





Belgian Inter-university Course in Geriatric Medecine

16/11/2018

Frailty

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« As geriatricians, our concern is the frail old patient »

Art. 3. Het zorgprogramma voor de geriatrische patiënt richt zich tot de populatie geriatrische patiënten van gemiddeld ouder dan 75 jaar, welke een specifieke aanpak behoeft om verschillende van de hierna vermelde redenen:

Art. 3. Le programme de soins pour le patient gériatrique s'adresse à la population de patients gériatriques ayant une moyenne d'âge de plus de 75 ans et qui requiert une approche spécifique pour plusieurs des raisons suivantes :

1° fragiliteit en beperkte homoïostase;

1° fragilité et homéostasie réduite ;

- · What is Frailty?
- Why identifying frail older people?
- How to identify frail older people?

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- 5. Clinical pictures
 - Cardio-geriatrics
 - Emergency
 - Oncology







2. Frailty and Functional decline and comorbidity?
Similarities- differences

Frailty, disability and comorbidity

• 3 different terms often used to described the « geriatric » population, a vulnerable, older population who required enhanced care.

But

These are distinct clinical entities.

Journal of Gerontology: MEDICAL SCIENCES 2004, Vol. 59, No. 3, 255-263 Copyright 2004 by The Gerontological Society of America

Review Article

Untangling the Concepts of Disability, Frailty, and Comorbidity: Implications for Improved Targeting and Care

Linda P. Fried, 1,2,3 Luigi Ferrucci, 3 Jonathan Darer, 4 Jeff D. Williamson, 5 and Gerard Anderson 2

Disability

Def: difficulty or dependency in carrying out activities in daily living

Physical disability is measured by performance test or assessed by self-report tools.

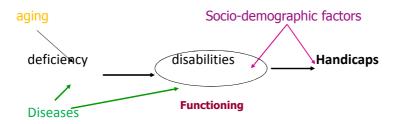
basic or instrumental ADL Mobility

Disability

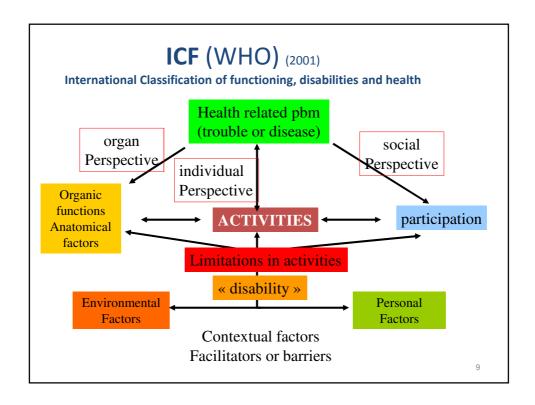
- Prevalence
- - After 65y, 1/7 persons in US
- Hébert et al. Incidence of FD and improvement in a community-dwelling very elderly population. Am J Epidemiol 1997;145:935-44
- Longitudinal, community-dwelling, 572 people 75 y and older, SMAF*
- Stable, no previous FD
 - Incidence of loss of functions :11,9 %/y
 - 6,2% improve
 - mortality 3,2%
- *Hébert R., Carrier R, Bilodeau A. Le Système de Mesure de l' Autonomie Fonctionnelle (SMAF). Rev Geriatr 1988 17;161-7.

Disability: conceptual background

Disability and functional decline 80'



Travaux de Wood , Verbrugge et Jette (1984), CIH (Classification Int. des Déficiences, Incapacités et Handicaps) et CIF (Classification Int. du Fonctionnement) (OMS, 1980 et 2001)



Theoretical classification of frailty factors, and markers (according to ICF WHO)

Primary contributors

- Body structure and organ function (impairments)
 - Muscle performance, nutrition
 - Cognitive performances, and sensory organs performance
 - Renal function, Hemoglobin level
- Diseases
 - Comorbidities and severity of chronic diseases
 - Medications (number, sedatives)
 - Health care utilization
- Contextual factors
 - Person's assets (AGE, health perception, income, education...)
 - Social support, family
 - Environment and home acomodation

Theoretical classification of frailty factors, and markers (according to ICF WHO)

Consequences of frailty as secondary contributors

- Limitations of individual's activities (disabilities)
 - Loss in iADL's performance
 - Loss of ADL's performance
- Health problems
 - Geriatric syndromes (falls, delirium)
 - Further health care utilization (hospital readmissions)
- Restrictions in individual's participation
 - Dependency
 - Institutionalization

Disability

2 different forms/pathways

 50% of disabilities develop chronically, progressively, in association with the development of chronic diseases and comorbidities and frailty.

?

- 50% develop acutely in association with an acute disease ?

Risk factors

for functional decline in community-living elderly people. Meta analysis

STUCK, Social Science and Medicine 1999;48:445-469

Cognitive impairment Depression

Comorbidity BMI

· Lower extremity f. limitation Low fr. of social contact

Low level of physical activity No alcohol use (vs moderate)

Poor self perceived health Smoking

Visual impairment

Factors often assessed in frailty measurements

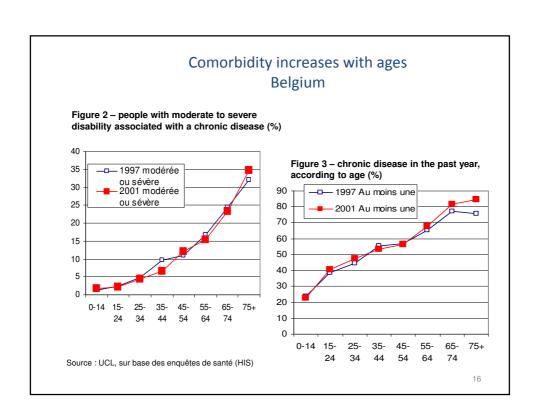
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Disability

- Disability is also a adverse health outcome in itself
- Mobility disability leads to dependence in IADL
- IADL dependence is a risk factor for dependence in bADL after a hospitalization, whatever the reasons for hospitalization
- Disability in ADL is a risk factor for increased mortality , institutionnalisation

Comorbidity

- Def: the concurrent presence of two or more medically diagnosed diseases in the same individual, with the diagnosis of each contributing disease based on established, widely recognized criteria
- But also a broader def: comorbidity involves interactions between any two conditions, even of clinical or subclinical
- Not only a cumulative effects of two conditions (at least) but their synergetic interactions



comorbidity

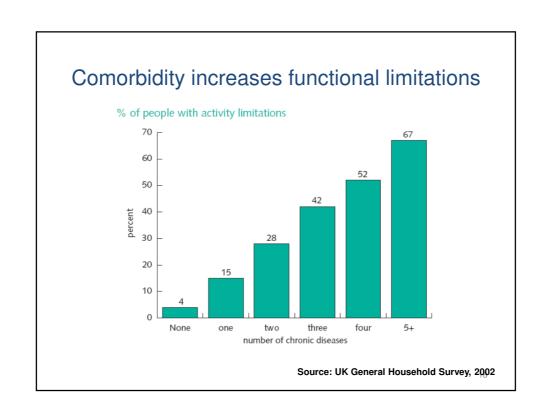
• At 65 y

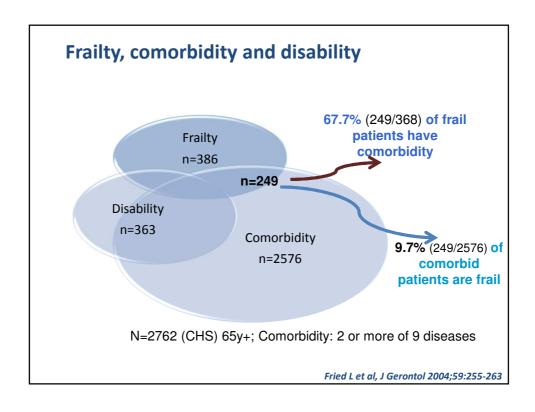
- HTA: 36%

Heart disease: 27%Diabetes: 10-15%

- Stroke: 6%

- Dementia 10,1% (Eur stat)











3. Physiopathologic basis of Frailty

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What does «frailty» mean for older people?

Puts MTE et al, J of Aging Studies, 2009

Qualitative study in frail and non frail older people (LASA Longitudinal Aging Study Amsterdam)

- Frailty?
 - Being in poor health
 - Having walking difficulties
 - Feeling down
 - Being anxious
 - Having few social contacts
 - Not being able to do things ones like to do
- Men described in more details the physical dimension
- Women elaborated in more depth the social and psy dimensions

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Definition

adapted from Studenski JAGS 2004;62:1560-66 and Ferrucci J Endocrinol Invest. 2002;25:10-5

Age-related alteration in physiology and pathology that leads to vulnerability with loss of organ system reserve, limited capacity to respond to internal and environmental stresses, unstable homeostasis

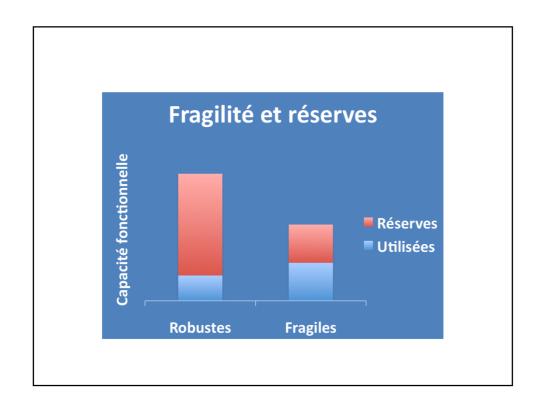
Leading to **poor medical and functional outcomes.**

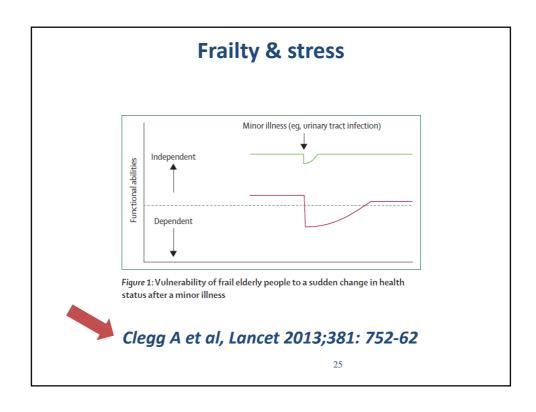
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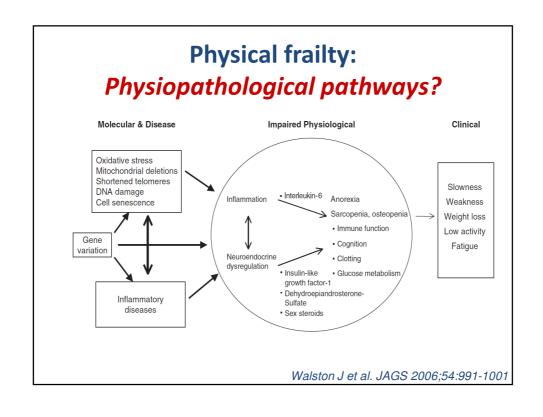
Main elements in the definition of frailty

- Biophysiological base that is age related
- Multiple system impairments
- Reduced reserve with diminished adaptative response
- Vulnerability to stressors and to challenges of the environment
- Increased risk for adverse outcomes
- Instability and change over time

Rockwood K, Drugs Aging 2000;17:295-302







Hypotheses for specific physiopathology of frailty → role of biomarkers?

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Biomarkers

- Highly sensitive and specific indicators of disease pathways
 - Used as substitutes for outcomes in clinical trials when evidence indicates that they predict clinical risk or benefit.
- Definition
 - "A characteristic that is objectively measured as an indicator of normal or pathogenic biological processes, or pharmacologic responses to a therapeutic intervention"
- Exemple: Chol & statine

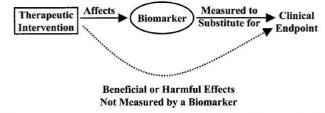
Biomarkers Definitions working Group, Clin Phar Ther 2001 Slide inspirited from Pr JM Degryse

Biological markers

- Many applications in disease detection and monitoring:
 - Use as a diagnostic tool
 For the identification of those patients with a disease or abnormal condition
 - Use as a tool for staging or extension of a disease
 - Use as an indicator of disease prognosis
 - Use for prediction and monitoring of clinical response to an intervention.

Definitions

- Clinical endpoint :
 - A characteristic or variable that reflects how a patient feels, functions, or survives.
 - They reflect the reflect of a therapeutic intervention.
- Surrogate endpoint :
 - A biomarker that is intended to substitute for a clinical endpoint. A surrogate endpoint is expected to predict clinical benefit or harm based on epidemiologic, or nathophysiologic evidence

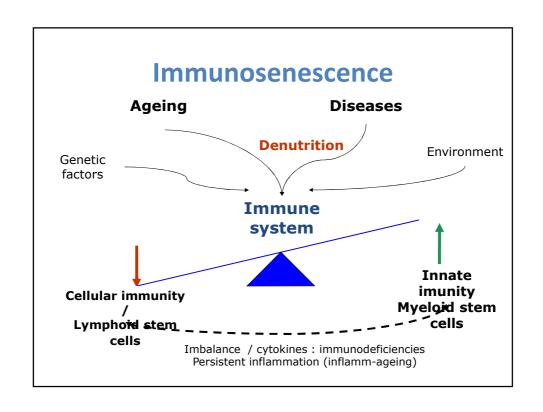


Biomarkers Definitions working Group 2010

Biomarqueurs

Immunosenescence	Chronic inflammation	++
	Cellular immunity	+
Neurohormonal dysregulation	IGF-1, DHEA	+
Replicative senescence	Telomer lenght	?
Oxydative stress	Anomalies RNAm, DNA	?
Proteic glycations	Advanced Glucation Endproducts	?

Fried & Walston, Principles of Geriatric Medicine (...), Hazzard 1999



Inflammation & ageing « Inflamm-ageing »

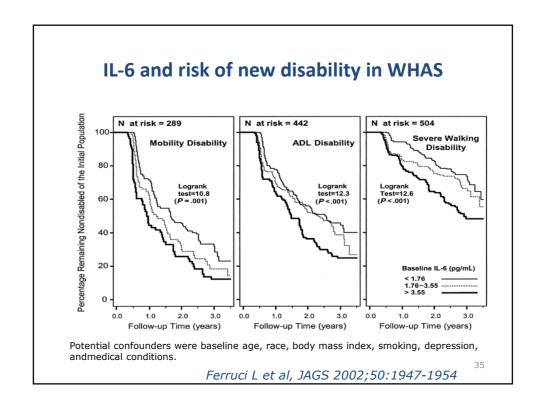
- C. Franceschi (2000)
- Imbalance between inflammatory and antiinflammator
- Etiology
 - Immunosenescence
 - Hormonal dysregulation (E2)...
 - Others: polymorphisms, tabac, obésité, HT,...
 - Role of CMV?

Pawelec et al., Immunol Rev 2005;205:257-268 Ershler W et Keller E, Annu Rev Med 2000;51-245-270 Schmaltz HN et al (Fried LP), JAGS 2005;53:747-754

II-6: results from the WHAS

- Women's Health & Aging Study
 - 65y+ recruted among the one-third most disabled women (community-dwelling)
 - Randomly sampled form the Medicare beneficiaries in Baltimore
 - Difficulty performing 1 or more tasks in at least 2 of the following 4 domains of functioning:
 - mobility/exercise tolerance, upper extremity abilities, basic self-care, and higher functioning tasks of independent living
- MMSE >18/30
- 5316 → 1002 inclusion, 620 with blood samples

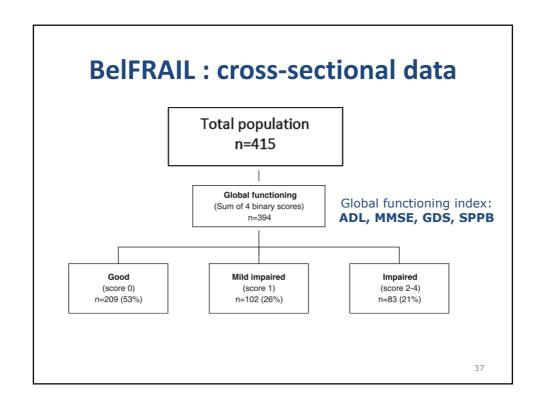
Ferruci L et al, JAGS 2002;50:1947-1954

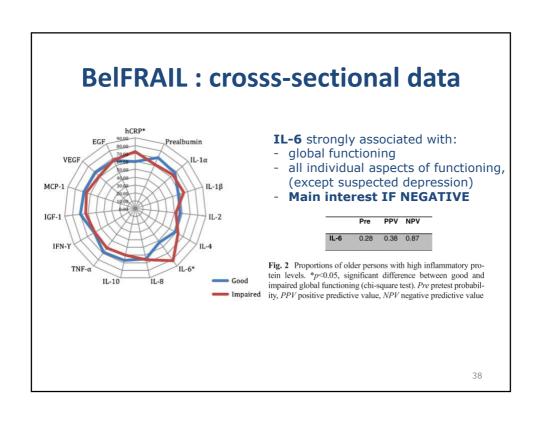


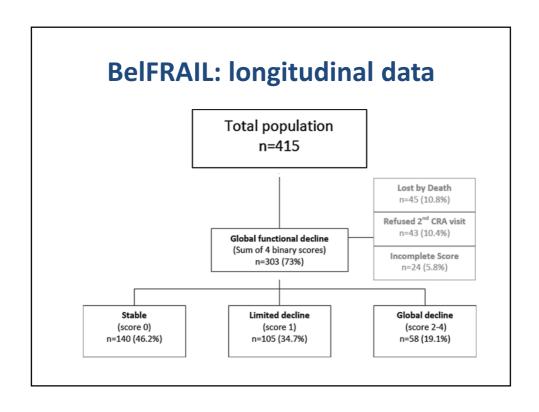


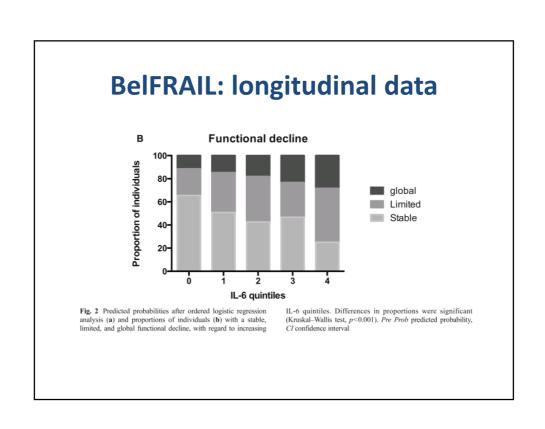
- Prospective population-based study,
- 80y+, selection by GPs
 - Exclusion criteria: severe dementia, palliative care and medical emergency
- Clinical and biological assessment
- N=567
- Cross-sectional and longitudinal analyses
 - Follow-up

Vaes B et al, BMC Geriatr 10:39. doi:10.1186/1471-2318-10-39







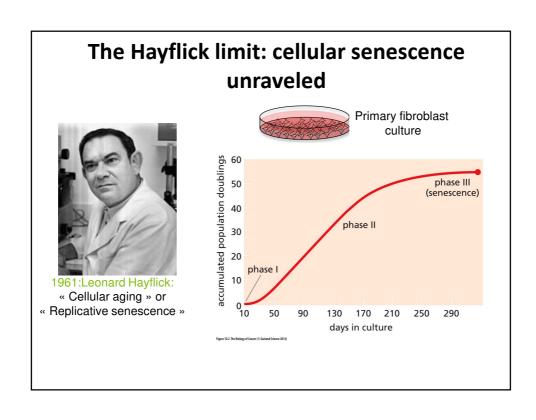


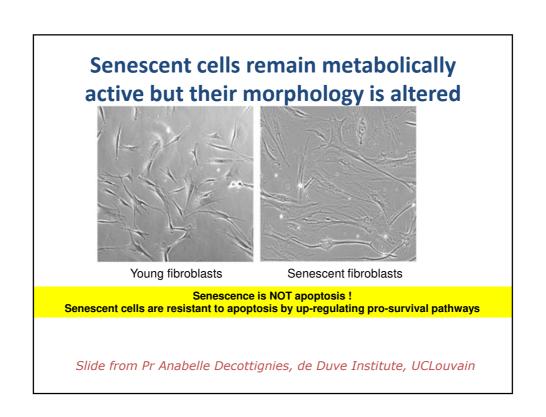
Other markers

- Inflammation: (us)CRP, TNF-a
- Coagulation: D-Dimers, fibrinogen
- IGF-1
- Telomer length
- Association:
 - Cross-sectional
 - Longitudinal (mortality)

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Telomer length & frailty telomères Woo et al, Mech Ageing Dev 129(11): 642-648.





Perspectives of future therapies?

- -Activation of endogenous telomerase gene expression:
- androgen therapy (e.g. Danazol)
 -Transient activation of exogenous telomerase:

AAVs



-Gene therapy:

From reprogrammed stem cells of the patient

-Targeting cellular senescence:

SASP modulators, senolytics, immune clearance

Slide from Pr Anabelle Decottignies, de Duve Institute, UCLouvain

Biogerontology https://doi.org/10.1007/s10522-018-9749-5

REVIEW ARTICLE

A review of telomere length in sarcopenia and frailty

Maria Lorenzi 🕝 · Stefano Bonassi · Teresa Lorenzi · Silvia Giovannini · Roberto Bernabei · Graziano Onder

Experimental Gerontology 106 (2018) 16-20



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Review

The association between telomere length and frailty: A systematic review and meta-analysis

Jianghua Zhou a,b,1 , Jiang Wang c,1 , Yanjiao Shen d , Ying Yang a,b , Pan Huange, Shanping Chen a,b , Chuan Zou a,b , Birong Dong f,g,*

The association between telomere length and frailty: A systematic review and meta-analysis

Jianghua Zhou^{a,b,1}, Jiang Wang^{c,1}, Yanjiao Shen^d, Ying Yang^{a,b}, Pan Huang^e, Shanping Chen^{a,b}, Chuan Zou^{a,b}, Birong Dong^{f,g,*}

Author, year	Design	Male (%)	n	Frailty preval (%)	Frailty def	Quality (NOS)
Marzetti, 2014	cs	40,8	142	51,7	Index	7
Yu 2015	cs	100	976	7,3	Fried	8
Yu, 2015	CS	0	1030	5,4	Fried	8
Collerton, 2012	CS	61,7	811	21,6	Fried	9
Pathai, 2013	СС	25	256	13,3	Fried	7
Brault 2014	cs	-	53	28,3	mFried	6

CS: cross-sectional; CC: case-control; TL: qPCR from leukocytes; NOS: Newcastle_Ottaxa Scale – assessment of quality of non-randomized studies in meta-analyses: http://www.ohri.ca/programs/clinical_epidemiology/oxford.asp

Zhou et al, Exp Gerontol 2018;106:16-20

The association between telomere length and frailty: A systematic review and meta-analysis

Jianghua Zhou a,b,1 , Jiang Wang c,1 , Yanjiao Shen d , Ying Yang a,b , Pan Huang e , Shanping Chen a,b , Chuan Zou a,b , Birong Dong f,g,*

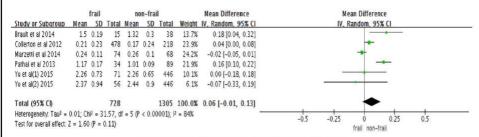


Fig. 2. Meta-analysis of the association between telomere length and frailty.

- No significant association between frailty and telomere length
- · Association TL and grip strength

Zhou et al, Exp Gerontol 2018;106:16-20

Maria Lorenzi : Stefano Bonassi · Teresa Lorenzi · Silvia Giovannini · Roberto Bernabei · Graziano Onder				
SARCOPENIA	Design	n	Main findings	
Marzetti, 2014	cs	142	TL↔ sarcop NOT GripS et GaitSpeed	
Woo, 2014	Prospect (5y)	2006	$TL \longleftrightarrow Grip \ S \ NOT \ sarcop \ or \ other \ phys \ perf$	
Batsis, 2017	cs	2672	TL NOT ↔ sarcop	
Frailty	Design	n	Main findings	
Woo, 2014	Prospect (5y)	2006*		
Collerton, 2012	CS	845*°		
Marzetti, 2014	CS	142*°	TL NOT \leftrightarrow frailty	
Saum, 2014	CS	3537°	·	
Yu, 2015	Prospect (5y)	2006*		
Breitling, 2016	cs	1820°	DNA methyl ↔ frailty NOT TL	

Take home message Physiopathology of frailty & biomarkers

- Physiopathology of frailty
 - Still debated
 - Frailty & ageing?
 - ...but lack of evidence!
- 2 differents positions
 - Biomarkers: ORGAN / CELLULE
 - Frailty: multidimensional syndrom & global assessment
 - Impact of cellular mechanisms PLUS
 - Proeminent environmental impact
 - → Causal or mechanistic association with frailty unclear

Woo, Mech Ageing Dev 2008; 129:642

Take home message Physiopathology of frailty & biomarkers

- Limits of biomarker measurements
 - Dynamics
 - Design (longitudinal), population and sample size
 - Measurement methods
 - Survival bias
- Perspectives
 - Recognize biomarkers and their limits (epiphenomen or causal relationship?)
 - Combination of biomarkers?
 - Evaluation of biomarkers / accuracy

Woo, Mech Ageing Dev 2008; 129:642







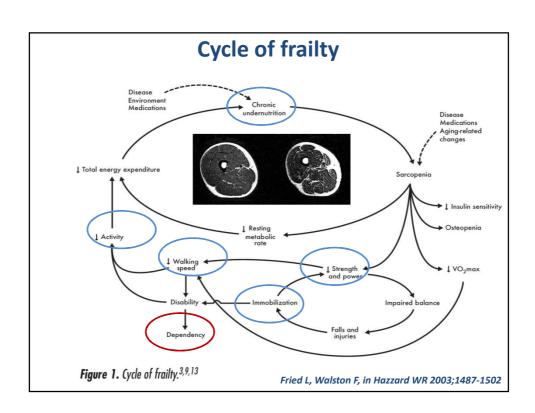
4. Is frailty a useful concept from a clinical point of view?

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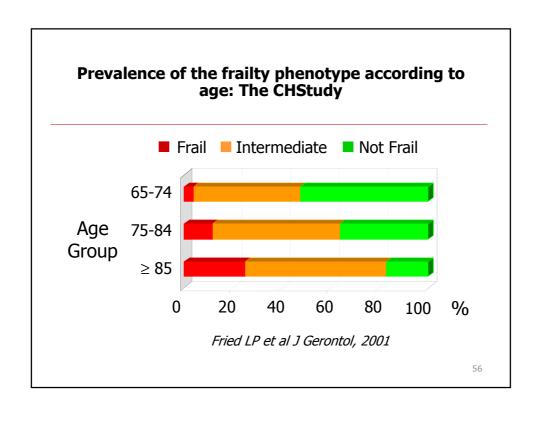
Assessment of frailty

- 2 majors approachs...
 - Physical frailty (« Ph-railty »)
 - Phenotype of frailty
 - Fried L
 - Functional fraily (« F-railty »)
 - Index of frailty
 - Rockwood
- ...Same outcomes

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Frailty phenotype Lower quintile at GRIP STRENGTH Lower quintile at WALKING SPEED WEIGHT LOSS more than 4,5 kg past year EXHAUSTION criteria Bottom quartile for PHYSICAL ACTIVITY > =3: frail 1-2: intermediate Cardiovascular Health Study Fried et al, J Gerontol Med Sc 2001;56A, M146–M156



Predictive validity of physical frailty

3 years (%)	Robusts (n=2469)	Intermediate (n=2480)	Frail (n=368)
Death	3	7	18
1st hospital.	33	43	59
1st fall	15	19	28
Worsening ADL disability	8	20	35
Worsening mobility disability	23	40	51

p<0.001 for all

Fried LP et al J Gerontol, 2001

Frailty index







- List of 20-30-70 (!) deficits (present =1, absent = 0)
- Multiple domains (function, cognitive, psychological, mobility, morbidity etc)
- Rapport deficits / nb max of deficits
- Proposed threshold: 0.25

Rockwood et al, CMAJ 2005;193:489-95

Table 1 46 deficits included in frailty index. Converbidities • Stroke • Thyroid condition • Cancer • Heart attack • Heart disease • Ever had high blood pressure • Onieroperosis • Diabetes • Arthrifs • Ever had broken hip • Cataract operation • Weakfailing kidneys • Difficulty using fork and knife • Difficulty using fork and knife • Difficulty standing up from armiess chair • Difficulty standing for long periods of time • Difficulty praparing meals • Difficulty piraping holding small objects • Difficulty lifting or carrying • Difficulty attending social event • Signs/symptoms • Heart atte at rest • Syntolic blood pressure • Sough regularly • Leaked/lost control or urine • Ceneral hearing • Canfusion or inability to remember things • Canfusion or

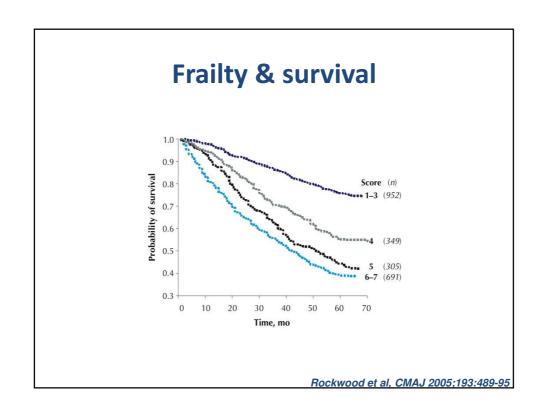
Levels of frailty

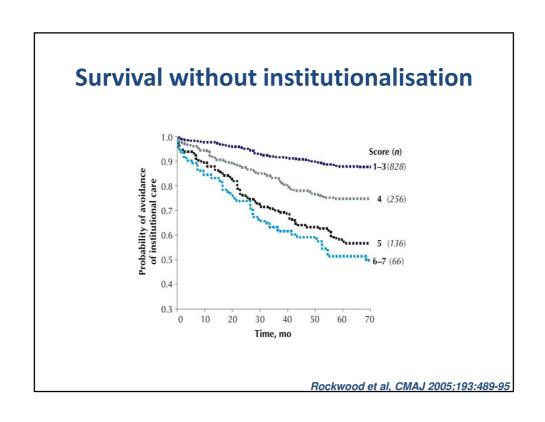
Box 1: The CSHA Clinical Frailty Scale

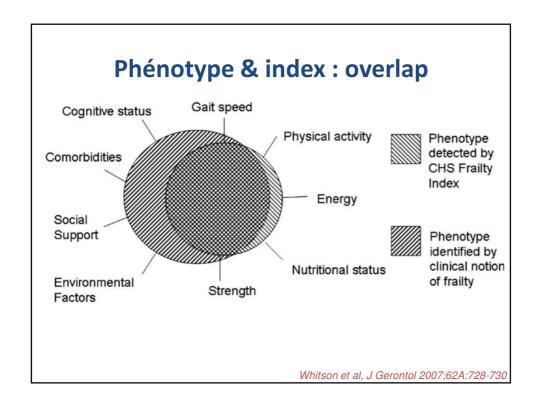
- 1 Very fit robust, active, energetic, well motivated and fit; these people commonly exercise regularly and are in the most fit group for their age
- 2 $\ensuremath{\textit{Well}}$ without active disease, but less fit than people in category 1
- 3 Well, with treated comorbid disease disease symptoms are well controlled compared with those in category 4
- 4 Apparently vulnerable although not frankly dependent, these people commonly complain of being "slowed up" or have disease symptoms
- 5 Mildly frail with limited dependence on others for instrumental activities of daily living
- 6 Moderately frail help is needed with both instrumental and non-instrumental activities of daily living
- 7 Severely frail completely dependent on others for the activities of daily living, or terminally ill

Note: CSHA = Canadian Study of Health and Aging.

Rockwood et al, CMAJ 2005;193:489-95







2 definitions, same outcomes

- Consensus on outcomes of frailty
 - Functional decline (disability, dependance)
 - Geriatric syndromes (big I 's)
 - Health care utilization (home care, H adm and readm.)
 - Institutionalisation
 - Mortality
- Is more predictive for clinical outcomes than diagnosis per se

Winograd, JAGS 2001;39:778-784

F phenotype or F Index?

• The 2 concepts are complementary

(Cesari et al, Age and Aging, 2014)

- « The frailty phenotype may be more suitable for an immediate identification of non-disabled elders at risk of negative events. »
- « The Frailty Index may summarise the results of a comprehensive geriatric assessment providing a marker of deficits accumulation. »

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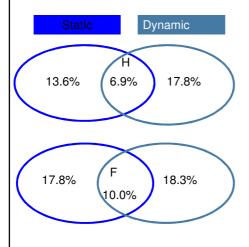
FRAILTY IN A DYNAMIC PERSPECTIVE...

Static or dynamic assessment?

- Static measure of frailty:
 - « Picture »
- Dynamic measure of frailty:
 - Evolution between 2 assessments
 - Identify a decline

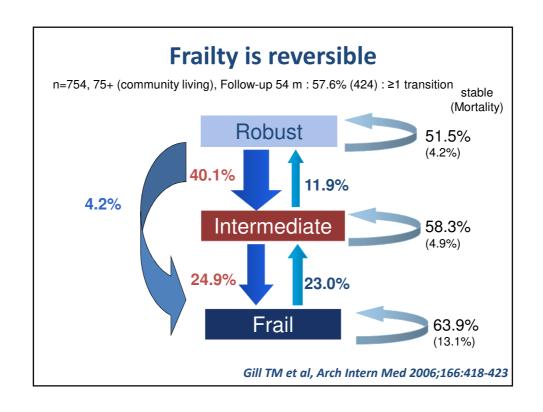
Puts MTE et al. JAGS 2005;53:40-47

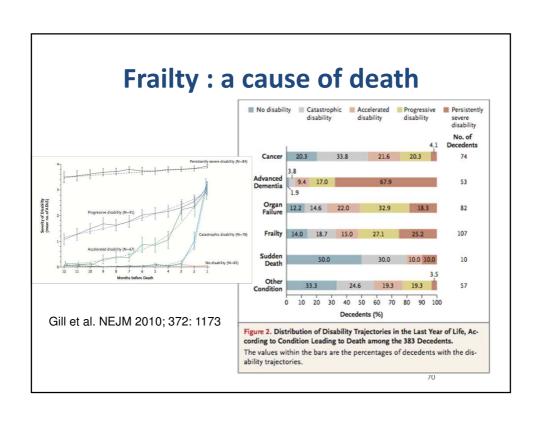
Static or dynamic assessment?



- 9 markers of frailty
 - BMI, VEMS, cognition, sens, continence, locus of control, depression, physical activity
- N=2257 (Amsterdam)
- First assessment (t1) + 3 years (t2)
- Static frailty: 3+ markers at t1
- Dynamic frailty: 3+ changes between t1 & t2

Puts MTE et al. JAGS 2005;53:40-47











Review

Physical Resilience in Older Adults: Systematic Review and Development of an Emerging Construct

Heather E. Whitson, ^{12,3,4} Wei Duan-Porter, ^{1,5} Kenneth E. Schmader, ^{1,2,3} Miriam C. Morey, ^{1,2,3} Harvey J. Cohen, ^{1,2,3} and Cathleen S. Colón-Emeric ^{1,2,3}

1. Change of paradigm

- Point of view focusing on the process leading from frailty to disabilities
- Predominant view of frailty: based on adverse outcomes
- Evolution in classic WHO health definition
 - Dynamic concept of health, linked to resilience and ability to cope

Boers et Cruz-Jentoft, Calcif Tissue Int (2015) 97:429-431

New perspectives

- Health is the resilience or capacity to cope, and to maintain and restore one's integrity, equilibrium, and sense of wellbeing in three domains: physical, mental, and social.
- Frailty is the weakening of (health; see above).



Boers et Cruz-Jentoft, Calcif Tissue Int (2015) 97:429–431 Invitational Conference 'Is health a state or an ability? 'Report of the meeting December 10–11, 2009. www.gezondheidsraad.nl/sites/default/files/bijlage%20A1004_1.pdf

2. Physical and cognitive frailty

 sarcopenia has been proposed to represent the biological substrate of the physical function impairment that characterizes physical frailty (PF)

Calvani R et al, Aging Clin Exp Res 2017;29:29-34

COGNITIVE FRAILTY: RATIONAL AND DEFINITION FROM AN (I.A.N.A./I.A.G.G.) INTERNATIONAL CONSENSUS GROUP

E. KELAIDITI¹, M. CESARI¹²³, M. CANEVELLI¹⁴, G. ABELLAN VAN KAN¹², P.-J. OUSSET¹, S. GILLETTE-GUYONNET¹², P. RITZ²⁵, F. DUVEAU⁶, M.E. SOTO¹², V. PROVENCHER⁷, F. NOURHASHEMI¹², A. SALVA⁸, P. ROBERT⁶, S. ANDRIEU^{12,3,10}, Y. ROLLAND¹², J. TOUCHON¹¹, J.L. FITTEN¹², B. VELLAS^{1,2,3}

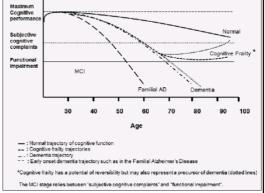
- Main stream based on physical frailty
- Frailty and cognitive impairment : studied separately
- Frailty: associated with low cognitive performance over time in older individuals with and without dementia
- CF : precursor of dementia?
 - PreMCI phase? Reversibility?
 - Lack of definitions & measures....

Kelaiditi E, et al, JNHA 2013;17:9

Cognitive frailty

- Heterogeneous clinical manifestation
- Key factors:
 - Presence of physical frailty and cognitive impairment (CDR=0.5);
 - Exclusion of concurrent AD dementia or other dementias.

Figure 2
Different trajectories of cognitive function according to specific conditions



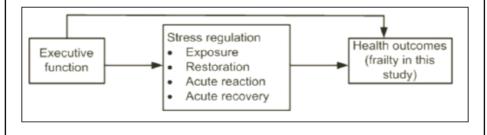
Kelaiditi E, et al, JNHA 2013;17:9

STRESS REGULATION AS A LINK BETWEEN EXECUTIVE FUNCTION AND PRE-FRAILTY IN OLDER ADULTS

R.A. ROILAND^{1*}, F. LIN^{2*}, C. PHELAN¹, B.P. CHAPMAN³

Figure 1

Conceptual Model (Developed based on Williams et al., 2009)



Rowland et al, JNHA 2015; 19:8

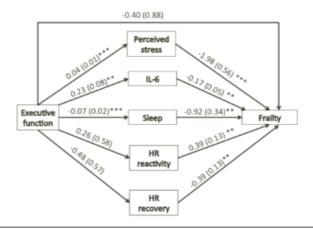
3. Role of executive function (EF)

- EF: cognitive processes involved in problem solving and the adjustment of behaviors in response to stress,
 - Important factor to consider when examining stress regulation.
- Stress exposure and restoration reflect chronic processes, whereas stress reactivity and recovery reflect more acute processes.

Stress regulation, executive function and frailty

- Design: Cross-sectional.
- Participants: 690 community-dwelling older adults ≥ 50 years of age.
- Measurements:
 - Pre-frailty: modified form of the Fried Frailty measure.
 - EF was assessed via telephone- based neurocognitive assessments.
 - Indicators of stress regulation :
 - stress exposure (measured by perceived stress),
 - reactivity and recovery (measured by heart rate) and
 - restoration (measured by serum IL-6 and sleep quality).

Figure 2 Statistical model of relationships between EF, Indicators of Stress Regulation, and Frailty Status (i.e., Pre-Frail or Non-Frail



Note. Parameter estimates (standard error) are presented. Age, gender, education, anti-hypertensives, anti-depressants, corticosteroids, smoking, and time lag between P3 and P4 were controlled. * p < .05, ** p < .01, **** p < .001.

4. Interventions: recommandations for further research

- Outcomes
- · Quality of studies
- Robust & validated measurments
- Representativity of participants
- Roles of caregivers

2 recent systematic reviews....

Age and Ageing 2017; **46:** 383–392 doi: 10.1093/ageing/afw247 Published electronically 7 January 2017 © The Author 2017. Published by Oxford University Press on behalf of the British Geriatrics Society.

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properly cited. For commercial re-use, please contact journals.permissions@oup.com

REVIEW

Interventions to prevent or reduce the level of frailty in community-dwelling older adults: a scoping review of the literature and international policies

Martine T. E Puts $^{\rm I}$, Samar Toubasi $^{\rm I}$, Melissa K. Andrew $^{\rm 2}$, Maureen C. Ashe $^{\rm 3.4}$, Jenny Ploeg $^{\rm 5}$, Esther Atkinson $^{\rm 6}$, Ana Patricia Ayala $^{\rm 7}$, Angelique Roy $^{\rm 8}$, Miriam Rodríguez Monforte $^{\rm I}$, Howard Bergman $^{\rm 9}$, Kathy McGilton $^{\rm 8}$

Results: fourteen studies were included: 12 randomised controlled trials and 2 cohort studies (mean number of participants 260 (range 51–610)), with most research conducted in USA and Japan. The study quality was moderate to good. The interventions included physical activity; physical activity combined with nutrition; physical activity plus nutrition plus memory training; home modifications; prehabilitation (physical therapy plus exercise plus home modifications) and comprehensive geriatric assessment (CGA). Our review showed that the interventions that significantly reduced the number of frailty markers present or the prevalence of frailty included the physical activity interventions (all types and combinations), and prehabilitation. The CGA studies had mixed findings.

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Interventions to limit frailty

Clinical Interventions in Aging

Dovepress



REVIEW

Effects of multi-domain interventions in (pre)frail elderly on frailty, functional, and cognitive status: a systematic review

This article was published in the following Dove Press journal: Clinical Interventions in Aging 24 May 2017 Number of times this article has been viewed

Lenore Dedeyne¹ Mieke Deschodt²⁻⁴ Sabine Verschueren⁵ Jos Tournoy^{1,3} Evelien Gielen^{1,3}

Background: Frailty is an aging syndrome caused by exceeding a threshold of decline across multiple organ systems leading to a decreased resistance to stressors. Treatment for frailty focuses on multi-domain interventions to target multiple affected functions in order to decrease the adverse outcomes of frailty. No systematic reviews on the effectiveness of multi-domain interventions exist in a well-defined frail population.

Conclusion: Evidence of beneficial effects of multi-domain compared to mono-domain interventions is limited but increasing. Additional studies are needed, focusing on a well-defined frail population and with specific attention to the design and the individual contribution of mono-domain interventions. This will contribute to the development of more effective interventions for frail elderly.

Multidomains interventions in (pre)frail elderly

- Multi-domain interventions improve frailty characteristics and physical functioning more effectively than mono-domain interventions
- Inconsistent effects on functional abilities, falls, and psychosocial outcomes?
- Physical exercise seems tp play an essential role in the multi-domain intervention

Dedeyne et al, Clin Interventions in Aging 2017

Multidomains interventions in (pre)frail elderly: perspectives

- Effects on cognition, social involvement, or some functional outcomes?
- · Optimal duration of intervention
- Core outcome set
 - 1) frailty status, score;
 - 2) muscle outcomes (mass and strength);
 - 3) physical outcomes;
 - 4) cognition, social outcomes, and/or psychological well-being.
- Heterogeneity of populations and frailty tools
- Understanding the contribution of each mono-domain intervention to optimize and prioritize the frailty syndrome management?
- Optimal moment for intervention?

Dedeyne et al, Clin Interventions in Aging 2017

Why to assess frailty?

- Older people are a heterogeneous group
 - Fit ⇔ Frail ⇔ dependent
- Frailty is an elevated state of risk...
 - ψ ability to deal with stressor events
 - Strong predictor of several adverse events
- · Frailty is reversible
 - Early diagnosis ⇔ preventing and treating
 - · Treatment decisions
- Intervention to maintain homeostasis, decrease consequence of frailty

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Clinical Public health Target Prevent Direct Decision Plan Research Include Intervene







5. Clinical pictures

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Objectives of this last part

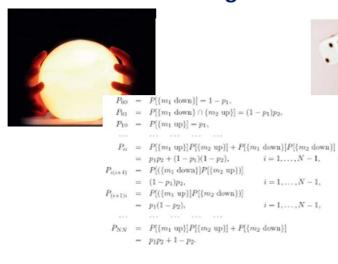
To show:

How frailty screening in different contexts of care help to identify a population of older patients who will benefit from geriatric intervention, in order to limit functional decline, geriatric syndroms,

How frailty screening followed by CGA is useful before a surgical intervention, a cancer treatment, ...

- Help to identify modifiable risk factors: polymedication, denutrition, delirium
- Better appreciation of risk/benefit balance
- Improve information to patients, carers.





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Two step approaches

- To target
 - The population who will benefit from interventions
 - Screening tools
- To assess
 - CGA

Frailty in specific settings

- Primary care
- Emergency department
- Oncology department
- Surgical department
- Cardiovascular department

9:

PRIMARY CARE

Robert's story

Robert -a 87-year old man- is sent to the emergency department where he is found to have suffered a myocardial infarction, developed hyponatremia, and become delirious. He is admitted to your geriatric ward.

Robert's story

During the past 2 days, Robert develops confusion and experiences repeated falls. He now needs assistance from 2 people to transfer and requires assistance with feeding and toileting. His wife is no longer able to manage his care needs.

Robert's story

Six month ago, Robert visited his GP because of worsening urinary incontinence and falls:

- Slow gait speed;
- 6-month weight loss (5%) and reduced muscle mass
- Normal physical examination
- Magnetic resonance imaging: central lumbar spinal stenosis.
- By urologist: Urinary retention after voiding treated by urinary catheter.
- During the following 2 days, Robert develops confusion and experiences repeated falls.

Robert's story

His wife tell you that Robert is a retired engineer, very nice with her and his children, but is slowing down progressively since 2 years. Previously physically active, he was walking more and more slowly and suffered recently from several recent falls. Because of resulted "lost strenght" and fear of falling, he was used a walker or cane to walk short distances. His reduced appetite has resulted in a 9-kg unintentional weight loss over theses 2 years. She does not describe any cognitive problem. He was cognitively intact and did not have any complain.

He has had urinary incontinence for several years, attributed to radiotherapy for prostate cancer in 1996. His medical history includes a stroke in 2003 involving mild left-sided hemiparesis, hypertension, and hyperlipidemia. He lives independently with his elderly wife in a 2-storey house. Robert takes the following medications: 20 mg of simvastatin once daily, 5 mg of ramipril once daily, 75 mg of clopidogrel once daily, and vitamin D.

- How common is frailty in GP practice?
- How relevant is frailty to GP practice?
- What can do a GP in his/her practice?
- When to refer to a specialist?

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Screening tools for frailty in primary care

- GPs are increasingly confronted with frail patients
 - 7% 65+; 25-40% 80+
 - Importance to distinguish normal ageing from frailty, potentially reversible/prevention of adverse outcomes
- GPs require a simple screening tool for frailty
 - Easy, reliable and inexpensive tool? + practical feasibility
- When screening positive: medical review
 - Medical evaluation (comorbidity, other underlying conditions) and medication review
 - Intervention related to malnutrition and physical activity (exercise program): cf. supra
 - Referral to geriatrician (...) and allied health professional
 - Advance Care Planning

Tool/study	No. items	Mode of administrat°	Language	Administrat° duration	Reference G assessment
Screening letter	9 simple items	Self-administered questionnaire	English	-	CGA by geriatrician
Sherbrooke postal questionnaire	6 simple items	Self-administered questionnaire	French	-	SMAF scale by nurse in the home
Functional assessment screening package	16 simple items or measures	Non-medical staff	English	8–12 min	CGA by geriatrician
Screening instrument	16 simple items	-	English	5 min	CGA by geriatrician
Strawbridge questionnaire	16 simple items	Self-administered questionnaire	English	-	CGA by geriatrician
					101

Tool/study	No. items	Mode of administrat°	Language	Administrat° duration	Reference G assessment
PRISMA-7	7 simple items	Self-administered questionnaire	French	3 min	SMAF scale
Bright tool	11 simple items	Self-administered questionnaire	English	-	MDS-HC by nurse in the home
Self-administered test	49 simple items	Self-administered questionnaire	Italian	-	MCPS by geriatricia
Tilburg frailty indicator	15 simple items	Self-administered questionnaire	Dutch	14 min	CGA by trained interviewers
SHARE-FI	5 simple items plus grip measured on a dynamometer	Non-medical staff	-	-	No CGA Mortality physical, social and cognitive data from the SHARE survey

Tool/study	Patient- reported fatigue	Physical performance	Walking	No. comorbidities	Nutritional state	Psychometric properties compared to CGA
Screening letter ²⁵	+	_			-	Se = 0.95/Sp = 0.68
Sherbrooke postal questionnaire ²⁶	-	-	+	-	-	Se = 0.75/Sp = 0.52
Functional assessment screening package ²⁷	-	+	+	-	+	Kappa = 0.77-1/Se = 0.70-0.95/Sp = 0.64-0.95
Screening instrument ²⁸	-	+	+	-	-	Se = 0.65-0.93/Sp = 0.50-0.96
Strawbridge questionnaire ²⁹	-	+	+	_	+	Inter-evaluation agreement = 0.67/kappa = 0.29
PRISMA-730	+	_	+	_	_	Se = 0.78/Sp = 0.74
Bright tool31	+	+	+	_	_	Kappa = $0.77/\alpha = 0.77/Se = 0.65/Sp = 0.84$
Self-administere test32	d –	-	+	+	+	Similar classification for 48% of the subjects
Tilburg frailty indicator 33	+	+	+	-	+	Kappa = $0.79/\alpha$ = $0.73/Pearson$'s corr. coeff. (r) significant ($P < 0.001$)
SHARE-FI34	+	+	+	-	+	Compared to non-frail odds ratio for mortality among frail >1/Spearman's corr. coeff. significan (P < 0.001)

The "PRISMA 7" Questions

- 1. Are you more than 85 years?
- 2. Male?
- 3. In general do you have any health problems that require you to limit your activities?
- 4. Do you need someone to help you on a regular basis?
- 5. In general do you have any health problems that require you to stay at home?
- 6. In case of need, can you count on someone close to you?
- 7. Do you regularly use a stick, walker or wheelchair to get about?

Part A Determinants of frailty 1. Which sex are you? 2. What is your age? 3. What is your marital status? 4. In which country were you bom? 4. In which country were you bom? 5. What is the highest level of education you have completed? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 7. Overall, how healthy would you say your lifestyle is? 7. Overall, how healthy would you say your lifestyle is? 7. Overall, how healthy would you say your lifestyle is? 9. Have you experienced one or more of the following events during the past year? 1. The death of a loved one 2. The death of a loved one 3. What is the highest level of education you have completed? 9. The work of the professional or university education 9. Have you experienced one or more of the following events during the past year? 1. The death of a loved one 2. The death of a loved one 3. What is your age? 4. In which country were you bom? 9. Have you experienced one or more of the following events during the past year? 1. The death of a loved one 3. What is your age? 4. In which country were you bom? 9. Have you experienced one or more of the following events during the past year? 1. The death of a loved one 3. What is your age? 4. In which country were you bom? 5. What is the highest level of education you have completed one one of the following events during the past year. 9. Year you experienced one or more of the following events during the past year. 1. The death of a loved one 3. Description with your home living environment? 9. Year on no 9. Ye	Tilburg Frailty Indicator (TFI)* Gobbens RJJ, van Assen MALM, Luikx KG, Wijnen-Sponselee MTh, Frailty Indicator: psychometric properties. J Am Med Dir Assoc 2010;	Schols JMGA. The Tilburg 11(5):344-355.	
2. What is your age? years 3. What is your marital status? 3. What is your marital status? 4. In which country were you born? 4. In which country were you born? 5. What is the highest level of education you have completed? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 7. Overall, how healthy would you say your lifestyle is? 7. Overall, how healthy would you say your lifestyle is? 9. Have you experienced one or more of the following events during the part of a serious litness your elf on yes 9. Have you experienced one or more of the following events during the past year? 10. The Metherlands and the part of the past your past of the past year? 11. Are you satisfied with your home living environment? 12. Are you satisfied with your home living environment? 13. What is your marital status? 14. In which country were you be partial status? 15. What is the highest level or equipment of the past your	Part A Determinants of frailty		
3. What is your marital status? 0 married/living with partner 0 unmarried 0 separated/divorced 0 vidow/widower 4. In which country were you born? 1 Permet Define East Indies 1 Summarried 0 Permet Define East Indies 0 Summarried 0 Netherlands Antilles 0 Turkey 0 Morocco 0 Other, namely	1. Which sex are you?	0 male 0 female	
4. In which country were you born? 4. In which country were you born? 5. What is the highest level of education you have completed? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 7. Overall, how healthy would you say your lifestyle is? 7. Overall, how healthy would you say your lifestyle is? 8. Do you have two or more diseases and/or chronic disorders? 9. Have you experienced one or more of the following events during the past year? 1. the death of a loved one 2. a serious liness in a loved one 3. a divorce or ending of an important intimate relationship 3. a traific accident 4. In which country were you be perienced one or more of the following events one a divorce or ending of an important intimate relationship 3. Over all, how healthy would you say your lifestyle is? 4. In the death of a loved one 5. Over all, how healthy would you say your lifestyle is? 6. Which category indicates your net monthly household income? 7. Overall, how healthy would you say your lifestyle is? 9. Heave you experienced one or more of the following events during the past year? 1. the death of a loved one 2. a serious liness your serious planes. 3. yes 4. In which category during the life your home living environment? 4. In which category experienced one or more of the following events during the past year? 2. the death of a loved one 3. yes 4. One 4. a serious liness your serious hour and year and year and year and year and year? 5. What is the third say the professional or university education 6. Which category education 6. Which	2. What is your age?	years	
O Former Dutch East Indies O Suriname O Netherlands Antilles O Turkey O Morocco O Other, namely	3. What is your marital status?	0 unmarried 0 separated/divorced	
6. Which category indicates your net monthly household income? 6. Which category indicates your net monthly household income? 9	In which country were you born?	Former Dutch East Indies Suriname Netherlands Antilles Turkey Morocco	
6601 - 6900 0 e801 - 61500 0 e801 - 61500 0 e1201 - 61500 0 e1301 - 62100 0 yes 0 no 0 yes 0 no 0 yes 0 no 0 yes 0 no 0 a serious lilness in a loved one a divorce or ending of an important intimate relationship a divorce or ending of an important intimate relationship a a timic accident 0 yes 0 no 10. Are you satisfied with your home living environment? 0 yes 0 no	5. What is the highest level of education you have completed?	0 secondary education 0 higher professional or	
O not healthy, not unhealthy 0 unhealthy 8. Do you have two or more diseases and/or chronic disorders? 9. Have you experienced one or more of the following events during the past year? - the death of a loved one - a serious liness yourself 0 yes 0 no - a serious liness yourself 0 yes 0 no - a divorce or ending of an important intimate relationship - a traffic accident 0 yes 0 no - a climic or ending of an important legistration of the control of	6. Which category indicates your net monthly household income?	0 €601 - €900 0 €901 - €1200 0 €1201 - €1500 0 €1501 - €1800 0 €1801 - €2100	
9. Have you experienced one or more of the following events during the past year? 1 - the death of a loved one 1 - a serious illness yourself 2 - a serious illness in a loved one 1 - a divorce or ending of an important intimate relationship 2 - a traffic accident 3 - a crime 4 - a crime 5 - a crime 5 - a crime 6 - a divorce or ending of an important intimate relationship 7 - a crime 7 - a crime 7 - a crime 9 - a crime	7. Overall, how healthy would you say your lifestyle is?	0 not healthy, not unhealthy	
during the past year? - the death of a loved one - a serious litness yourself 0 yes 0 no - a serious litness in a loved one - a divorce or ending of an important intimate relationship 0 yes 0 no - a traffic accident 0 yes 0 no - a crime 0 yes 0 no 10. Are you satisfied with your home living environment? 0 yes 0 no	8. Do you have two or more diseases and/or chronic disorders?	0 yes 0 no	
	during the past year? - the death of a loved one - a serious illness yourself - a serious illness in a loved one - a divorce or ending of an important intimate relationship - a traffic accident	0 yes 0 no 0 yes 0 no 0 yes 0 no 0 yes 0 no	
105	10. Are you satisfied with your home living environment?	0 yes 0 no	
103			105

Part	B Components of frailty				Scoring Part B Components of frailty (range: 0 - 15		
B1	Physical components						
11.	Do you feel physically healthy?	0 yes		0 no	Question 11:	yes = 0, no = 1	
12.	Have you lost a lot of weight recently without wishing to do so?	0 yes		0 no	Question 12 – 18:	no = 0, yes = 1	
	('a lot' is: 6 kg or more during the last six months, or 3 kg or more during the last month)				Question 19:	no and sometimes = 0, ye	
Do y	ou experience problems in your daily life due to:				Question 20 and 21:	no = 0, yes and sometime	
13.	difficulty in walking?	0 yes		0 no	Question 22:	yes = 0, no = 1	
14.	difficulty maintaining your balance?	0 yes		0 no	Question 23:	no = 0, yes = 1	
15.	poor hearing?	0 yes		0 no	Question 24:	no = 0, yes and sometime	
16.	poor vision?	0 yes		0 no	Question 25:	yes = 0, no = 1	
17.	lack of strength in your hands?	0 yes		0 no	Cutpoint: 5		
18.	physical tiredness?	0 yes		0 no			
B2	Psychological components						
19.	Do you have problems with your memory?	0 yes	0 sometimes	0 no			
20.	Have you felt down during the last month?	0 yes	0 sometimes	0 no			
21.	Have you felt nervous or anxious during the last month?	0 yes	0 sometimes	0 no			
22.	Are you able to cope with problems well?	0 yes		0 no			
ВЗ	Social components						
23.	Do you live alone?	0 yes		0 no			
24.	Do you sometimes miss having people around you?	0 yes	0 sometimes	0 no			
25.	Do you receive enough support from other people?	0 yes		0 no			

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EMERGENCY DEPARTMENT

Personal story

You are admitted because of a stupid fall at sport. What happened in emergency department (ED) ?

Robert is admitted because of a stupid fall at home. What happened in the ED?

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Main way of admission: the emergency department

Qualitative and quantitative concerns: the emergency dept. (ED) constrains and the geriatric complexity

Rapid management of an acute illness

Unforeseeable nature and 24/7 availability

Overcrowding

Multiple comorbidities and complex care needs (ψsocial)

Atypical presentations

Longer LOS in ED Fragmentation of care

Aminzadeh & al. Annals of Emergency Medicine 2002 Salvi F & al. Intern Emerg Med, 2007 Samaras & al. Annals of Emergency Medicine 2010

Older Patients in the Emergency Dept: adverse outcomes after discharge

- Discharged OP to community:
 - 1 patient in 2 readmitted to ED at 6 months
 - 1 patient in 3 with functional decline (FD) at 3 months
 - 1 patient in 10 : death
 - Risk for hospitalization : x3

Salvi et al, Intern Emerg Med 2011

- Hospitalized OP:
 - Early FD (48h)
 - 1 patient in 3 with persistent FD at 3 months
 - ↑mortality,
 - ↑LOS, ↑%unplanned readmissions,
 - †institutionalization, †use health care resources

Ellis et al. BMJ 2011; Ellis et al. Cochrane 2010

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Older Patients in the Emergency Dept: Effectiveness of interventions

· Discharged OP to community: promising results

Hastings et al. Acad Emerg Med October 2005 Fealy et al. Journal of Advanced nursing, 2009 Graf et al. Aging Clin Exp Res. October 5, 2010 Sinha et al. Annals Emerg Med, 2011

- Hospitalized OP: Geriatric Evaluation and Management Unit:
 - Higher likelihood of being alive and in their own homes
 - Less likely to be institutionalized, to suffer death or deterioration
 - More likely to experience improved cognition
 - Potential cost reduction

Ellis et al. BMJ 2010 Ellis et al. Cochrane 2011

CGA and Case-finding in ED

- CGA following by appropriate interventions could improves outcomes
 - CGA is time-consuming and cannot be applied routinely in ED
- Screening of at-risk patient more efficient than age-based screening
 - Identifying older people that would benefit the most from G intervention
 - Time and resources saving

Graf et al. Aging Clin Exp Res. October 5, 2010

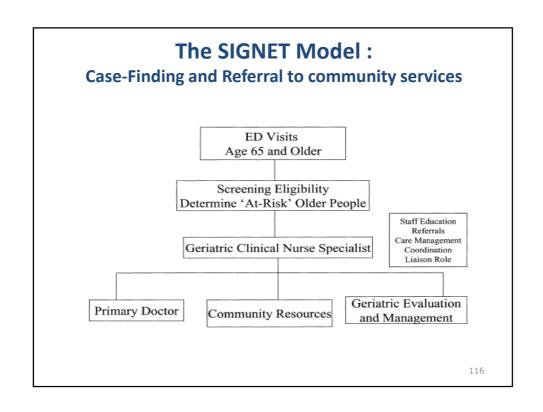
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Rapid Emergency Dpt Intervention for « senior at risk » of FD

- Case-finding:
 - Screening tool ISAR to 65+: positive ≥ 2/6
 - Self-reported or nurse evaluation
- Intervention: CGA and referral to community services for high-risk patients
- Outcomes : death, institutionalization and increased functional dependence at 6-month

Mc Cusker et al. JAGS 2001

The	The Identification Senior At Risk Tool: Mc Cusker et al. JAGS 1999								
			Hospital use only						
1.	. Before the illness or injury that brought you to the Emergency, did you need someone to help you on a regular basis?	☐ YES	1 0						
2	Since the illness or injury that brought you to the Emergency, have you needed more help than usual to take care of yourself?	☐ YES	1 0						
3	. Have you been hospitalized for one or more nights during the past 6 months (excluding a stay in the Emergency Department)?	☐ YES	1 0						
4	. In general, do you see well?	☐ YES	0 1						
5	. In general, do you have serious problems with your memory?	☐ YES	1 0						
6.	. Do you take more than three different medications every day?	☐ YES	1 0						
		TOTAL:		· 					
				115					



The Triage Risk Screening Tool: Meldon et al. Acad Emerg Med 2003					
History of cognitive impairment (poor recall or not oriented)					
2. Difficulty walking / transferring or recent falls					
3. ☐ Five or more medications					
4. ED use in previous 30 days or hospitalization in previous 90 days					
5. Lives alone and/or no available caregiver					
6. ☐ ED staff professional recommendations: ☐ Suspected abuse/neglect ☐ Problems with iADL ☐ Non compliant patient with < 5 med ☐ Others : specify ☐ Suspected substance abuse					
If 2 or more factors identified: Referral to GEM Nurse Referral to GEM Nurse not indicated Referral to Social Work when GEM nurse not available					

The Flemish TRST GRP* RISICO JA NEE 2 Aanwezigheid van een cognitieve stoomis 0 2. Alleenwonend of geen hulp mogelijk door inwonende partner/familie 0 1 3. Moeilijkheden bij stappen/transfers of gevallen in de afgelopen 6 maanden 0 1 4. Hij/Zij werd gehospitaliseerd in de afgelopen 3 maanden 0 1 5. De patient gebruikt ≥ 5 geneesmiddelen 0 Totaalscore Kenis et al. Crit Rev Oncol Hematol 2006 $_{18}$

G Syndromes predict postdischarge outcomes among older patients



ORIGINAL CONTRIBUTION

Geriatric Syndromes Predict Postdischarge Outcomes Among Older Emergency Department Patients: Findings From the interRAI Multinational Emergency Department Study

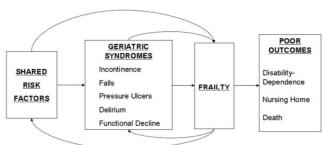
Andrew P. Costa, PhD, John P. Hirdes, PhD, George A. Heckman, MD, MSc, Aparajit B. Dey, MD, Palmi V. Jonsson, MD, Prabha Lakhan, RN, PhD, Gunnar Ljunggren, MD, PhD, Katrin Singler, MD, MME, Fredrik Sjostrand, MD, PhD, Walter Swoboda, MD, Nathalie I.H. Wellens, PhD, and Leonard C. Gray, MD, PhD

Costa et al. Acad Emerg Med April 2014

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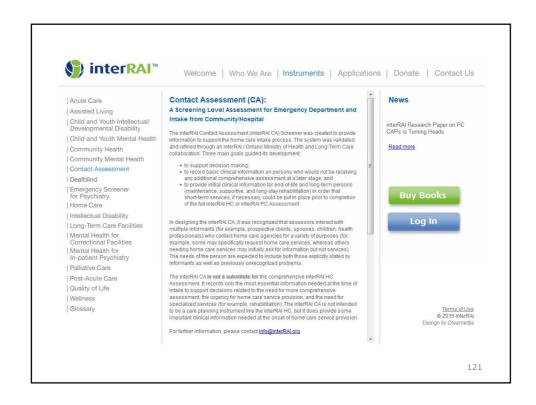
Frailty and Geriatric Syndromes

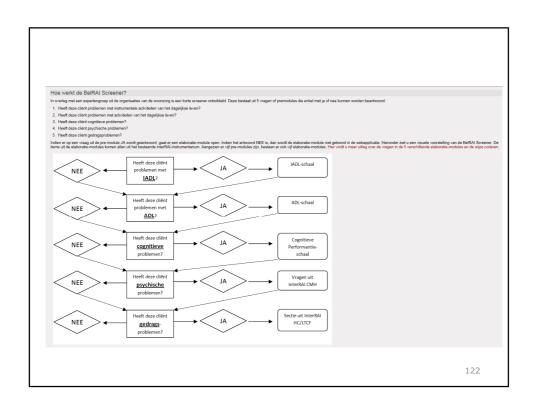




A unifying conceptual model demonstrates that shared risk factors may lead to geriatric syndromes, which may in turn lead to frailty, with feedback mechanisms enhancing the presence of shared risk factors and geriatric syndromes. Such self-sustaining pathways may result in poor outcomes involving disability-dependence, nursing home placement, and ultimately death, thus holding important implications for elucidating pathophysiologic mechanisms and designing effective intervention strategies.

Inouye et al. JAGS 2007

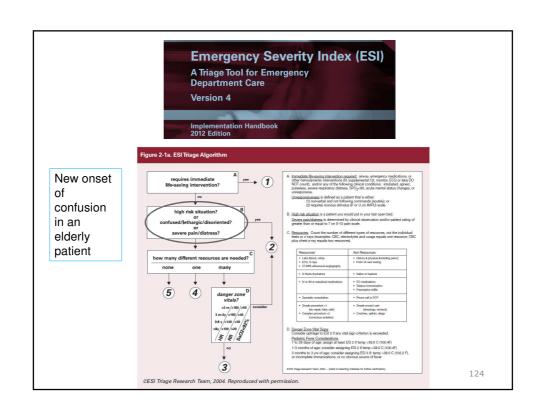




G Syndromes predict postdischarge outcomes among older patients

- Common G conditions influence the p! of some adverse events among older patients
- Geriatric clinical features may help to refine and focus existing clinical reasoning
 - In-patients: psychosocial, locomotion and trauma
 - GEMU, co-management (hip #) elder-friendly care
 - Out-patients: recent ED visits, bADL impairment, psy
 - Unmet needs, mental health services, coordination with primary care in suba-cute cases

•Costa et al. Acad Emerg Med 2014



A novel multidimensional geriatric screening tool in the ED: evaluation of feasibility and clinical relevance

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ABSTRACT

Purposes: Geriatric problems frequently go undetected in older patients in emergency departments (EDs), thus increasing their risk of adverse outcomes. We evaluated a novel emergency geriatric screening (EGS) tool designed to detect geriatric problems.

Basic procedures: The EGS tool consisted of short validated instruments used to screen 4 domains (cognition, falls, mobility, and activities of daily living). Emergency geriatric screening was introduced for ED patients 750 years or older throughout a 4-month period. We analyzed the prevalence of abnormal EGS and whether EGS increased the number of EGS-related diagnoses in the ED during the screening, as compared with a preceding control period. control period.

control period.
Main findings: Emergency geriatric screening was performed on 338 (42.5%) of 795 patients presenting during screening. Emergency geriatric screening was unfeasible in 175 patients (22.0%) because of life-threatening conditions and was not performed in 282 (35.5%) for logistical reasons. Emergency geriatric screening took less than 5 minutes to perform in most (85.8%) cases. Among screened patients, 285 (84.3%) had at least 1 abnormal EGS finding, In 270 of these patients, at least 1 abnormal EGS finding did not result in a diagnosis in the ED and was reported for further workup to subsequent care. During screening, 142 patients (42.0%) had at least 1 diagnosis listed within the 4 EGS domains, significantly more than the 29.3% in the control period (odds ratio 1.75; 95% confidence interval, 1.34-2.29; P < .001). Emergency geriatric screening predicted nursing home admission after the in-hospital stay (odds ratio for ≥3 vs <3 abnormal domains 12.13; 95% confidence interval, 2.79-5.27; P = .001).

Principal conclusions: The novel EGS is feasible, identifies previously undetected geriatric problems, and predicts determinants of subsequent care.

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Cognition		
Instruction: Ask the patient the following questions. If the patient does not respond, the question is rated incorrect.		
What day is today? What is the date today? (± 1 day is correct) What year is this? Spell "radio" backward. Evaluation consistent with impairment of cognition (if one single response was incorrect):	Incorrect* Incorrect* Incorrect Incorrect Yes	Correct Correct Correct No
Falls		
Instruction: Rate the following questions considering all available sources (patient, proxy, observation, reports).		
Did the patient present to the ED because of a fall? Did the patient have one or more falls during the last 12 months? Evaluation consistent with patient history of falls (if one single response was yes): Mobility	Yes* Yes Yes	No No No
2000-000-000-000-000-000-000-000-000-00		
Instruction: Rate the following question considering all available sources (patient, proxy, observation, reports).		
Did the patient require walking aids (cane, wheeled walker, or helping person) indoors or outdoors before presenting to the	ED? Yes	No
Instruction: Rate the following questions according to the current situation in the ED.		
Is the patient currently confined to bed? Does the patient currently need help (walking aids or helping person) to get out of bed? Does the patient need ≥20 seconds for the Timed Up and Go Test? Evaluation consistent with impairment of mobility (if one single response was yes):	Yes Yes Yes Yes	No No No
ADLs		
Instruction: Rate the following question considering all available sources (patient, proxy, observation, reports).		
Did the patient require assistance for personal hygiene (sponge bath, tub bath, or shower) before presenting to the ED?	Yes*	No
Instruction: Rate the following questions according to the current situation in the ED.		
is the patient currently confined to bed or does he need help (walking aid or helping person) to get out of bed? Does the patient require assistance (for direct help or instruction) for dressing (clothes or shoes)? Does the patient require assistance (for direct help or instruction) for toileting? Does the patient require assistance (for direct help or instruction) for feeding? Evaluation consistent with impairment in ADL (for single response was yes):	Yes* Yes* Yes Yes Yes	No No No No







Oncogeriatrics

Use of Geriatric Assessment for Older Adults in Oncology Setting: A Systematic Review
Puts et al. J Natl Cancer Inst 2012

International Society of Geriatric Oncology Consensus on Geriatric Assessment in Older Patients With Cancer Wildiers et al. J Clinical Oncol 2014

Four Screening instruments for frailty in older patients with and without cancer: a diagnosis study

Smets et al. BMC Geriatrics 2014

Frailty screening methods for predicting outcome of a comprehensive geriatric assessment in elderly patients with cancer: a systematic review

Hamaker et al. The Lancet Oncology 2012

Older Patients in the Oncology dpt: Rationale

- >50% newly diagnoses 65+, heterogeneous group
 - Cancer type, stage, disease & R/ trajectories
 - Ageing process: not only chronologic age
- Debilitating disease: preserve QOL
 - Prevent functional decline
 - Live in own home
- Treatment toxicity & decision: decompensation of other comorbidity/psychosocial factors

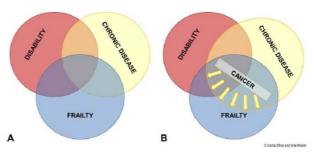
Older Patients in the Oncology Dpt: Rationale

- Detection of potentially reversible G problems
 - Identify opportunities for intervention
- Prediction of treatment toxicity/↓ in QOL
 - Preventive measures/Intervention to 1 QOL and compliance
 - Treatment decision
- Prognostic information
 - P! to die because of / with the cancer
- · Appraising objective health
 - Comorbidity/ψsocial factors that may decompensate

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Oncogeriatrics: prevalence of frailty

CA CANCER J CLIN 2017;67:362-377



62% of older cancer patients are « frail » or « pre-frail »

Handforth et al. Ann Oncol 2015;26:1091

Oncogeriatrics Organisation

- Several ways of implementation
- Preference should be given to models that fit the local health care structure and setting
- Interaction with multidisciplinary G teams is highly recommended... for selected patients

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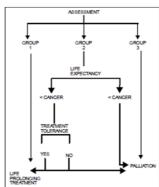
How to identify frailty in older cancer patients?

CGA = gold standard in oncologic literature for 20y

Management of Cancer in the Older Person: A Practical Approach

LODOVICO BALDUCCI, MARTINE EXTERMANN

The Oncologist 2000;5:224-237



VOLUME 32 · NUMBER 24 · AUGUST 20 2014

JOURNAL OF CLINICAL ONCOLOGY

REVIEW ARTICLI

International Society of Geriatric Oncology Consensus on Geriatric Assessment in Older Patients With Cancer

Hans Wildiers, Pieter Heeren, Johan Flamaing, Cindy Kenis, and Koen Milisen, University Hospitals Leuven, KU Leuven, Leuven, Leuven, Belgium; Hans Wildiers, Pieter Heeren, Martine Puts, Eva Topinkova, Maryska L.G. Janssen-Heijnen, Martine Extermann, Claire Falandry, Andrew Artz, Etienne Brain, Giuseppe Colloca, Johan Flamaing, Theodora Karnakis, Cindy Kenis, Riccardo A. Audisio, Supriya Mohile, Lazzaro Repetto, Barbara Van Leeuwen, Koen Milisen, and Arti Hurria

Oncogeriatrics: Which domains and tools?

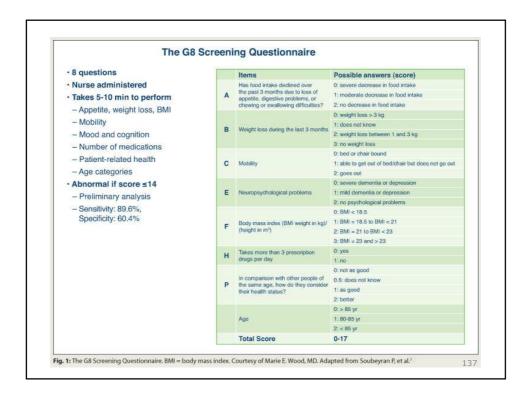
- Important domains are:
 - Functional status
 - Fatigue
 - Comorbidity
 - Cognition and mental health status
 - Social support
 - Nutrition
 - G syndromes

Oncogeriatrics Screening tools

- Screening for relevance of CGA
 - abbreviated CGA (aCGA)
 - Vulnerable Elders Survey-13 (VES-13),
 - Groningen Frailty Indicator (GFI)
 - Geriatric 8 (G8)
- Various tools available, no superiority proven

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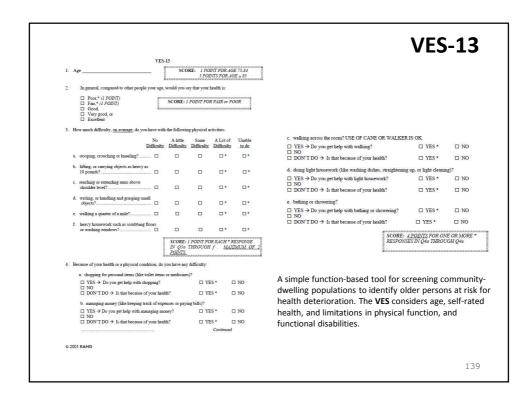
	GFI ⁵	G836	VES-136	aCGA ^{38*}	Fried ⁴	Barber ²⁴	TRST ³⁷
Functional status	27%	12%	60%	60%	60%		
ADL impairment	13%	**	-	20%		33%	
IADL impairment	7%	**		40%		11%	
Mobility and falls	7%						20%
Psychosocial domain	40%	12%	w.	40%			
Cognitive disorder	7%	**		27%			20%
Mood and anxiety	13%		(m)	13%	**	**	
Social support	20%	***	995	**		11%	
Neurosensory deficits	13%		***			22%	
Nutritional status and weight loss	7%	46%	-		20%		
Polypharmacy	7%	6%					20%
Comorbidity	960		m (10	**		
Recent hospital admission			**			11%	20%
Geriatric syndromes							20%
Self-reported health	7%	12%	10%		20%	11%	Ti de la companya de
Age	page .	12%	30%		**		
Optimum score	0	17	0		0	0	0
Poorest score	15	0	10		5	9	5
Standard cutoff value†	4+	≤14	3+		3+	1+	2+
Population designed for	Various	Patients with cancer	Community- dwelling elderly	Patients with cancer		Patients in primary care	Patients in emergency roon
GFI=Groningen Frailty Index. G8=Geriatric 8 ADL=activities of daily living. IADL=instrum be considered frail.			aCGA=abbreviated	comprehensive g	eriatric assessr	nent. TRST=triag	e risk screening tool



The Groningen Frailty Indictor

Are you able to carry out these tasks single-handed without any help? (The use of aids such as a walking stick, walking frame, wheelchair, is considered as independent)

- 1 Shopping,
- 2 Walking around outside (around the house or to the neighbors)
- 3 Dressing and undressing
- 4 Going to the toilet
- 5 What score do you give yourself for physical fitness? (scale 0 to 10)
- 6 Do you experience problems in daily life due to poor vision?
- 7 Do you experience problems in daily life due to poor hearing?



Impact of CGA on oncological decision treatment Patients 70 years or older with newly diagnosed cancer (N = 656) Treatment proposed by the oncologist: initial cancer treatment plan (n = 656) Adaptation of oncological treatment Referred to geriatrician Comprehensive Geriatric Assessment (n = 392) in 20% of patient following CGA Undefined initial cancer treatment plan (n = 17) Multidisciplinary meeting: decision about the cancer treatment plan: final cancer treatment plan (n = 375)No change in the initial cancer Change in the initial cancer treatment plan (n = 78) (n = 297) Caillet et al, J Clin Oncol 2011.

Impact of CGA on oncological decision treatment

• ADL

Nutrition

 Table 5. Multivariate Analysis to Identify Factors Independently Associated

 With Changing the Cancer Treatment

Factor	OR	95% CI	P
ECOG PS 1-point increase	1.07	0.72 to 1.59	.74
Inappropriate social environment	1.34	0.61 to 2.95	.46
ADL .5-point decrease	1.25	1.04 to 1.49	.016
Walking problems: risk of falls	1.27	0.53 to 3.03	.54
Malnutrition	2.99	1.36 to 6.58	.007
Cognitive impairment	0.93	0.44 to 2.00	.86
Depressive disorder	1.84	0.89 to 3.80	.10
Polypharmacy	1.72	0.72 to 4.14	.22
Urinary and/or fecal incontinence	1.09	0.45 to 2.64	.84
No. of comorbidities by 1-point increase	1.09	0.98 to 1.22	.11

NOTE. Multivariate analysis using a logistic regression model that included factors listed in the table: ECOG PS, inappropriate social environment, 1-point ADL score decrease, walking problems/risk of falls, malnutrition, cognitive impairment, depressive disorder, polypharmacy, incontinence, and 1 point per additional comorbidity.

Abbreviations: OR, odds ratio; ECOG PS, Eastern Cooperative Oncology Group performance status; ADL, Activities of Daily Living.

Caillet et al, J Clin Oncol 2011.

VOLUME 30 · NUMBER 15 · MAY 20 2012

JOURNAL OF CLINICAL ONCOLOGY

Predictors of Early Death Risk in Older Patients Treated With First-Line Chemotherapy for Cancer

Pierre Soubeyran, Marianne Fonck, Christèle Blanc Bisson, Jean-Frédéric Blanc, Joël Ceccaldi, Cécile Mertens, Yves Imbert, Laurent Cany, Luc Vogt, Jerôme Dauba, Francis Andriamampionona, Nadine Houédé, Anne Floquet, Francois Chomy, Véronique Brouste, Alain Ravaud, Carine Bellera, and Muriel Rainfray

Soubeyran et al, J Clin Oncol 2012.

	Nonmetas: 0-1	tatic/IPI	Metastatic/IPI 2-		
Cancer Type	No. of Patients	%	No. of Patients	9	
Non-Hodgkin's lymphoma*	61	57.0	44	41	
Stomach/colon cancer	39	30.2	90	69	
Other solid tumors	21	18.8	91	81	
Lung	9		27		
Primary unknown	0		4		
Ovarian	1		19		
Bladder	3		15		
Prostate	0		18		
Pancreas	8		15		

Table 4. Logistic Regression Model Analysis for Early Deaths (within 6 months) That Occurred for All Patients Who Received First-Line Chemotherapy (n = 339)

Risk Factor*	Odds Ratio	95% CI	P
Sex			
Female	1	Reference	
Male	2.40	1.20 to 4.82	.013
Tumor stage			
Localized	1	Reference	
Advanced	3.9	1.59 to 9.73	.003
Mini Nutritional Assessment			
Good nutrition, score > 23.5	1	Reference	
At risk/poor nutrition, score ≤ 23.5	2.77	1.24 to 6.18	.013
Timed Get Up and Go			
No impairments (≤ 20 seconds)	1	Reference	
Impaired	2.55	1.32 to 4.94	.006

NOTE: Model was adjusted for treatment site (regional and teaching hospi-

NOTE: Model was adjusted for treatment site pregional and becoming four-tals v community hospitals).

*Age, tumor site, Activities of Daily Living, Mini-Mental State, platelet count, and performance status were also included in the model but not retained because they were not significant.

In patients greater than 70 years of age with cancer, advanced disease, a low MNA score, and poor mobility predicted early death. We recommend that the MNA and GUG, performed by a trained nurse, be maintained as part of routine pretreatment workup in these patients to identify at-risk patients and to inform the decision-making process for chemotherapy.

Soubeyran et al, J Clin Oncol 2012.

CGA in older cancer patients...

- Identify underlying diseaese at risk of decompensation during oncological treatment
- · Assess preferences and motivation of patients
- Screen for caregivers'burden
- · Help to maintain QOL
- Plan follow up
- Talk about palliative care

Clinical picture in oncogeriatric

- M. JL, born in 1935
- Urologic symptoms in 08/2018 (pain, pollakiuria et hamaturia)
- Cystoscopie: High grade adenocarcinoma (G3), pT2, negative extension assessment.
- OMC: neo-adjuvant chemotherapy+ radical cystoprostatectomy (+Bricker) versus Radiotherapie + chemotherapy.
- Comorbidity: ischemic cardiovascular disease (MI 2015 with low residual EF 40%)
- R/ Bisoprolol, AAS, lisinopril, aldactone

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JL, 1935

- G8:13/17
- ECOG: 0
- Isar 1/6
- Social: live at home, MD in oncology!
- ADL: Katz 6/6; lawton 7/7
- Mobility : no previous fall, Timed Up&Go : 13 sec
- Pain: miction 2/10
- Fatigue: 4/10, sleep disturbances due to nycturia (7-8x)
- QOL (Eortc Qlq-C30): 5/7
- No sensoriel limitations (audition, vision)
- MMSE: 29/30
- Geriatric Depression Scale: 3/15
- MNA 11/14 risk of denutrition, loss of 2 kg last 2 months (72 kg, BMI 24)
- · CCLs:
- OncoG MC:







Cardiovascular and surgical department

Importance of frailty in patients with cardiovascular disease

Mandeep et al. European Heart Journal 2014

Role of frailty assessment in patients undergoing cardiac interventions

Rowe et al. Open Heart 2014

Importance of frailty in CV patients

- CVDs are the leading cause of morbidity and mortality
 - 82% CVD † are 65+; 46% CVD † are 75+
 - 1 CVD hospital admission
 - Non-cardiac predictors for 1-yr survival (TAVI)
- G prognosis determinants are seldom measured into clinical decision-making
 - High-risk interventions : CABG vs. PCI vs. TAVI vs. conservative treatment

Frailty in the older surgical patient

- Higher increase of older surgical people than the rate of population ageing
- Adverse post-operative outcomes, despite surgical, anaesthetic and medical advances
- Frailty is an independant risk factors for morbidity, mortality, 1 LOS and institutional discharge
 - Preoperative risk stratification tool
 - Identifying potentially modiafiable factors

Importance of frailty in CV patients

Table 5 Reasons for evaluating whether frailty is present in patients with cardiovascular diseases

- CV
 - m
- G p cli
- 1 Population ageing is increasing the number of frail patients with CVD
- 2 Eye ball or end of the bed assessments of frailty may not be reliable
- 3 Frailty increases the risks of cardiac surgery and other cardiovascular interventions
- 4 Frailty increases the risk of cardiovascular and non-cardiovascular mortality and the need for future institutional care
- 5 Frail patients may have more complications from medical treatments
- 6 The benefits of some cardiac interventions may be less in frail elderly patients because of competing risks. Non-cardiac deaths dominate following TAVR, PCI, and CABG

into

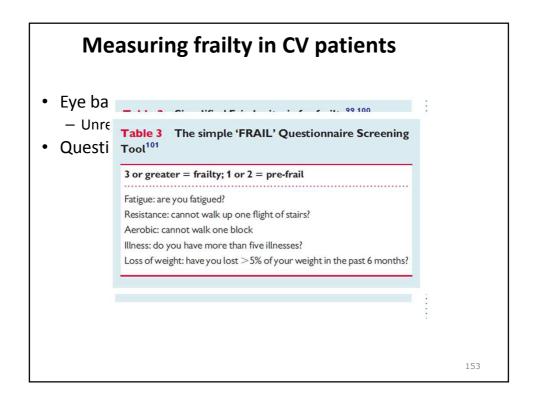
Measuring frailty in CV patients

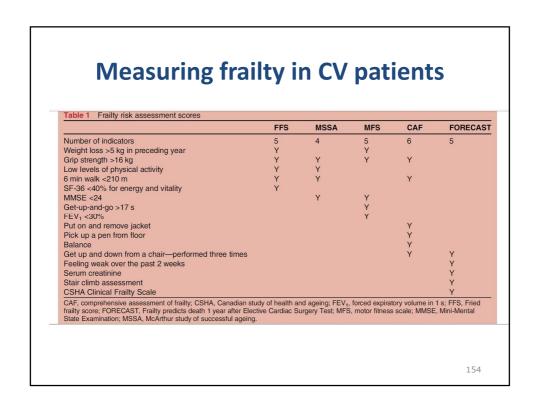
- · Eye ball test at the end of the bed
 - Unreliable & prone to bias
- Questionnaires or simple measurements

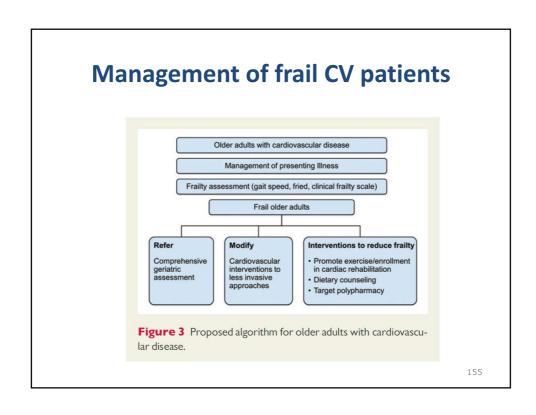
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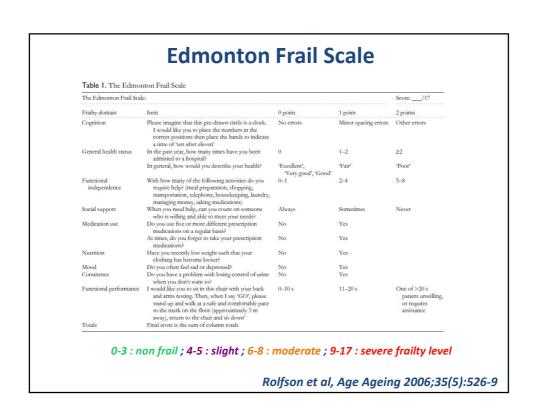
Measuring frailty in CV patients

Table 2 Simplified Fried criteria for frailty 99,100 Eye l 1 Unintentional >4.5 kg in the past year weight loss 2 Exhaustion For at least 3 days during the last week 'I felt Que: that everything I did was an effort' or 'I could not get going' 3 Physical activity No physical activity, spend most of the time sitting or rarely a short walk during the last Time to walk 4 m > 6 s4 Walk time 5 Grip strength Grip strength by dynamometer Frail = 3 or more criteria present, pre-frail = 1 or 2 criteria.



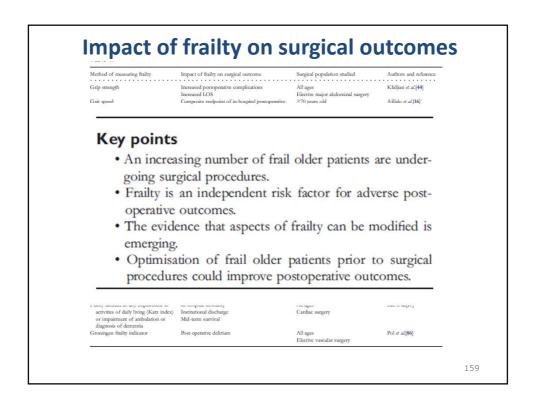


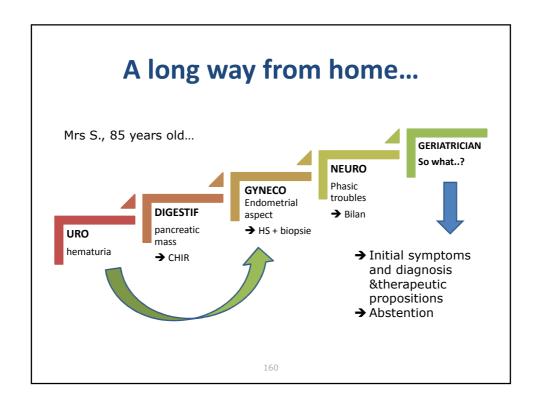


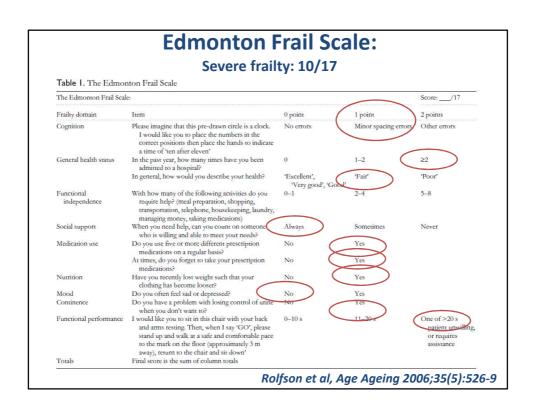


Star cancer in companies	Table 1. The Edmonton Frail Scale				
The Edmonton Frail Scal	le:			Score:/17	
Frailty domain	Item	0 point	1 point	2 points	
Cognition	Please imagine that this pre-drawn circle is a clock. I would like you to place the numbers in the correct positions then place the hands to indicate a time of 'ten after eleven'	No errors	Minor spacing error	s Other errors	
General health status	In the past year, how many times have you been admitted to a hospital?	0	1-2	≥2	
PathoAssoc	f inflammatory bi physiology of frailty iation with post-ope ectal surgical patient	rative c	·	ŕ	
PathoAssoc	physiology of frailty iation with post-ope	rative c	·	ŕ	
PathoAssoc colore	physiology of frailty iation with post-ope ectal surgical patient dothing has become looser? Do you often feel said or depressed? Do you have a problem with losing control of urine	rative c s	omplicatic	ŕ	
PathoAssoc colore	physiology of frailty iation with post-ope ectal surgical patient clothing has become looser? Do you often feel sad or depressed?	rative c s	omplicatio Yes	ŕ	

Method of measuring frailty	Impact of frailty on surgical outcome	Surgical population studied	Authors and reference
Grip strength	Increased postoperative complications Increased LOS	All ages Elective major abdominal surgery	Klidjian et al.[44]
Gait speed	Composite endpoint of in-hospital postoperative mortality or major morbidity (as defined by Society of Thoracic Surgeons criteria) Cardia: surgery	≥70 years old	Afilalo et al.[16]
Edmonton Frail Scale	Postoperative complications Prolonged LOS	≥70 years old Lower limb orthopaedic surgery	Dasgupta et al.[13]
	Increased institutionalisation rate	Spinal surgery Abdominal surgery Vascular surgery	
Frailty score based on frailty	Postoperative complications	≥65 years old	Makary et al.[14]
phenotype	Prolonged LOS New institutionalisation at discharge	Elective surgery (major and minor)	
Comprehensive Assessment of Frailty Score	Increase in 30-day mortality	Cardiac surgery	Sundermann et al[15]
8 'markers' of frailty (age, cognition, recent weight loss, BMI, serum albumin, falls, depression, haematocrir)	Increase in 6-month mortality (although underpowered for this)	≥65 years old	Robinson et al. [46]
	Post-discharge institutionalisation	General, thoracic, urology and vascular surgery (patients undergoing major elective surgery necessitating postoperative surgical ICU admission)	
14 frailty 'characteristics' in 6 domains (comorbidity, function, cognition, geriatric syndromes, extrinsic frailty)	Institutionalisation at hospital discharge	≥ 65 years old	Robinson et al. [47]
NB: most closely associated were TUAG ≥ 15 seconds and functional dependence		Elective general, cardiac, thoracic, urology and vascular surgery (patients undergoing major elective surgery necessitating postoperative surgical ICU admission)	
Frailty defined as any impairment in activities of daily living (Katz index) or impairment of ambulation or diagnosis of dementia	In-hospital mortality Institutional discharge Mid-term survival	All ages Cardiac surgery	Lee et al.[17]
Groningen frailty indicator	Post-operative delirium	All ages Elective vascular surgery	Pol et al.[86]









Take HOmes Messages THOMS

- · Heterogeneity in ageing
 - Fit ⇔ Frail ⇔ dependent
- Frailty is associated with adverse outcomes in different settings
- Frailty is often clinically recognizable
 - $\bullet \longleftrightarrow$
 - •Variability in operational criteria

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THOMS

- No consensus about the best form of screening/assessment
 - Translation of the G 6th sense for non G caregivers (MD, nurses, ...)
- · Point out on red flags
 - Frailty is dynamic/reversible
 - R/ decision for people at risk of adverse events
 - Settings and patient preferences