

Susan Coetzer

Geriatrician at Wits Donald Gordon Medical Centre







FRAILTY

- Definition
- Epidemiology
- Pathophysiology
- Identification
- Differential diagnosis
- Outcomes
- Management/Prevention





DEFINITION

- CLINICAL DEFINITION: "a state of increased vulnerability to stressors due to agerelated declines in physiologic reserve across neuromuscular, metabolic and immune systems" (Walston, Hadley et al. 2006)
- CELLULAR DEFINITION: Homeostenosis decreased physiologic reserves to meet the challenges to homeostasis.(Resnick and Marcantonio 1997)
- Thus: excess demand on a reduced capacity (Ahmed, Mandel et al. 2007)



DISABILITY VS FRAILTY

- Disability inability to perform ADL's
- Disability does not affect body across multiple organ systems
- Frailty can lead to disability and vice versa, but they are distinct entities.



- Age, chronic conditions and disability do not establish the diagnosis (Walston 2017)
- It's a spectrum biological disorder
 - Robust pre-frail frail failure to thrive (advanced/end-stage)-death (Ahmed, Mandel et al. 2007)





PREVALENCE

- Difficult to determine due to different definitions used in studies.
 - Physical frailty definitions 9.9% for frailty and 44.2% for pre-frailty (Collard, Boter et al. 2012)
 - Cardiovascular health study 6.9%
 - Broad phenotype definitions 13.6%
- Women more than men (9.2% vs 5.2%)
- SA rural population 7%
- Tanzania (community dwelling) 19.1%



PATHOPHYSIOLOGY

Hypothesized model of frailty and adverse health outcomes



CRP: C-reactive protein; IL: interleukin; IGF: insulin-like growth factor; DHEA-S: dehydroepiandrosterone sulfate.

Reproduced with permission from: Walston J, Hadley EC, Ferrucci L, et al. Research Agenda for Frailty in Older Adults: Towards a Better Understanding of Physiology and Etiology. J Am Geriatr Soc 2006; 54:991. Copyright © 2006 Wiley-Blackwell.



RISK FACTORS

- Older age
- Current smoker
- Lower education level
- Current use of HRT (Fugate Woods, LaCroix et al. 2005)
- Depression or use of anti-depressants (Lakey, LaCroix et al. 2012)
- Intellectual disability
- Not married
- Poor nutritional status and decreased protein intake (Beasley et al 2010; Bollwein et al 2013)
- Alzheimer's disease pathology



Clinical Frailty Scale*

I Very Fit – People who are robust, active, energetic and motivated. These people commonly exercise regularly. They are among the fittest for their age.

2 Well – People who have no active disease symptoms but are less fit than category 1. Often, they exercise or are very active occasionally, e.g. seasonally.

3 Managing Well – People whose medical problems are well controlled, but are not regularly active beyond routine walking.

4 Vulnerable – While not dependent on others for daily help, often symptoms limit activities. A common complaint is being "slowed up", and/or being tired during the day.

5 Mildly Frail – These people often have more evident slowing, and need help in high order IADLs (finances, transportation, heavy housework, medications). Typically, mild frailty progressively impairs shopping and walking outside alone, meal preparation and housework.

6 Moderately Frail – People need help with all outside activities and with keeping house. Inside, they often have problems with stairs and need help with bathing and might need minimal assistance (cuing, standby) with dressing.



7 Severely Frail – Completely dependent for personal care, from whatever cause (physical or cognitive). Even so, they seem stable and not at high risk of dying (within ~ 6 months).

8 Very Severely Frail – Completely dependent, approaching the end of life, Typically, they could not recover even from a minor illness.



9. Terminally III - Approaching the end of life. This category applies to people with a life expectancy
6 months, who are not otherwise evidently frail.

Scoring frailty in people with dementia

The degree of frailty corresponds to the degree of dementia. Common **symptoms in mild dementia** include forgetting the details of a recent event, though still remembering the event itself, repeating the same question/story and social withdrawal.

In moderate dementia, recent memory is very impaired, even though they seemingly can remember their past life events well. They can do personal care with prompting.

In severe dementia, they cannot do personal care without help.

 I. Canadian Study on Health & Aging Revised 2008.
K. Rockwood et al. A global clinical measure of fitness and frailty in eldenly people. CMAJ 2005;173:489-495.

© 2007-2009 Version 12: All rights reserved, Genatic Medicine Research, Dahousie University, Halifax, Canada, Permission granted to copy for research and educational purposes only





Comprehensive Frailty Assessment Instrument CFAI

- Self-reporting
- Domains physical, psychological, social and environmental, (cognitive)
- Helps towards a more holistic approach to frailty and appropriate interventions



3 Linda Fried – cardiovascular health study –

Fried Frailty Index (FFI)

- Weight loss (>5% of body weight in the last year)
- Exhaustion (positive response to the question regarding effort required for activity)
- Weakness (decreased grip strength)
- Slow walking speed (>6-7 seconds to walk 15feet (3m))
- Decreased physical activity (kcals spent per week males <383 kcals; females <270kcal)



4 Study for osteoporotic fractures frailty tool

- Weight loss of >5% in last year
- Inability to rise from chair five times without use of arms
- "No" response to "do you feel full of energy"

5 Rockwood/CSHA study

- Cumulative deficit
 - 92 problems the more you have, the higher risk for frailty



5. FRAIL scale

- F Are you fatigued?
- R Resistance (can you climb a flight of stairs?)
- A Activity (can you walk around the block?)
- I Number of Illnesses (\geq 5)
- L Loss of weight (>5% in 6 months)



DIFFERENTIAL DIAGNOSIS (WEIGHT LOSS, WEAK AND IMPAIRED FUNCTIONAL ABILITIES)

- Depression
- Malignancy
- Rheumatologic disease (PMR, vasculitis)
- Endocrine disease (DM, thyroid)
- Cardiovascular disease
- Renal disease
- Hematologic disease
- Nutritional deficits
- Neurologic disease



LABORATORY TESTS

- FBC
- Urea/creat and electrolytes
- Calcium and albumin
- Basic liver enzymes
- Vit B12
- Vit D
- TSH





WHY IS IT IMPORTANT?

- In community:
 - Hospitalization
 - Institutionalisation
 - Death
 - Depression
 - Impaired cognition
 - Falls risk (and increased risk of hip fractures)
- In hospital:
 - Predicts surgical outcome
 - 20x less likely to go home
 - 2.5x more complications
 - Increased risk of developing disabilities (7x)
 - Early re-admission
 - Impaired mobility



OTHER USEFUL TOOLS

RANKIN scores

- 1 - no significant disability
- 2 slight disability, but able to look after own affairs without help
- 3 - moderate disability, but can walk without help
- 4 unable to attend to own bodily needs without help
- 5 - bedridden, requiring constant nursing care
- 6 - death



IN URBAN SA – HOSPITALISED PATIENTS







	Year	Country	Participants (n)	Length of follow-up (years)	Falls (HR*/OR† [95% CI])		Worsening disability (HR*/OR† [95% CI])		Hospitalisation (HR*/OR† [95% CI])		Care home admission (HR*/OR† [95% CI])		Mortality (HR*/OR† [95% CI])	
					Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty
Cardiovascular Health Study (CHS) ³	2001	USA	5317	7	1·12* (1·00- 1·26)	1·23* (0·99– 1·54)	1·55* (1·38- 1·75)	1·79* (1·47- 2·17)	1·11* (1·03- 1·19)	1·27*, (1·11- 1·46)	NA	NA	1·32* (1·13- 1·55)	1.63* (1.27- 2.08)
Canadian Study of Health and Aging (CSHA) ⁹⁷	2004	Canada	9008	5	NA	NA	NA	NA	NA	NA	2·54 † (1·67– 3·86)	2·60† (1·36- 4·96)	2-54† (1-92– 3-37)	3-69† (2-26– 6-02)
Women's Health and Aging Study (WHAS) ⁹³	2006	USA	1438	3	0·92* (0·63- 1·64)	1·18* (0·63- 2·19)	NA	NA	0·99* (0·67– 1·47)	0·67* (0·33- 1·35)	5·16* (0·81- 32·79)	23·98* (4·45- 129·2)	3·50* (1·91– 6·39)	6-03* (3-00- 12-08)
Study of Osteoporotic Fractures (SOF) ⁹⁴	2008	USA	6701	4.5	1·23† (1·02– 1·48)	2·44† (1·95- 3·04)	1·89†, (1·66– 2·14)	2·79† (2·31– 3·37)	NA	NA	NA	NA	1·54† (1·40– 1·69)	2-75* (2-46- 3-07)

HR=hazard ratio. NA=not available. OR=odds ratio. *Hazard ratio. †Odds ratio. The comparator for hazard ratios and odds ratios is people who are not frail.

Table: Covariate-adjusted associations between frailty and adverse outcomes (falls, disability, hospitalisation, care home admission, and mortality) from four large prospective cohort studies



		Participants (n)	Length of follow-up (years)	Falls (HR*/OR† [95% CI])		Worsening disability (HR*/OR† [95% CI])		Hospitalisation (HR*/OR† [95% CI])		Care home admission (HR*/OR† [95% CI])		Mortality (HR*/OR† [95% CI])	
				Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty	Inter- mediate frailty	Severe frailty
001	USA	5317	7	1·12* (1·00- 1·26)	1·23* (0·99– 1·54)	1·55* (1·38- 1·75)	1·79* (1·47- 2·17)	1·11* (1·03- 1·19)	1·27*, (1·11- 1·46)	NA	NA	1-32* (1-13- 1-55)	1.63* (1.27- 2.08)
004	Canada	9008	5	NA	NA	NA	NA	NA	NA	16	23.00	2-54† (1-92- 3-37)	3-69† (2-26– 6-02)
16	USA	1438	3	0·92* (0·63- 1·64)	1·18* (0·63- 2·19)	NA	NA	0·99* (0·67– 1·47)	0-67 5 (0-3 (1-35 5	0.81-	(4.45-	3-50* 1-91- 5-39)	6-03* (3-00- 12-08)
80	USA	6701	4.5	1·23† (1·02- 1·48)	2·44† (1·95- 3·04)	1·89†, (1·66– 2·14)	2·79† (2·31– 3·37)	NA	NA	~.79)	129.21	1·54† (1·40– 1·69)	2·75* (2·46- 3·07)
	001 004 06 08	01 USA 004 Canada 06 USA 08 USA	01 USA 5317 004 Canada 9008 06 USA 1438 08 USA 6701	01 USA 5317 7 04 Canada 9008 5 06 USA 1438 3 08 USA 6701 4.5	Intermediate frailty 01 USA 5317 7 1.12* (1.00-1.26) 04 Canada 9008 5 NA 06 USA 1438 3 0.92* (0.63-1.64) 08 USA 6701 4.5 1.23† (1.02-1.48)	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	$\begin{bmatrix} \text{Inter-} & \text{Severe} & \text{Inter-} \\ \text{mediate} & \text{frailty} & \text{frailty} & \text{mediate} \\ \text{frailty} & \text{frailty} & \text{frailty} \\ 101 & \text{USA} & 5317 & 7 & 1.12^* & 1.23^* & 1.55^* \\ (1.00- & (0.99- & (1.38- \\ 1.26) & 1.54) & 1.75) \\ 1.26) & 1.54) & 1.75) \\ 104 & \text{Canada} & 9008 & 5 & \text{NA} & \text{NA} & \text{NA} \\ 106 & \text{USA} & 1438 & 3 & 0.92^* & 1.18^* & \text{NA} \\ 1438 & 3 & 0.92^* & (0.63- \\ 1.64) & 2.19) & \text{NA} \\ 108 & \text{USA} & 6701 & 4.5 & 1.23^\dagger & 2.44^\dagger & 1.89^\dagger, \\ (1.02- & (1.95- & (1.66- \\ 1.48) & 3.04) & 2.14) \\ \end{bmatrix}$	Inter- mediate frailty Severe frailty Inter- mediate frailty Severe mediate frailty Inter- mediate frailty Severe frailty 101 USA 5317 7 1.12* (1.00- (0.99- (1.36) 1.23* (1.38- (1.47- 1.26) 1.55* (1.38- (1.47- 1.26) 1.75) 2.17) 104 Canada 9008 5 NA NA NA NA 16 USA 1438 3 0.92* (0.63- 1.64) 1.18* (0.63- (0.63- 1.64) NA NA NA 18 USA 6701 4.5 1.23† (1.02- 1.48) 2.44† 3.04) 1.89†, 2.14) 2.79† (2.31- 3.37)	Inter- mediate frailty Severe frailty Inter- mediate frailty Severe frailty Inter- mediate frailty Severe frailty Inter- mediate frailty Inter- mediate frailty 01 USA 5317 7 1.12* (1.00- 1.26) 1.23* 1.54) 1.55* 1.79* 1.79* (1.47- 1.47- 1.19) 1.11* (1.03- 1.26) 04 Canada 9008 5 NA NA NA NA NA 06 USA 1438 3 0.92* (0.63- 1.64) 1.18* 2.19) NA NA NA 0.99* (0.67- 1.47) 08 USA 6701 4.5 1.23† (1.02- 1.48) 2.44† 3.04) 1.89†, 2.14) 2.79† 3.37) NA	$ \begin{array}{ c c c c c c c c c c c c c c c c c c c$	Inter- mediate frailty Severe frailty Inter- mediate frailty 001 USA 5317 7 1.12* 1.23* 1.55* 1.79* 1.11* 1.27*, (1.03- NA 04 Canada 9008 5 NA NA NA NA NA NA 06 USA 1438 3 0.92* 1.18* NA NA NA NA 08 USA 6701 4.5 1.23† 2.44† 1.89†, (1.02- 2.79† NA 3.37) 2.79† NA 3.37) 3.37	$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	$\begin{array}{c c c c c c c c c c c c c c c c c c c $

WHY IS IT IMPORTANT?

- In specific illnesses/conditions:
 - Coronary artery disease increased mortality
 - Worse clinical outcomes with chemotherapy
 - HIV accelerated immune-function deterioration



Potential interventions along the spectrum of frailty in older adults



ACE unit: Acute Care for Elders unit; GEM: Geriatric Evaluation and Management; PACE: Program for All-Inclusive Care of the Elderly.

Modified with permission from: Walston JD, Fried LP. Frailty and its Implications for Care. Chapter 9. In: Geriatric Palliative Care, Morrison RS, Meire DE. Oxford University Press, New York 2003. p.93. Copyright ©2003 Oxford University Press.

INTERVENTIONS

Identify the goal of care

WHAT DOESN'T WORK...

- Routine testosterone replacement
- Growth hormone/GHRF supplementation
- DHEA-S





WHAT DOES WORK ...

- Comprehensive Geriatric Assessment
- Exercise especially resistance training (Chan et al 2012)
- Nutritional supplementation
 - vit D 800-1000 IU daily in deficient patients
 - Avoid weight loss caloric and protein supplementation
 - Mediterranean diet (Talegawkar SA et al 2012)
- Medication review
 - Stop unnecessary drugs
 - Consider side-effects as a potential cause for a new symptom
 - Consider non-pharmacologic approaches
 - Substitute with safer alternatives
 - Reduce the dose
 - Use beneficial therapies when indicated
- Palliative care





CONCLUSION

- Frailty = a biological syndrome with low reserve and therefore low resistance to stressors
- It is due to cumulative decline across multiple physiological systems
- It leads to increased vulnerability to adverse outcomes ...disability, poor quality of life and death
- It is important to identify it before it is too late

