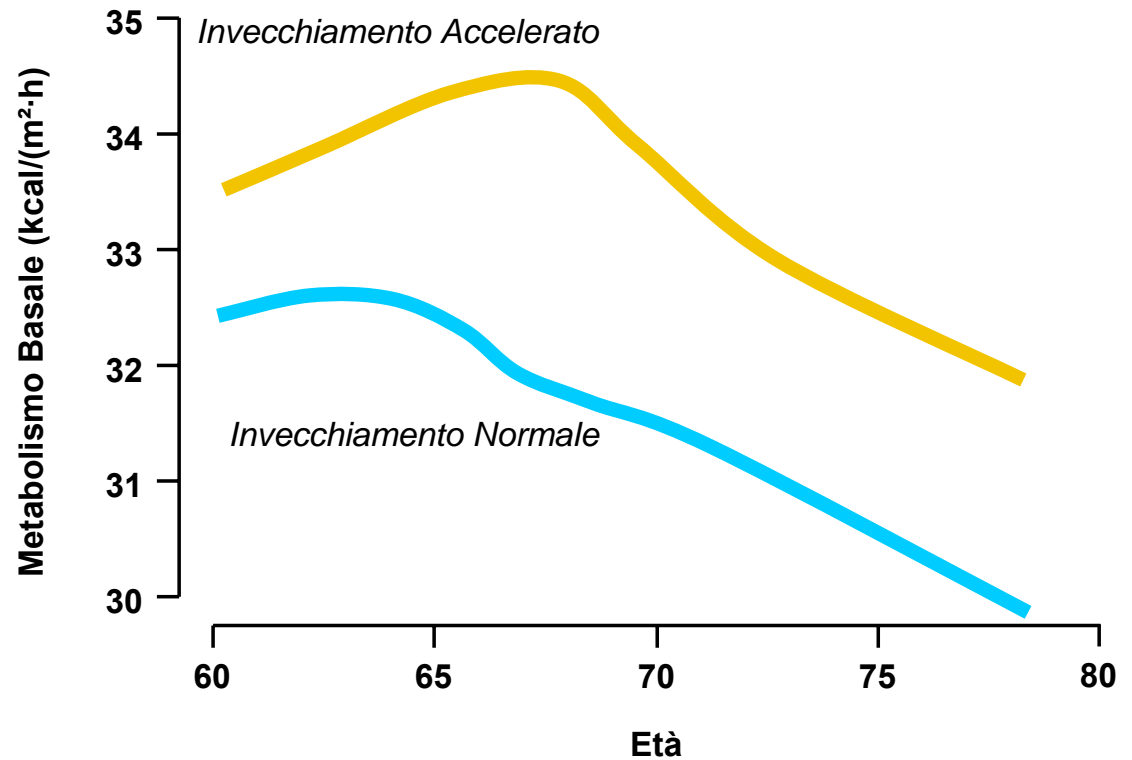
## Variazioni Longitudinali della Glicemia 120 min dopo Carico di Glucosio in Soggetti con Invecchiamento Normale ed Accelerato

(BLSA, n=184, Visits=2757)

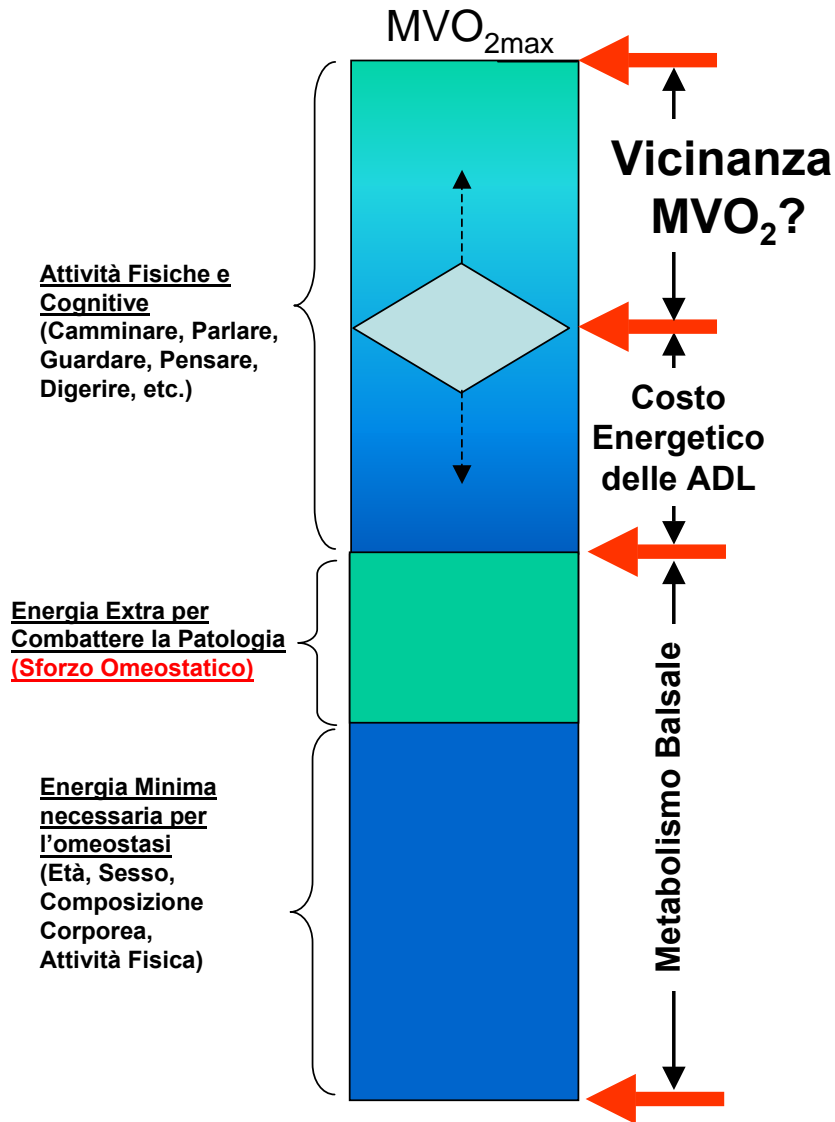


# Variazioni Longitudinali del Metabolismo Basale in Soggetti con Invecchiamento Accelerato e Normale

(BLSA, n=184, Visits=2757)



# Modello di Fragilità Energetica



## Fatica

Distanza tra costo energetico delle ADL e limiti dell'energia disponibile?

La relazione reciproca dipende da tre fattori:

1.  $MVO_{2max}$
2. Metabolismo Basale (MB)
3. Costo Energetico delle ADL (CEA)  
(Efficienza =  $1/CEA$ )

$MVO_{2max}$  inversamente correlato alla **Fatica**

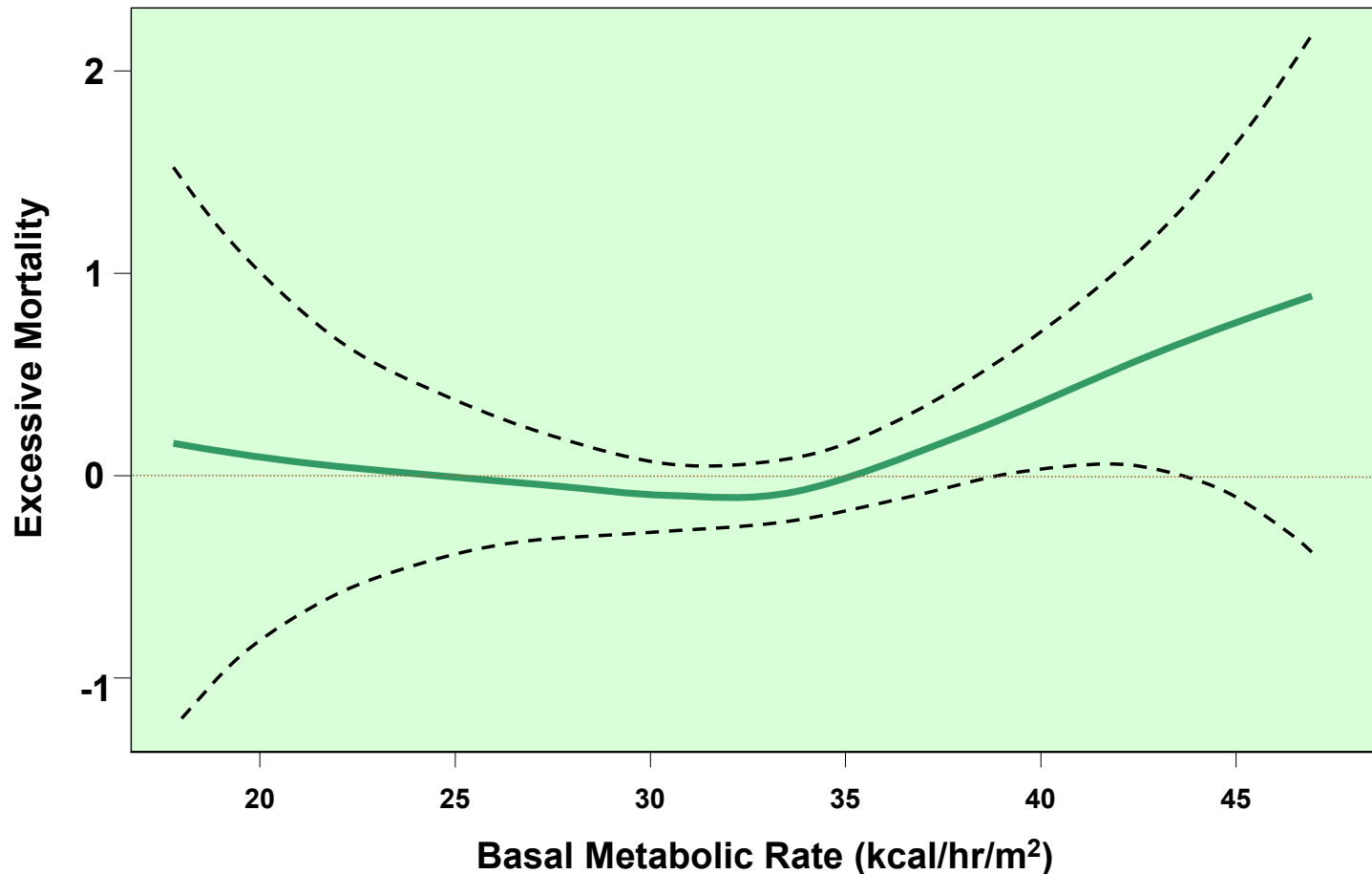
MB direttamente correlato alla **Fatica**

Efficienza direttamente correlata alla **Fatica**

+ modulazione della componente percettiva (infiammazione, depressione etc.).

# High Basal Metabolic Rate Is a Risk Factor for Mortality: The Baltimore Longitudinal Study of Aging

Carmelinda Ruggiero,<sup>1,4</sup> E. Jeffrey Metter,<sup>1</sup> Vojtech Melenovsky,<sup>2</sup> Antonio Cherubini,<sup>4</sup> Samer S. Najjar,<sup>3</sup> Alessandro Ble,<sup>1</sup> Umberto Senin,<sup>4</sup> Dan L. Longo,<sup>1</sup> and Luigi Ferrucci<sup>1</sup>



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