Impact of Frailty in Critically Ill Patients: Does It Add Any Value?

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- I am not a geriatrician nor necessarily an expert in frailty





- 1. Define and review the biologic concept of "frailty"
- 2. Understand the tools to capture frailty and identify a vulnerable population
- 3. Understand the impact of frailty on outcomes after acute stress and critical illness



FRAILTY is a multi-dimensional "syndrome" or "state" related to ageing first described in elderly patients

Characterized by:

Loss of reserve (energy, physical, cognitive, health) and the accumulation of "deficits"

(individually reversible but collectively insurmountable)

Consequence:

Heightened vulnerability or "state-at-risk" to adverse outcomes





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Singer J et al AATS 2016

Patterns of Functional Decline at End of Life



Definitions: Sudden death = cardiac arrest or trauma; Terminal illness = cancer; Organ failure = HF/COPD; Frailty: residence in nursing home



Lunney JR et al JAMA 2003

Trajectories of Disability in the Last Year of Life

Thomas M. Gill, M.D., Evelyne A. Gahbauer, M.D., M.P.H., Ling Han, M.D., Ph.D., and Heather G. Allore, Ph.D.

n=383 elderly decedents

UNIVERSITY



FRAILTY ~ most common condition leading to death

Gill TM et al. N Engl J Med 2010;362:1173-1180

Cumulative Deficit Model

CSHA captured 92 variables such as symptoms, signs, laboratory values, disease states and disabilities – collectively termed "deficits"

- Changes in everyday activities
- Head and neck problems
- Poor muscle tone in neck
- Bradykinesia, facial
- Problems getting dressed
- · Problems with bathing
- Problems carrying out personal grooming
- · Urinary incontinence
- Toileting problems
- Bulk difficulties
- Rectal problems
- Gastrointestinal problems Problems cooking
- Sucking problems
- Problems going out alone
- Impaired mobility
- Musculoskeletal problems
- Bradykinesia of the limbs
- Poor muscle tone in limbs
- · Poor limb coordination
- · Poor coordination, trunk
- Poor standing posture
- Irregular gait pattern
- Falls

- Mood problems
- Feeling sad, blue, depressed · History of depressed mood
- Tiredness all the time
- Depression (clinical impression)
- Sleep changes
- Restlessness
- Memory changes
 - · Short-term memory impairment
 - · Long-term memory impairment
 - · Changes in general mental functioning
 - · Onset of cognitive symptoms
 - Clouding or delirium
- - or loss

- History of Parkinson's disease

- Seizures, partial complex

- - History of stroke
 - History of diabetes mellitus

- Paranoid features
- · History relevant to cognitive impairment
- Family history relevant to cognitive
- impairment or loss
- Impaired vibration
- Tremor at rest
- Postural tremor
- Intention tremor
- · Family history of degenerative disease

- Seizures, generalized Syncope or blackouts
- Headache
- Cerebrovascular problems
- · Arterial hypertension
- Peripheral pulses
- Cardiac problems · Myocardial infarction
- - Arrhythmia
 - · Congestive heart failure Lung problems
 - Respiratory problems
 - · History of thyroid disease
 - Thyroid problems Skin problems
 - Malignant disease
 - Breast problems
 - Abdominal problems
 - · Presence of snout reflex
 - · Presence of the palmomental reflex
 - · Other medical history

Frailty Index (FI) = # deficits / total variables Frailty represents the "cumulative effects of individual deficits" **Reinforces the concept of** "physiologic/homeostatic reserve" and "biological gradation"

FI strongly correlates with risk of death and institutionalization



Jones et al J Am Ger Soc 2004

Physical Phenotype Model

Frailty Characteristics	CHS Measure		
Shrinking, weight loss (unintentional), sarcopenia	>10 lb lost unintentionally in prior 1 year		
Weakness	Grip strength: lowest 20% (by sex, BMI)		
Poor endurance, exhaustion, slowness	"exhaustion" (self-reported); walking time/15 ft: slowest 20% (by sex, height)		
Low activity	Kcal/week: lowest 20% (males <383 Kcal/wk; females <270 Kcal/wk)		

- 1. Shrinking: weight loss, unintentional, of ≥ 10 lbs in prior year)
- 2. Weakness: grip strength in lowest 20% at baseline (age/BMI)
- 3. Poor endurance/energy: self report of exhaustion (CES-D scale)
- 4. Slowness: slowest 20% at baseline for time to walk 15 ft (sex/height)
- 5. Low physical activity: lowest quintile of physical activity (gender)





Fried et al (Cardiovascular Health Study)_J Gerontol Biol Sci Med Sci 2001

Domains to Define, Measure and Operationalize

Domain	Operational Measures
General Health Status	Hospitalizations, global assessment of functioning scale, self-rated health
Physical Function	BADL, IADL, functional independence measure (FIM)
Cognitive Function	MMSE, Montreal Cognitive assessment, clock drawing test
Mobility	Short physical performance battery, gait speed, TGUG, chair rise, mobility aid
Strength	Grip strength, stair climb, subjective assessment of weakness
Energy	Fatigue severity scale, subjective assessment of exhaustion or fatigue
Nutritional Status	BMI, weight, albumin/prealbumin, mini-nutritional assessment, weight loss
Skeletal Muscle Mass	Anthropometry, bioelectrical impedance, MRI/CT/ultrasound
Mood	Geriatric depression scale, HADS, self-reported depression/anxiety
Social relations/support	Availability of social resources, subjective assessment of loneliness or isolation
Laboratory Markers	Inflammatory mediators (IL-6, IL-1, TNF, CRP), oxidized LDL, creatinine



Rajabali et al Can J Cardiol 2016; De Vries et al Ageing Research Reviews 2011

Methods to Screen and "Diagnose" Frailty

Method	Description	Items	Example	Evaluated in ICU
Frailty Index (FI)	Deficit accumulation	30-70	CSHA FI (70 items)	YES
Physical phenotype	\geq 3 physical features	3-5	Fried (CHS) criteria	YES
Physical performance measures	Single measure	1	Gait speed, grip strength, chair stand	NO
Judgement-Based tools	Global subjective assessment	1	Clinical Frailty Scale	YES
Multidimensional tools	Battery of assessments across domains	5-20	EFS, FRAIL, CAF, Groningen Frailty Indicator	NO
Sarcopenia	Imaging to assess skeletal muscle	1	CT scan psoas or rectus femoris	YES

Gold Standard: Comprehensive Geriatric Assessment (CGA)



Mortality in Relation to Frailty in Patients Admitted to a Specialized Geriatric Intensive Care Unit



Each 1% ↑ in FI associated with 11% ↑ in 30-day mortality (OR 1.11; 95% CI, 1.07-1.15)

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Zeng et al J Genrontol A Biol Sci Med Sci 2015

Frailty as a Predictor of Surgical Outcomes in Older Patients

n=594	Non-Frail (58.2%)	Intermediate (31.3%)	Frail* (10.4%)
Age (yr)	71 (67-94)	75 (65-92)	76 (65-94)
Female Sex (%)	67.6	52.7	41.9
Post-operative complications	1.0	2.06 (1.2-3.6)	2.54 (1.1-5.8)
Length of stay	1.0	1.49 (1.2-1.8)	1.69 (1.3-2.2)
Institutionalized	1.0	3.2 (1.0-9.9)	20.5 (5.5-76)



* FRAILTY defined by the physical phenotype criteria proposed by Fried

Makary et al JACS 2010

Prevalence and impact of frailty on mortality in elderly ICU patients: a prospective, multicenter, observational study





Le Maguet et al ICM 2014

A global clinical measure of fitness and frailty in elderly people

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.</p>

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.
Z. K. Rockwood et al. A global clinical measure of fitness and frailty in elderly people. CMAI 2005; 173:489-495.

© 2007-2009. Version 1.2. All rights reserved. Geriatric Medicine Research Darbouise University. Halfas: Canada. Remission granted to copy for research and educational purposes only. Inspiring Mindle Table 2: Cox proportional hazard ratios (HR) for time until death and until the requirement for institutional care

Factor	Death, HR (95% CI)	Entry into institution, HR (95% CI)
Age	1.08 (1.07–1.08)	1.15 (1.10–1.13)
Sex	0.83 (0.78-0.89)	1.38 (1.21–1.58)
Education level*	0.98 (0.97-0.99)	0.98 (0.97-0.99)
Modified Mini-Mental State Examination	0.84 (0.82–0.86)	0.65 (0.60-0.70)
Cumulative Illness Rating Scale	1.14 (1.11–1.17)	1.22 (1.16–1.27)
CSHA measuring tools Rules-based definition	1 17 (1 12 1 20)	
	1.17(1.13 - 1.20)	1.27 (1.19–1.35)
Frailty Index	1.26 (1.24–1.29)	1.56 (1.48–1.65)
Function Scale	1.16 (1.13–1.20)	1.29 (1.20–1.39)
Clinical Frailty Scale	1.30 (1.27–1.33)	1.46 (1.39–1.53)

CFS score and mathematically derived FI highly correlated (Pearson 0.80, p<0.01)



Available at: http://geriatricresearch.medicine.dal.ca/clinical_frailty_scale.htm

Rockwood et al CMAJ 2005

A global clinical measure of fitness and frailty in elderly people



Institutionalization



For each 1-category \uparrow in CFS score ~ 21.2% \uparrow death and 23.9% \uparrow institutionalization



Rockwood et al CMAJ 2005

Frailty – The "What" and "When"



ICU admission \rightarrow \rightarrow Identify Contributors \rightarrow \rightarrow Precision Recovery



Adapted from D. Rolfson





Bagshaw et al CMAJ 2014

	Group; no. of patient		no. (%) ients*	Association,
	Outcome	Frail <i>n</i> = 138	Not frail <i>n</i> = 283	difference in medians (p valuet)
↑ vulnerability	Adverse event‡	54 (39.1)	83 (29.3)	1.54 (1.01–2.37)
	Death			
	In ICU	16 (11.6)	27 (9.5)	1.37 (0.72–2.62)
↑ risk for death	In hospital	44 (31.9)	45 (15.9)	1.81 (1.09–3.01)
	Duration of stay, d, median (IQR)			
↑ time for recovery	In ICU	7 (4–13)	6 (3–10)	1 d (0.02)
	In hospital	30 (10–64)	18 (10–40)	12 d (0.02)
	Discharge disposition§	<i>n</i> = 91	<i>n</i> = 235	
	Home, living independently	20 (22.0)	104 (44.3)	0.35 (0.20–0.61)
	Home, living with help	33 (36.3)	58 (24.7)	1.67 (1.00–2.81)
	Other¶	38 (41.8)	73 (31.1)	1.51 (0.92–2.48)
functional impairment	Discharged newly dependent**	24 (70.6)	96 (51.6)	2.25 (1.03–4.89)
	Hospital readmission§	51 (56.0)	92 (39.1)	1.98 (1.22–3.23)

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Bagshaw et al CMAJ 2014

Survival 1-year after ICU Admission





Bagshaw et al CMAJ 2014

Secondary Analysis of BRAIN/MIND ICU Studies



RTA n=1040

Courtesy of Brummel N et al In Press

But...Does It Add Value?

- 1. Better informed triage decisions \sim
 - Regarding to suitability and likely benefit for ICU support

2. Guide and inform patient-centered decision-making \sim

- Regarding scope/duration of ICU support (i.e., time-limited trials)
- Regarding establishing/revisiting goals of care
- Regarding managing post-ICU survivorship expectations and experience (i.e., impact on HRQL, new disability, institutionalization, rehospitalization)



Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT Acute Skeletal Muscle Wasting in Critical Illness

63 critically ill mechanically ventilated patients (age 54.7; APACHE II 23.5)

Change in rectus femoris (RF) cross-sectional area (CSA) over 10 d Measures of muscle wasting in patients assessed by all 3 measures Α В on both day 1 and day 7 (n = 28)100 Percentage Change in CSA 50 -10 Loss, % -20 -50 -17.7% -30 -100 Ratio of Protein 2 g 10 **RF CSA** Fiber CSA to DNA Time From Admission, d



Puthucheary et al JAMA 2014

Timing of onset and burden of persistent critical illness in Australia and New Zealand: a retrospective, population-based, observational study



At ~ 10 days after ICU admission, acuity did not predict mortality better than antecedent characteristics (age, sex, comorbid disease) (variable transition point by case-mix and acuity)

Prevalence only 5.0% but accounted for 32.8% of ICU-bed days and only 46.5% returned home



And...How Else May It Add Value?

3. Transitions of care \sim

- Priorities/specialized needs for transition from ICU to ward setting
- Priorities/specialized needs for hospital to community (i.e., CGA)

4. Interventions (recognizing vulnerability)~

- Focused on maximizing physical recovery (i.e., minimizing avoidable disability)
- Focused on cognitive, psycho-social, and emotional recovery
- Focused on care-giver burden/experience
- Focused strategy towards palliation



Frailty – An Integrated Model



ICU admission $\rightarrow \rightarrow$ ICU Transition $\rightarrow \rightarrow$ Community Transition



Summary

- **Frailty** is a multi-dimensional syndrome contributing to vulnerability to adverse events:
 - can be measured in critically ill patients
 - is associated with \uparrow risk adverse events, death, re-hospitalization
 - is associated with \downarrow HRQL, new disability and \uparrow functional dependence
 - identifies a vulnerable population

• Frailty Assessment

- at ICU admission should focus on "case-finding" for <u>near-term</u> prognostication, guiding clinical care and decision-making and
- <u>after</u> ICU should start to focus more comprehensive assessments, care transitions and specialist referral



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Questions?

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