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Practical approach on frail older patients attended for acute heart failure.

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Keywords:

Frailty; Older; Acute heart failure.
Highlights

- Frailty is a treatable and potentially reversible syndrome.
- Frailty increases the risk of disability and/or other adverse health outcomes.
- Frailty identification is critical in older patients with acute heart failure.
- Best tool to determine frailty remains to be established.
- Comprehensive Geriatric Assessment is gold standard instrument for diagnosis of frailty.
Abstract

Acute heart failure (AHF) is a multi-organ dysfunction syndrome. In addition to known cardiac dysfunction, non-cardiac comorbidity, frailty and disability are independent risk factors of mortality, morbidity, cognitive and functional decline, and risk of institutionalization. Frailty, a treatable and potential reversible syndrome very common in older patients with AHF, increases the risk of disability and other adverse health outcomes. This position paper highlights the need to identify frailty in order to improve prognosis, the risk-benefits of invasive diagnostic and therapeutic procedures, and the definition of older-person-centred and integrated care plans.
1.-INTRODUCTION

The proportion of people over the age of 65 years is dramatically rising worldwide. It has been estimated that more than 30% of Europeans will be over this age in 2050. One consequence of longer life expectancy is the increased use of health care services. Although greater age does not necessarily imply poorer health, the heterogeneity of the intrinsic capacity varies enormously as a function of age between individuals. Ultimately, the different levels of health in aging are better contemplated in terms of frailty, rather than years alive that are arbitrary and predominately centred on socio-demographic aspects.

Heart failure (HF) is a chronic disease. It has an incidence and prevalence that are highly age-dependent. Three out of 4 patients over the age of 75 years suffer with HF. Acute HF (AHF) is multi-organ dysfunction syndrome that involves cardiac, renal, pulmonary, cerebral, and hepatic injury. It is one of the most frequent causes of visits to emergency departments (ED) and hospitalization. Older patients with AHF require a more complex evaluation and have a worse short-term prognosis across the spectrum of morbidity, cognitive and functional decline, and the risk of institutionalization and mortality, compared to younger adults. Poorer outcomes in older individuals are probably more related to comorbidities, frailty and disability than with chronologic age.

The terms comorbidity, frailty and disability are associated with aging, and although commonly used interchangeably, they are unique entities with different prognoses and health care implications. Comorbidity describes a framework where one specific disease is the primary focus, and the other distinct entities are comorbid conditions modifying the course and the treatment of an individual with the index disease. In HF patients, the prevalence of comorbidity has increased in the last 2 decades and is associated with adverse events.

Disability is defined as difficulty or dependency in performing activities of daily living. Functional disability is prevalent in HF, and the level of functional dependence determines a poor prognosis in older patients with AHF.

Frailty is clinically characterized by a reduction in physiological capacity not necessarily related to a specific disease process and typically involves alterations in multiple systems. Frailty may be reversible or attenuated by interventions. It is more frequent in patients with comorbidity and chronic diseases, particularly HF, than in the general population. It is associated with higher risks of hospitalization, disability and mortality. Approximately 50-70% of older patients admitted for AHF present with some degree of frailty. This contributes to adverse short and long-term outcomes both in those managed medically and in relation to interventional procedures.

Therefore, evaluation of older patients with AHF requires more than assessment of pump failure alone. The identification of frailty and its degree is critical to improve prognosis, optimize
the risk-benefit relationship of invasive diagnostic and therapeutic procedures. An evaluation of frailty is also necessary to accurately define older-person-centred and integrated care plans whose main goal is to maintain or reverse the potential decline in physical and cognitive capacities (add life to your years and not years to your life).

This position paper highlights the need to recognize that frailty, as a syndrome, is different from aging, comorbidity and disability. We will review the definition and diagnosis of frailty and will present a practical evaluation and management of it and other possible concurrent circumstances in older patients attended with AHF.
2.-DEFINITIONS OF FRAILTY

2.1.-Concept.

Frailty is a dynamic and nonlinear process. It describes a state of vulnerability to stressors in terms of systems reserves and capacity of response to stress situations (i.e., decompensation of AHF) in older populations\textsuperscript{13}. This concept may help to identify patients at increased risk of disability and/or other adverse health outcomes (i.e. death, reduced physical performance, functional decline, hospitalization or institutionalization)\textsuperscript{13,25}.

2.2.-Models of frailty.

Two main models have been used to conceptualize frailty. These are based on different theoretical constructs: 1) a biologic syndrome model and 2) an accumulation of deficiencies model\textsuperscript{26}.

a.-Frailty phenotype: This model, based on data from the Cardiovascular Health Study, considers frailty as a biologic syndrome characterized by the presence of 3 or more of 5 components: 1) exhaustion, 2) unintentional weight loss, 3) impaired grip strength, 4) slowness, and 5) low physical activity\textsuperscript{27} (Table 1). Subsequently, some authors have proposed variations of the original model by introducing new criteria (i.e. cognitive impairment) or even reducing the number of components required (i.e. slow gait speed, low physical activity and weight loss)\textsuperscript{25,28}.

b.-Frailty index (FI): This model, derived from the Canadian Study of Health and Aging, is based on deficit accumulation; that is, a measure of the cumulative burden of non-specified age-associated health deficits (i.e. diseases, impairments in cognition, mood, mobility, or function) associated with poor outcomes. The frailty index was originally comprised of 70 measures and conceptualized frailty as a continuum. This model counts disabilities and comorbidities and is able to quantitatively summarize vulnerability\textsuperscript{29}.

Numerous other frailty definitions have been developed, but have largely been based on these two basic conceptual approaches\textsuperscript{30}.
3.-TOOLS FOR IDENTIFYING FRAILTY IN OLDER PATIENTS WITH AHF

3.1.-Screening of frailty in the emergency setting.

Several screening tools for frailty, based on a multi-domain approach, have been proposed\textsuperscript{31}. The tools most frequently studied are the Identification of Seniors at Risk (ISAR)\textsuperscript{32} and Triage Risk Screening Tool (TRST)\textsuperscript{33}. Both are validated for older patients attending in the Emergency Department (ED). These are self-reported (or obtained by a nurse) questionnaires, take approximately 1 minute to administer, and are composed of six items related to different domains. The score ranges from 0 to 6 (0 = low risk, 6 = high risk). A score of two or more is associated with a greater probability of presenting short- and long-term adverse outcomes (hospital readmission, ED revisit, and functional decline and mortality)\textsuperscript{32,33}. These tools have a high sensitivity and relatively low specificity, although the predictive capacity for adverse outcomes of the ISAR seems better than that of the TRST\textsuperscript{31}. Regarding the frailty syndrome, the ISAR has shown a good predictive capacity in identifying frail older patients in the ED\textsuperscript{34}.

The ISAR is considered a useful screening tool for frailty in older patients presenting to the ED\textsuperscript{34} (Table S1 in supplementary material). Taking into account its limitations\textsuperscript{35}, the use of the ISAR alone is inadequate and the cut-off of ≥2 may be useful to select older patients most likely to benefit from geriatric assessment\textsuperscript{34}. Some authors have suggested a higher cut-off point, or that consideration as a continuous variable, may facilitate more efficient use of care resources\textsuperscript{31}. They suggest a cut-off of 3 as better in terms of discriminative capacity for adverse outcomes\textsuperscript{36}.

3.2.-Diagnosis of frailty in inpatient units.

Many of tools have been developed to diagnose frailty in the older population with substantial differences in respect to their ability to predict adverse outcomes\textsuperscript{37}. Nonetheless, the reliability and validity of these tools have rarely been evaluated\textsuperscript{38}. The few studies in which they were tested were epidemiological and their application in the cardiovascular setting, and specifically in HF, is limited\textsuperscript{39} (Table 2). Indeed, the best tool to determine frailty for use in research and clinical practice remains to be established\textsuperscript{38}. Therefore, when selecting a frailty assessment tool, one must consider where it was validated (setting and population), the mode of administration in relation to time-consumption and personnel, and the specialized equipment required\textsuperscript{38}.

Most studies in HF are based on tools derived from the frailty phenotype (Fried Scale) or from some of its domains (Physical Performance test)\textsuperscript{40}. There is less evidence on accumulations of deficits instruments (i.e. Frailty Index-Comprehensive Geriatric Assessment (FI-CGA) or Tilburg Frailty Indicator (TFI)) in the cardiovascular setting\textsuperscript{39,41} (Table 2).

The Fried Scale requires 3 or more criteria (exhaustion, unintentional weight loss, impaired strength, slowness, and low physical activity) for the diagnosis of physical frailty (Table 1)\textsuperscript{27}. It is
important to take into account that any modification from the original scale proposed by Fried et al 2001 may influence the results. In order to compare the different studies available using the frailty phenotype, minimum requirements of the measurements must be reported\textsuperscript{42}.

Several \textbf{Physical Performance tests} have been described, including the Short Physical Performance Battery, gait or walking speed, timed-up-and-go test, handgrip strength, and 4 or 6-minute walk test, each of which may identify physical frailty or preclinical disability in the older population\textsuperscript{43}.

The \textbf{Short Physical Performance Battery (SPPB)} encompasses slowness, weakness, and balance. This test assesses lower extremity function using three physical performance tests that include standing balance (the ability to stand with the feet together in the side-by-side, semi-tandem, and tandem positions), gait speed (time to walk 8 feet or 2.4 metres), and strength and endurance (time to rise from a chair and return to the seated position 5 times). The total score ranges from 0-12, with higher scores indicating better performance\textsuperscript{44}. The SPPB predicts incident activities of daily living disability, worsening mobility and death in older community HF subjects\textsuperscript{45} (Figure 2).

Gait speed is a part of the SPPB, but as a single parameter it has been associated with survival in older adults\textsuperscript{46}. It is an important risk factor for 1-year mortality in older community population with HF\textsuperscript{47}. The 5-m distance is a good balance between the walking speed achieved and cardiopulmonary limitations\textsuperscript{39}.

Alternatively, the \textbf{timed get-up-and-go test} measures the time needed to complete a series of functionally important tasks such as standing up from a chair, walking a short distance, turning around, returning to the chair, and sitting down again\textsuperscript{48}. This test appears to be a reliable and valid functional measurement in patients with HF\textsuperscript{49}. A gait speed <0.8m/s and a timed-up-and-go test >10s are markers of possible frailty in community-dwelling older patients\textsuperscript{50}. In patients with lower limb conditions, the handgrip test may be an alternative option\textsuperscript{51}.

The SPPB is easily applied, preferable to other performance tests (i.e. 4-m walk test, 6-minute walk test, and handgrip strength) in community-dwelling older patients with HF\textsuperscript{45}, and it is currently considered the best instrument to characterize frailty in clinical trials\textsuperscript{52}. In older patients with HF, a total SPPB score ≤4 applied at hospital admission is an independent predictor of the length of hospital stay\textsuperscript{53}. Further, its measurement at hospital discharge is an independent predictor of 30-day mortality\textsuperscript{54} and rehospitalisation\textsuperscript{55}.

\textbf{Comprehensive Geriatric Assessment (CGA)} is the instrument recommended for the evaluation and care of frail older patients in clinical practice\textsuperscript{56}. This holistic evaluation is performed by a multidisciplinary team that usually includes a geriatrician or other physician knowledgeable in the care of older adults, nurse, social worker, pharmacist, and an occupational or physical therapist. This team assesses comorbidity, polypharmacy, and cognitive, functional, nutritional, and socio-economic
areas in order to develop treatment planning and follow-up. The main limitations of the GCA include the need to have experts of diverse disciplines and enough time for the evaluation. The CGA has demonstrated ability to predict in-hospital and long-term adverse outcomes in older patients admitted with HF\textsuperscript{57-60}. Major geriatric syndromes (frailty, severe disability, cognitive, depression) are associated with poor intra-hospital and 1-year results in older patients with acute cardiac conditions\textsuperscript{58}. The CGA is currently the gold standard to detect frailty and should be used when making complex decisions regarding invasive procedures. Because of the limitations mentioned above, some authors have proposed a \textbf{Brief Geriatric Assessment} adapted to non-geriatricians using a combination of screening scales that approach different domains of the patient\textsuperscript{61} (Table 3).

Various \textbf{instruments derived from CGA} have been published (i.e. Multidimensional Prognostic Index [MPI], CGA score, and Edmonton Frail Scale [EFS]) that have a high predictive value of adverse short-term results. The MPI predicts the 1-month mortality in patients aged 65 years and older admitted with AHF\textsuperscript{57}. The CGA score estimates the in-hospital and 2-year mortality in older patients hospitalized for AHF\textsuperscript{59,60}. The EFS tool may be useful to identify frailty when considering a surgical intervention in order to help with pre-operative optimisation\textsuperscript{60}.

With regard to the last-mentioned instrument derived from CGA, \textbf{EFS} is a brief multidimensional tool that may be applied in older admitted patients by non-geriatricians. It includes the domains of cognition, mood, mobility, functional independence, drugs, social support, nutrition, health attitudes, continence, medical disease load and quality of life\textsuperscript{62}. The examination takes less than 5 minutes and the maximum score (total 17) represents the highest level of frailty\textsuperscript{62}. The new version of this scale, the Reported Edmonton Frail Scale (REFS), adapted from the EFS, substitutes the get up and go test with self-reporting of physical function before the current illness. The REFS is scored from 1 to 18\textsuperscript{63}. With respect to the need for major cardiac interventional or surgical decisions in invasive cardiovascular procedures, recent findings have shown that poor agreement among clinicians when using the REFS to diagnose frailty, and therefore a geriatric assessment is recommended in these cases\textsuperscript{64}. 
4.-EVALUATION OF FRAILTY IN OLDER PATIENTS WITH AHF

In all older patients presenting with AHF, the level of frailty must be determined through assessment and monitoring of physical and cognitive status during acute management, during convalescence and, above all, at the time of hospital discharge. Some circumstances, such as the clinical presentation (i.e. delirium, falls or acute functional decline) or the presence of some level of baseline functional dependence in the basic activities of daily living, may be used to indicate possible frailty. Biomarkers are only able to capture single aspects of frailty and are weakly associated with clinically meaningful outcomes. In the absence of universal recommendation as to how to detect vulnerable older patients in clinical practice, we suggest a simplistic approach in the ED and ward settings (Figure 3).

4.1.- Emergency setting.

In the ED, where personnel and time resources are limited, we recommend to screen for frailty, particularly in patients with non-apparent disability discharged directly from the ED. We recommend using ISAR as a continuous variable, with a cut point ≥2 for maximum sensitivity and ≥3 for maximum discrimination, to provide an individualized care plan that includes a CGA program.

4.2.- Inpatient units.

On inpatient units, we recommend that information about comorbidity (Charlson Comorbidity Index) (Table S2 in supplementary material) and baseline functional status (Barthel Index) (Table S3 in supplementary material), be collected at admission to establish the grade of disability. In older patients with established disability (moderate and severe disability), measurement of physical frailty should be focused on basic activities of daily living and mobility. The Barthel index has shown a greater sensitivity to change and may detect the onset of disability earlier than other scores (Table). Previous studies have shown that severe baseline functional dependence (Barthel index <60 points) in older patients attended with AHF is associated with an increase in 30-day mortality, and its inclusion in the HF risk stratification models (Bi-EFFECT) has improved the prediction of 30-day mortality. In this profile patient’s information about other domains (e.g., comorbidity, medications, cognitive, nutritional and social support) should be included since these variables influence short- and long-term prognosis, and care planning decisions.

We recommend using instruments based on the frailty phenotype (i.e. the Fried scale) or physical performance (i.e. SPPB) to diagnose physical frailty in older patients with non-established disability (pre-disabled or mild disability). The presence of frailty is associated with in-hospital, as well as short and long-term outcomes. As mentioned above, there is no single feasible, valid tool to diagnose frailty in AHF inpatients and neither has the best time to perform these tests been
established. Multi-domain tools do not necessarily provide incremental value above single-domain tools, and the ease of implementation may be an important factor for adoption. Taking into account the acute phase of the heart failure condition, self-reported instruments may be more appropriate at hospital admission, while objective performance measures would be better suited at hospital discharge. Finally, some authors have also suggested the addition of cognitive and nutritional status to improve the diagnosis of frailty.

It is also important to monitor the cognitive and functional situation during hospitalization since delirium and acute functional decline are markers of frailty. Delirium is the main manifestation of cognitive frailty and frequently appears in hospitalised elderly patients with cognitive impairment. Its presence in patients with decompensated HF has been associated with 30-day mortality. The Confusion Assessment Method (CAM) is a good tool for the identification of delirium. The CAM for the Intensive Care Unit (CAM-ICU) has shown to have a better capacity to diagnose delirium in older patients in the ED. It is recommended that cognitive status is evaluated, after ruling-out delirium, at the time of the first visit, or failing that, on ward admission. Montreal Cognitive Assessment (MoCA) is the best method to screen for cognitive impairment in patients with HF in both clinical practice and trials.
5.-FRAILTY-BASED MANAGEMENT IN OLDER PATIENTS WITH AHF.

The management of older patients with AHF should be based on clinical guidelines taking into account that older patients, and particularly frail older patients, have often been excluded from clinical trials. We recommend measurement of the degree of frailty as well as the documentation of the presence or absence of concurrent comorbidity and disability (Figure 4).

The identification of frailty involves early treatment of the frailty syndrome and close monitoring of patient capacities during and after hospitalization in order to minimize disability. The most commonly used interventions to treat frailty include, comorbidity optimization, exercise, protein-calorie supplementation, and the development of an individualised care and support plan based on a CGA. Regarding vitamin D3, it was not demonstrated to improve physical performance in spite of the increase in serum 25OHD in older patients with HF. These interventions can reverse frailty, but may have no effect on adverse outcomes (hospitalizations, falls, or performance of activities of daily living) in community-living older persons. The presence of significant functional decline or delirium in non-disabled older patients with AHF should be considered as a high-risk situation that needs CGA. Frailty should be determined with the currently used risk models for decision making. Regarding invasive procedures, the identification of frailty, using frailty criteria and performance tests (i.e. 5m-gait speed and the timed-get up-and-go test) has helped to predict short- and long-term adverse events in patients undergoing transcatheter aortic valve replacement, cardiac surgery, cardiac resynchronization therapy and post percutaneous coronary interventions.

The presence of comorbidities and renal failure may make clinical (i.e. chronic obstructive pulmonary disease (COPD) and biochemical (i.e. renal failure) diagnosis of AHF even more difficult in frail older patients. Some comorbidities such as anaemia, renal failure and hyperglycaemia, may influence the short and long-term prognosis. The treatment of concurrent conditions in the frail older patients with AHF should be optimized by balancing the risk-benefit relation (prioritizations, interactions or contraindications) and making adjustments according to creatinine clearance (MDRD-4). Polypharmacy should be minimized because of the increased risk of adverse events and the consequence of potentially reduced adherence. The application of evidence-based medication review checklists (e.g. STOPP/START criteria) can be helpful to reduce inappropriate medicine use. Regarding heath care, multi-provider or multi-settings should be avoided, or failing this, they should be well coordinated, with close monitoring of active morbidities during both hospitalization and after discharge.

Disabled patients represent the highest risk scenario and require more complex decision-making regarding treatment and care planning. One out of three patients aged 85 years and older (one of six if ≥75 years) attending with AHF in the ED has a moderate or severe disability. To facilitate
the determination of frailty, we suggest to distinguishing between patients with middle or moderate and severe disability. In moderate disability there may be a thin line dividing consideration for therapeutic invasive procedures and their indications (Bathel index 90-40 points). These decisions should be based on CGA integrating risk scores, and frailty and disability components.

Palliative care, ethical constructs, advanced directives, and the rationalisation of medications should be considered in patients with non-acute severe disability. There is no evidence to guide end-of-life decisions for older patients with HF. The usual medications such as beta-blockers, diuretics, angiotensin-converting enzyme inhibitors, angiotensin receptor blockers, nitroglycerin and digoxin are recommended to maintain symptom relief and improve quality of life94. Other treatments such as lipid-lowering medications and antiplatelet agents, with the exception of aspirin, may never be indicated and anticoagulants and antiarrhythmics may rarely be appropriate95.

Regarding the transition of care, the discharge of all frail older patients with AHF should include a comprehensive care and support plan. This should involve plans for optimisation and maintenance, self-care, escalation (what to look for and who to call), and emergency responses that may include whether or not hospital care is appropriate/desirable and what alternatives are in place50. All these aspects are important in order to improve subjective and objective quality of life in older patients with AHF96.
6.-CONCLUSIONS.

In conclusions, 1) AHF is a multi-organ dysfunction syndrome. In addition to cardiac, renal, pulmonary, cerebral, and hepatic injuries, as well as non-cardiac comorbidity, frailty and disability are independent factors predicting mortality, morbidity, cognitive and functional decline, and the risk of institutionalization in older patients with AHF; 2) Frailty (or state of vulnerability to stressors) is a treatable and potentially reversible syndrome which increases the risk of disability and/or other adverse health outcomes; 3) Frailty identification is critical in older patients with AHF in order to improve the stratification of prognosis (disposition), the evaluation of the risk-benefits of invasive diagnostic and therapeutic procedures and the development of older-person-centred and integrated care plans (person-centred coordinated care) which have the main goal of maintaining or reversing potential declines in physical and cognitive capacities; 4) Though the best tool to determine frailty for use in research and clinical practice remains to be established, we recommend the ISAR for the screening of frailty in ED, and the Fried phenotype (i.e. Fried Scale) and Physical Performance Test (i.e. SPPB) for the diagnosis of frailty during the hospitalization of older patients with AHF; 5) CGA (or instruments derived from CGA) is the instrument recommended for the evaluation and care of frail older patients in clinical practice.
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20.-McMurray JJ, Adamopoulos S, Anker SD, Auricchio A, Böhm M, Dickstein K, et al. ESC Guidelines for the diagnosis and treatment of acute and chronic heart failure 2012: The Task Force for the Diagnosis and Treatment of Acute and Chronic Heart Failure 2012 of the European Society of Cardiology. Developed in collaboration with the Heart Failure Association (HFA) of the ESC. Eur Heart J. 2012;33:1787-847.


**TABLES AND FIGURES.**

### Table 1.-Fried Criteria.

<table>
<thead>
<tr>
<th></th>
<th>Original frailty phenotype (as proposed by Fried et al. 2001)</th>
<th>Self-Reported Deficit</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Exhaustion</strong></td>
<td>How often in the last week did you feel this way? (a) I felt that everything I did was an effort; (b) I could not get going</td>
<td>Self-report of fatigue or felt unusually tired or weak in the past month.</td>
</tr>
<tr>
<td></td>
<td>A moderate amount of the time (3–4 days) or most of the time = 1; rarely or none of the time (&lt;1 day) or some or a little of the time (1–2 days) = 0</td>
<td></td>
</tr>
<tr>
<td><strong>Weight loss</strong></td>
<td>In the last year, have you lost more than 5kg unintentionally (i.e., not due to dieting or exercise)? yes=1, no=0</td>
<td>Self-report weight loss &gt; 5kg unintentionally in the past year.</td>
</tr>
<tr>
<td><strong>Physical activity</strong></td>
<td>Minnesota Leisure Time Activity Questionnaire (past 2 weeks): walking, chores (moderately strenuous), mowing the lawn, raking, gardening, hiking, jogging, biking, exercise cycling, dancing, aerobics, bowling, golf, singles tennis, doubles tennis, racquetball, calisthenics, swimming. Deficit given to (adjusted by sex) <strong>-Men:</strong> Kcals/week &lt; 383; <strong>-Women:</strong> Kcals/week &lt; 270</td>
<td>Self-report frequency and duration of physical activities.</td>
</tr>
<tr>
<td></td>
<td>Average grip strength score in dominant hand (3 trials) using JAMAR hand held dynamometer Deficit given to (adjusted by sex and BMI quartile based on CHS population by Fried et al. 2001) <strong>-Men:</strong> BMI ≤24kg and strength &lt;29kg; BMI 24.1–26 and strength &lt;30; BMI 26.1–28kg and strength &lt;30kg; BMI &gt;28 and strength &lt;32 kg <strong>-Women:</strong> BMI ≤23kg and strength &lt;17kg; BMI 23.1–26 and strength &lt;17.3; BMI 26.1–29kg and strength &lt;18kg; BMI &gt;29 and strength &lt;21kg</td>
<td>Self-report of difficulty standing up from a chair.</td>
</tr>
<tr>
<td><strong>Walking time</strong></td>
<td>Walking speed score (15 ft (5m) test, usual pace, one trial) Deficit given to (adjusted by sex and median height based on CHS population by Fried et al. 2001) <strong>-Men:</strong> height ≤173cm and speed ≤0.6531 m/s; height &gt;173cm and speed ≤0.762 m/s <strong>-Women:</strong> height ≤159cm and speed ≤0.6531 m/s; height &gt;159cm and speed ≤0.762 m/s</td>
<td>Self-report of any difficulty for walking 100m.</td>
</tr>
</tbody>
</table>

5 items: 0 deficits: nonfrail; 1-2 deficits: prefrail; ≥3 deficits: frail.
## Table 2: Principal studies about frailty in older patients with acute heart failure.

<table>
<thead>
<tr>
<th>Setting</th>
<th>Items</th>
<th>Score (cut-off)</th>
<th>Administer</th>
<th>Domains</th>
<th>Subjects</th>
<th>Outcomes</th>
</tr>
</thead>
<tbody>
<tr>
<td>Emergency Department</td>
<td></td>
<td></td>
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</tr>
<tr>
<td>ISAR Mc.Clusker 1999</td>
<td>6</td>
<td>0-6 (≥ 2 = frailty)</td>
<td>Self-reported</td>
<td>Cognition, ADL, Medications, Vision, Recent hospitalization</td>
<td>Patients ≥ 65 years discharged from ED</td>
<td>30-day mortality 30-day hospital readmission 30-day functional decline</td>
</tr>
<tr>
<td>SPPB Volpato 2008 2011</td>
<td>3</td>
<td>0-12 (&lt; 5 = frailty)</td>
<td>Physical performance</td>
<td>5-m gait speed test Chair rise test Balance test</td>
<td>Patients ≥ 65 years admitted for AHF</td>
<td>Length of stay Incident disability 15-month mortality</td>
</tr>
<tr>
<td>TFI Uchmanowicz I 2015</td>
<td>15</td>
<td>0-15 (≥5 = frailty)</td>
<td>Self-reported</td>
<td>Physical (8), Psychological (4), Social (3).</td>
<td>Patients ≥ 60 years admitted for AHF</td>
<td>Self-care capabilities Health-related quality of life</td>
</tr>
<tr>
<td>MPI Pilotto 2010</td>
<td>63</td>
<td>0-1 (0.34–0.66 = moderate risk; 0.67-1 = high risk)</td>
<td>Data abstracted out of CGA by geriatrician</td>
<td>Cognition, ADL, Nutrition, Comorbidities, Medications, Decubitus, Social support</td>
<td>Patients ≥ 65 years admitted for AHF</td>
<td>1-month mortality</td>
</tr>
<tr>
<td>CGA Score Rodriguez Pascual 2012 2014</td>
<td>5</td>
<td>0-10 (≥2: lower risk; 3-4; 5-6; ≥7: higher risk)</td>
<td>Data abstracted out of CGA by geriatrician</td>
<td>Cognition, ADL, Mobility, Comorbidity, Medications</td>
<td>Patients ≥ 75 years admitted for AHF</td>
<td>In-hospital mortality 2-year mortality</td>
</tr>
</tbody>
</table>

IADL: instrumental activities of daily living
Table 3.-Brief Geriatric Assessment based on Comprehensive Geriatric Assessment.

<table>
<thead>
<tr>
<th>Domain</th>
<th>Tool</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cognitive</td>
<td>Montreal Cognitive Assessment (MOCA)</td>
</tr>
<tr>
<td>Depression</td>
<td>5-item Geriatric Depression Scale (5-GDS)</td>
</tr>
<tr>
<td>Functional</td>
<td>Lawton index (LI) (8 instrumental activities of daily living)</td>
</tr>
<tr>
<td></td>
<td>Barthel index (BI) (8 basic activities of daily living and 2 of mobility).</td>
</tr>
<tr>
<td>Nutrition</td>
<td>Mini-Nutritional Status – Short Form (MNA-SF)</td>
</tr>
<tr>
<td></td>
<td>Serum albumin</td>
</tr>
<tr>
<td>Polypharmacy</td>
<td>START and STOPP Criteria</td>
</tr>
<tr>
<td>Comorbidity</td>
<td>Charlson Comorbidity Index</td>
</tr>
<tr>
<td>Hearing</td>
<td>Whispering test</td>
</tr>
<tr>
<td>Visual</td>
<td>Snellen test</td>
</tr>
<tr>
<td>Socio-economic</td>
<td>Gijon’s social-familial evaluation scale (SFES)</td>
</tr>
</tbody>
</table>
### Identification of Seniors At Risk (ISAR) Tool

<table>
<thead>
<tr>
<th>Sensorial</th>
<th>Drugs</th>
</tr>
</thead>
<tbody>
<tr>
<td>In general, do you see well?</td>
<td></td>
</tr>
<tr>
<td>• No = 1</td>
<td></td>
</tr>
<tr>
<td>• Yes = 0</td>
<td></td>
</tr>
<tr>
<td>Do you take more than three different medications every day?</td>
<td></td>
</tr>
<tr>
<td>• No = 0</td>
<td></td>
</tr>
<tr>
<td>• Yes = 1</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Mental</th>
<th>Use of hospital services</th>
</tr>
</thead>
<tbody>
<tr>
<td>In general, do you have serious problems with your memory?</td>
<td></td>
</tr>
<tr>
<td>• No = 0</td>
<td></td>
</tr>
<tr>
<td>• Yes = 1</td>
<td></td>
</tr>
<tr>
<td>Have you been hospitalized for one or more nights during the past 6 months (excluding a stay in ED)?</td>
<td></td>
</tr>
<tr>
<td>• No = 0</td>
<td></td>
</tr>
<tr>
<td>• Yes = 1</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Functional</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Before the illness that brought you to the ED, did you need someone to help you on a regular basis?</td>
<td></td>
</tr>
<tr>
<td>• No = 0</td>
<td></td>
</tr>
<tr>
<td>• Yes = 1</td>
<td></td>
</tr>
<tr>
<td>Since the illness that brought you to ED, have you needed more help than usual to take care of yourself?</td>
<td></td>
</tr>
<tr>
<td>• No = 0</td>
<td></td>
</tr>
<tr>
<td>• Yes = 1</td>
<td></td>
</tr>
</tbody>
</table>

**Total Score:**

*Answer in bold = 1; ED = Emergency Department.*
Table S2.-Charlson Comorbidity Index.

<table>
<thead>
<tr>
<th>Charlson Comorbidity Index</th>
<th>Neurologic disorders</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Cardiovascular system</strong></td>
<td></td>
</tr>
<tr>
<td>Myocardial infarct</td>
<td>Dementia</td>
</tr>
<tr>
<td>• Yes = 1</td>
<td>• Yes = 1</td>
</tr>
<tr>
<td>Congestive heart failure</td>
<td>Cerebrovascular disease</td>
</tr>
<tr>
<td>• Yes = 1</td>
<td>• Without hemiplegia = 1</td>
</tr>
<tr>
<td>Peripheral vascular disease</td>
<td>• Hemiplegia = 2</td>
</tr>
<tr>
<td>• Yes = 1</td>
<td></td>
</tr>
<tr>
<td><strong>Respiratory system</strong></td>
<td></td>
</tr>
<tr>
<td>Chronic pulmonary disease</td>
<td>Connective tissue disease</td>
</tr>
<tr>
<td>• Yes = 1</td>
<td>• Yes = 1</td>
</tr>
<tr>
<td><strong>Kidney and urinary tract</strong></td>
<td></td>
</tr>
<tr>
<td>Moderate or severe renal disease</td>
<td>AIDS</td>
</tr>
<tr>
<td>• Yes = 2</td>
<td>• Yes = 6</td>
</tr>
<tr>
<td><strong>Gastrointestinal system</strong></td>
<td></td>
</tr>
<tr>
<td>Ulcer disease</td>
<td>Leukemia</td>
</tr>
<tr>
<td>• Yes = 1</td>
<td>• Yes = 2</td>
</tr>
<tr>
<td>Liver disease</td>
<td>Lymphoma</td>
</tr>
<tr>
<td>• Mild = 1</td>
<td>• Yes = 2</td>
</tr>
<tr>
<td>• Moderate or severe = 3</td>
<td>Malignant solid tumor</td>
</tr>
<tr>
<td><strong>Endocrinology</strong></td>
<td>• Non-metastatic = 2</td>
</tr>
<tr>
<td>Diabetes</td>
<td>• Metastatic = 6</td>
</tr>
<tr>
<td>• Without end-organ damage = 1</td>
<td></td>
</tr>
<tr>
<td>• With end-organ damage = 2</td>
<td></td>
</tr>
<tr>
<td><strong>Total Score:</strong></td>
<td></td>
</tr>
</tbody>
</table>
### Table S3.- Barthel Index.

<table>
<thead>
<tr>
<th>Basic activities of daily living</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Bathing (0-5)</strong></td>
<td><strong>Grooming (0-5)</strong></td>
</tr>
<tr>
<td>0 = dependent</td>
<td>0 = needs help with personal care</td>
</tr>
<tr>
<td>5 = independent (or in shower)</td>
<td>5 = independent face / hair / teeth / shaving</td>
</tr>
<tr>
<td><strong>Dressing (0-10)</strong></td>
<td><strong>Toilet use (0-10)</strong></td>
</tr>
<tr>
<td>0 = dependent</td>
<td>0 = dependent</td>
</tr>
<tr>
<td>5 = needs help but can do about half unaided</td>
<td>5 = needs some help but can do something alone</td>
</tr>
<tr>
<td>10 = independent (including buttons, zips,...)</td>
<td>10 = independent (on and off, dressing, wiping)</td>
</tr>
<tr>
<td><strong>Transfer from bed to chair (0-15)</strong></td>
<td><strong>Feeding (0-10)</strong></td>
</tr>
<tr>
<td>0 = dependent – no sitting balance</td>
<td>0 = dependent</td>
</tr>
<tr>
<td>5 = mayor help (physical, one or two people)</td>
<td>5 = needs help (cutting, spreading butter,...)</td>
</tr>
<tr>
<td>10 = minor help (verbal o physical)</td>
<td>10 = independent</td>
</tr>
<tr>
<td>15 = independent</td>
<td></td>
</tr>
<tr>
<td><strong>Continence Bowels (0-10)</strong></td>
<td><strong>Continence Bladder (0-10)</strong></td>
</tr>
<tr>
<td>0 = incontinent</td>
<td>0 = incontinent</td>
</tr>
<tr>
<td>5 = occasional accident</td>
<td>5 = occasional accident</td>
</tr>
<tr>
<td>10 = continent</td>
<td>10 = continent</td>
</tr>
<tr>
<td><strong>Mobility</strong></td>
<td><strong>Stairs (0-10)</strong></td>
</tr>
<tr>
<td><strong>Mobility (0-15)</strong></td>
<td>0 = unable</td>
</tr>
<tr>
<td>0 = immobile</td>
<td>5 = needs help</td>
</tr>
<tr>
<td>5 = wheelchair independent</td>
<td>10 = independent to go up and down</td>
</tr>
<tr>
<td>10 = needs help of one person</td>
<td></td>
</tr>
<tr>
<td>15 = independent (may use any aid)</td>
<td></td>
</tr>
</tbody>
</table>

Total Score: 0-100
Figure 1.-Practical approach to assess the vulnerability in older patients with acute heart failure.

- **Frailty phenotype / Physical Performance Assessment**
  - 100 BI: Very fit, Well and Managing well
  - 100-90 BI: Vulnerable and Mild Frail

- **Basic ADL / Mobility dependence Assessment**
  - 90-40 BI: Moderate and Severely Frail
  - 40-0 BI: Very Severely and Terminally Frail

**Clinical Frailty Scale**

**Instrumental activities of daily living (IADL):** ability to use telephone, mode of transportation, ability to handle finances, responsibility for own medications, shopping, food preparation, housekeeping, laundry.

**Basic activities of daily living (BADL):** bathing, dressing, transferring, toileting, continence, and feeding.

**Mobility:** physical activity performed without help; requires a person, walking frame, or wheelchair; bedridden.
Figure 2.- Short Physical Performance Battery.

*See the videos in the supplementary electronic material.
Figure 3.- Approach to assess the frailty in older patients with acute heart failure.

- Patients with Acute Heart Failure
- Screening of frailty
  - Age 65 years and older.
  - Clinical presentation (i.e. delirium, falls or acute functional decline).
  - Some level of baseline functional dependence in the basic ADL.
- Emergency Department Identification of Seniors at Risk (ISAR)
  \( \geq 2 = \text{possible and } \geq 3 = \text{probable frailty} \)
  \( 6 = \text{the highest risk of frailty} \)
- Diagnosis of frailty
  - Fried Scale (FS) or Short Performance Physical Battery (SBPP)
    - Fried Scale \( \geq 3 = \text{frailty} \)
    - Performance test (SPPB): \(< 5 = \text{frailty} \)
- Comprehensive Geriatric Assessment (CGA)*

*Instruments derived from CGA: Multidimensional Prognostic Index, CGA score, Edmonton Frail Scale and Brief Geriatric Assessment.
Figure 4.-Management of older patients with acute heart failure based on frailty.

Comorbidity
- Diagnosis of AHF more complex (i.e. chronic pulmonary disease)
- Early detection and treatment of the active comorbidities (i.e. hyperglycaemia, anaemia, renal failure, chronic pulmonary disease,...)
- Minimize polypharmacy (i.e. START and STOPP).
- Avoid multi-provider and multi-setting care.
- Risk for higher degree of frailty or disability.
- Close monitoring of morbidities during hospitalization and after discharge.

Frailty
- Diagnosis of AHF complex (i.e. physiological age changes).
- Minimize the risk of delirium and functional decline.
- Early detection and treatment of frailty syndrome (i.e. exercise, nutrition supplementation, individualized care plan,...)
- Minimize risk for disability.
- High risk of short and long adverse outcomes.
- Assess the need for rehabilitative, community or supportive service.
- Close monitoring of capacities during hospitalization and after discharge.

Disability
- Diagnosis of AHF more complex (i.e. dementia)
- Risk of delirium and functional decline.
- Early detection and treatment of acute decline intrinsic capacities (i.e. rehabilitation, cognitive stimulation,...)
- Assess risk-benefit relation of invasive procedures.
- Very high risk of short and long adverse outcomes.
- Evaluate the need for palliative care (severe disability).
- Assess the need for long-term care.
- Minimize risk for emergency visits and hospitalization.