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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 aspects¹.

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 HF². 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^{3,4}. 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^{5,6}. Poorer outcomes in older individuals are probably more related to comorbidities, frailty and disability than with chronologic age⁵.

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^{7,8} (Figure 1).

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 disease⁹. In HF patients, the prevalence of comorbidity has increased in the last 2 decades¹⁰ and is associated with adverse events¹¹.

Disability is defined as difficulty or dependency in performing activities of daily living⁷. Functional disability is prevalent in HF¹⁰, and the level of functional dependence determines a poor prognosis in older patients with AHF¹².

Frailty is clinically characterized by a reduction in physiological capacity not necessarily related to a specific disease process and typically involves alterations in multiple systems¹³. Frailty may be reversible or attenuated by interventions¹³. It is more frequent in patients with comorbidity and chronic diseases¹⁰, particularly HF, than in the general population¹⁴. It is associated with higher risks of hospitalization, disability and mortality¹⁵. Approximately 50-70% of older patients admitted for AHF present with some degree of frailty¹⁶⁻¹⁸. This contributes to adverse short and long-term outcomes both in those managed medically and in relation to interventional procedures¹⁸⁻¹⁹.

Therefore, evaluation of older patients with AHF requires more than assessment of pump failure alone²⁰⁻²⁴. 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 FRAILITY

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 populations¹³. 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)^{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 model²⁶.

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 activity²⁷ (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)^{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 vulnerability²⁹.

Numerous other frailty definitions have been developed, but have largely been based on these two basic conceptual approaches³⁰.

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 proposed³¹. The tools most frequently studied are the Identification of Seniors at Risk (ISAR)³² and Triage Risk Screening Tool (TRST)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)^{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 TRST³¹. Regarding the frailty syndrome, the ISAR has shown a good predictive capacity in identifying frail older patients in the ED³⁴.

The **ISAR** is considered a useful screening tool for frailty in older patients presenting to the ED³⁴ (Table S1 in supplementary material). Taking into account its limitations³⁵, 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 assessment³⁴. Some authors have suggested a higher cut-off point, or that consideration as a continuous variable, may facilitate more efficient use of care resources³¹. They suggest a cut-off of 3 as better in terms of discriminative capacity for adverse outcomes³⁶.

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 outcomes³⁷. Nonetheless, the reliability and validity of these tools have rarely been evaluated³⁸. The few studies in which they were tested were epidemiological and their application in the cardiovascular setting, and specifically in HF, is limited³⁹ (Table 2). Indeed, the best tool to determine frailty for use in research and clinical practice remains to be established³⁸. 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 required³⁸.

Most studies in HF are based on tools derived from the frailty phenotype (Fried Scale) or from some of its domains (Physical Performance test)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^{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)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 reported⁴².

Several **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 population⁴³.

The **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 performance⁴⁴. The SPPB predicts incident activities of daily living disability, worsening mobility and death in older community HF subjects⁴⁵ (Figure 2).

Gait speed is a part of the SPPB, but as a single parameter it has been associated with survival in older adults⁴⁶. It is an important risk factor for 1-year mortality in older community population with HF⁴⁷. The 5-m distance is a good balance between the walking speed achieved and cardiopulmonary limitations³⁹.

Alternatively, the **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 again⁴⁸. This test appears to be a reliable and valid functional measurement in patients with HF⁴⁹. A gait speed <0.8m/s and a timed-up-and-go test >10s are markers of possible frailty in community-dwelling older patients⁵⁰. In patients with lower limb conditions, the handgrip test may be an alternative option⁵¹.

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 HF⁴⁵, and it is currently considered the best instrument to characterize frailty in clinical trials⁵². In older patients with HF, a total SPPB score ≤ 4 applied at hospital admission is an independent predictor of the length of hospital stay⁵³. Further, its measurement at hospital discharge is an independent predictor of 30-day mortality⁵⁴ and rehospitalisation⁵⁵.

Comprehensive Geriatric Assessment (CGA) is the instrument recommended for the evaluation and care of frail older patients in clinical practice⁵⁶. 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 HF⁵⁷⁻⁶⁰. Major geriatric syndromes (frailty, severe disability, cognitive, depression) are associated with poor intra-hospital and 1-year results in older patients with acute cardiac conditions⁵⁸. 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 **Brief Geriatric Assessment** adapted to non-geriatricians using a combination of screening scales that approach different domains of the patient⁶¹ (Table 3).

Various **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 AHF⁵⁷. The CGA score estimates the in-hospital and 2-year mortality in older patients hospitalized for AHF^{59,60}. The EFS tool may be useful to identify frailty when considering a surgical intervention in order to help with pre-operative optimisation⁵⁰.

With regard to the last-mentioned instrument derived from CGA, **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 life⁶². The examination takes less than 5 minutes and the maximum score (total 17) represents the highest level of frailty⁶². 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 18⁶³. 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 cases⁶⁴.

4.-EVALUATION OF FRAILITY 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 discharge⁵⁰. 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 frailty⁵⁰. Biomarkers are only able to capture single aspects of frailty and are weakly associated with clinically meaningful outcomes⁶⁵. 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)⁶⁶ and baseline functional status (Barthel Index) (Table S3 in supplementary material)⁶⁷, 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 scores⁸ (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 mortality¹², and its inclusion in the HF risk stratification models (Bi-EFFECT) has improved the prediction of 30-day mortality⁶⁸. 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 decisions⁵⁸⁻⁶⁰.

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)⁶⁹. The presence of frailty is associated with in-hospital, as well as short and long-term outcomes⁵³⁻⁵⁵. 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 frailty¹³.

It is also important to monitor the cognitive and functional situation during hospitalization since delirium and acute functional decline are markers of frailty⁵⁰. 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 mortality⁷⁰. The Confusion Assessment Method (CAM) is a good tool for the identification of delirium⁷¹. The CAM for the Intensive Care Unit (CAM-ICU)⁷² has shown to have a better capacity to diagnose delirium in older patients in the ED⁷³. 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^{74,75} in both clinical practice and trials⁵².

5.-FRAILITY-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^{76,77}. 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^{50,59}. Regarding vitamin D3, it was not demonstrated to improve physical performance in spite of the increase in serum 25OHD in older patients with HF⁷⁸. 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 persons⁷⁹. 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 criteria⁸⁰⁻⁸⁴ and performance tests (i.e. 5m-gait speed^{85,86} and the timed-get up-and-go test⁸⁷) has helped to predict short- and long-term adverse events in patients undergoing transcatheter aortic valve replacement^{80,82}, cardiac surgery⁸³, cardiac resynchronization therapy⁸⁴ and post percutaneous coronary interventions⁸¹.

The presence of comorbidities and renal failure may make clinical (i.e. chronic obstructive pulmonary disease (COPD)⁸⁸ and biochemical (i.e. renal failure)⁸⁹ 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 prognosis⁹⁰⁻⁹³. 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 use⁵⁰. Regarding health 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 disability⁵. 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.

REFERENCES.

- 1.-World Health Organization. World report on ageing and health, 2015.
- 2.-Llorens P, Escoda R, Miró O, Herrero Puente P, Martín-Sánchez FJ, Jacob J, et al. Characteristics and clinical course of patients with acute heart failure and the therapeutic measures applied in Spanish emergency departments: based on the EAHFE registry (Epidemiology of Acute Heart Failure in Emergency Departments). *Emergencias*. 2015;27:11-22.
- 3.-Go AS, Mozaffarian D, Roger VL, Benjamin EJ, Berry JD, Blaha MJ, et al. Heart disease and stroke statistics – 2014 update: A report from the American Heart Association. *Circulation* 2014; 129: e28–e292.
- 4.-Heidenreich PA, Albert NM, Allen LA, Bluemke DA, Butler J, Fonarow GC, et al. Forecasting the impact of heart failure in the United States: A policy statement from the American Heart Association. *Circ Heart Fail* 2013; 6: 606–619.
- 5.-Martín-Sánchez FJ, Marino-Genicio R, Rodríguez-Adrada E, Jacob J, Herrero P, Miró O, et al. Management of acute heart failure in spanish emergency departments based on age. *Rev Esp Cardiol (Engl Ed)*. 2013;66:715-20.
- 6.-Teixeira A, Parenica J, Park JJ, Ishihara S, AlHabib KF, Laribi S, et al. Clinical presentation and outcome by age categories in acute heart failure: results from an international observational cohort. *Eur J Heart Fail*. 2015;17:1114-23.
- 7.-Fried LP, Ferrucci L, Darer J, Williamson JD, Anderson G: Untangling the concepts of disability, frailty and comorbidity: implications for improved targeting and care. *J Gerontol Biol Sci Med Sci* 2004;59:255–263.
- 8.-Abizanda P, Romero L, Sanchez-Jurado PM, Martinez-Reig M, Gomez-Arnedo L, Alfonso SA. Frailty and mortality, disability and mobility loss in a Spanish cohort of older adults: the FRADEA study. *Maturitas*. 2013; 74:54-60.
- 9.-Boyd CM, Ritchie CS, Tipton EF, Studenski SA, Wieland D. From Bedside to Bench: summary from the American Geriatrics Society/National Institute on Aging Research Conference on Comorbidity and Multiple Morbidity in Older Adults. *Aging Clin Exp Res*. 2008;20:181-8.
- 10.-Wong CY, Chaudhry SI, Desai MM, Krumholz HM. Trends in comorbidity, disability, and polypharmacy in heart failure. *Am J Med*. 2011;124:136-43.
- 11.-Rushton CA, Satchithananda DK, Jones PW, Kadam UT. Non-cardiovascular comorbidity, severity and prognosis in non-selected heart failure populations: A systematic review and meta-analysis. *Int J Cardiol*. 2015;196:98-106.
- 12.-Miro O, Llorens P, Martin-Sanchez FJ, Herrero P, Pavon J, Perez-Dura MJ, et al. Short-term prognostic factors in elderly patients seen in emergency departments for acute heart failure. *Rev Esp Cardiol*. 2009;62:757-64.

- 13.-Rodríguez-Mañas L, Fearnt C, Mann G, Viña J, Chatterji S, Chodzko-Zajko W, et al. Searching for an operational definition of frailty: a Delphi method based consensus statement: the frailty operative definition-consensus conference project. *J Gerontol A Biol Sci Med Sci*. 2013;68:62-7.
- 14.-Altimir S, Lupón J, González B, Prats M, Parajón T, Urrutia A, et al. Sex and age differences in fragility in a heart failure population. *Eur J Heart Fail*. 2005;7:798-802.
- 15.-McNallan SM, Chamberlain AM, Gerber Y, Singh M, Kane RL, Weston SA, et al. Measuring frailty in heart failure: a community perspective. *Am Heart J*. 2013;166:768-74.
- 16.-Chiarantini D, Volpato S, Sioulis F, et al. Lower extremity performance measures predict long-term prognosis in older patients hospitalized for heart failure. *J Card Fail* 2010;16:390-5.
- 17.-Vidán MT, Sánchez E, Fernández-Avilés F, Serra-Rexach JA, Ortiz J, Bueno H. FRAIL-HF, a study to evaluate the clinical complexity of heart failure in nondependent older patients: rationale, methods and baseline characteristics. *Clin Cardiol*. 2014;37:725-32.
- 18.-Vidán MT, Blaya-Novakova V, Sánchez E, Ortiz J, Serra-Rexach JA, Bueno H. *Eur J Heart Fail*. 2016. In press. doi: 10.1002/ejhf.518.
- 19.-Chaudhry SI, McAvay G, Ning Y, Allore HG, Newman AB, Gill TM. Risk factors for onset of disability among older persons newly diagnosed with heart failure: the Cardiovascular Health Study. *J Card Fail*. 2011;17:764-70.
- 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.
- 21.-Mebazaa A, Yilmaz MB, Levy P, Ponikowski P, Peacock WF, Laribi S, et al. Recommendations on pre-hospital & early hospital management of acute heart failure: a consensus paper from the Heart Failure Association of the European Society of Cardiology, the European Society of Emergency Medicine and the Society of Academic Emergency Medicine. *Eur J Heart Fail*. 2015;17:544-58.
- 22.-Mueller C, Christ M, Cowie M, Cullen L, Maisel AS, Masip J, et al. European Society of Cardiology-Acute Cardiovascular Care Association Position paper on acute heart failure: A call for interdisciplinary care. *Eur Heart J Acute Cardiovasc Care*. 2015. In press.
- 23.-Llorens P, Manito Lorite N, Manzano Espinosa L, Martín-Sánchez FJ, Comín Colet J, Formiga F, et al. Consensus on improving the care integrated of patients with acute heart failure. *Emergencias*. 2015;27:245-66.
- 24.-Miró Ò, Peacock FW, McMurray JJ, Bueno H, Christ M, Maisel AS, et al. European Society of Cardiology - Acute Cardiovascular Care Association position paper on safe discharge of acute

heart failure patients from the emergency department. *Eur Heart J Acute Cardiovasc Care*. 2016. In press.

25.-Sternberg SA, Wershof Schwartz A, Karunananthan S et al. The identification of frailty: A systematic literature review. *J Am Geriatr Soc* 2011;59:2129–38.

26.-Cigolle CT, Ofstedal MB, Tian Z, Blaum CS. Comparing models of frailty: the Health and Retirement Study. *J Am Geriatr Soc*. 2009;57:830-9.

27.-Fried LP, Tangen CM, Walston J et al. Frailty in older adults: Evidence for a phenotype. *J Gerontol A Biol Sci Med Sci* 2001; 56A:M146–M156.

28.-Rothman MD, Leo-Summers L, Gill TM. Prognostic significance of potential frailty criteria. *J Am Geriatr Soc* 2008;56:2211–2216.

29.-Rockwood K, Mitnitski A. Frailty in relation to the accumulation of deficits. *J Gerontol A Biol Sci Med Sci* 2007;62A:722-7.

30.-de Vries NM, Staal JB, van Ravensberg CD, Hobbelen JS, Olde Rikkert MG, Nijhuis-van der Sanden MW. Outcome instruments to measure frailty: a systematic review. *Ageing Res Rev*. 2011;10:104-14.

31.-Warnier RM, van Rossum E, van Velthuisen E, Mulder WJ, Schols JM, Kempen GI. Validity, Reliability and Feasibility of Tools to Identify Frail Older Patients in Inpatient Hospital Care: A Systematic Review. *J Nutr Health Aging*. 2016;20:218-30.

32.-McCusker J, Bellavance F, Cardin S, Trepanier S, Verdon J, Ardman O. Detection of older people at increased risk of adverse health outcomes after an emergency visit: the ISAR screening tool. *J Am Geriatr Soc*. 1999;47:1229–37.

33.-Meldon SW, Mion LC, Palmer RM, Drew BL, Connor JT, Lewicki LJ, et al. A brief risk stratification tool to predict repeat emergency department visits and hospitalizations in older patients discharged from the emergency department. *Acad Emerg Med*. 2003;10:224–32.

34.-Salvi F, Morichi V, Grilli A, Lancioni L, Spazzafumo L, Polonara S, Abbatecola AM, De Tommaso G, Dessi-Fulgheri P, Lattanzio F. Screening for frailty in elderly emergency department patients by using the Identification of Seniors At Risk (ISAR). *J Nutr Health Aging*. 2012;16:313-8.

35.-Yao JL, Fang J, Lou QQ, Anderson RM. A systematic review of the identification of seniors at risk (ISAR) tool for the prediction of adverse outcome in elderly patients seen in the emergency department. *Int J Clin Exp Med*. 2015;8:4778-86.

36.-Singler K, Heppner HJ, Skutetzky A, Sieber C, Christ M and Thiem U. Predictive validity of the identification of seniors at risk screening tool in a german emergency department setting. *Gerontology* 2013; 60: 413-419.

- 37.-Theou O, Brothers TD, Mitntiski A et al. Operationalization of frailty using eight commonly used scales and comparison of their ability to predict all cause mortality. *J Am Geriatr Soc* 2013;61:1537-51.
- 38.-Sutton JL, Gould RL, Daley S, Coulson MC, Ward EV, Butler AM, et al. Psychometric properties of multicomponent tools designed to assess frailty in older adults: A systematic review. *BMC Geriatr*. 2016;16:55.
- 39.-Afilalo J, Alexander KP, Mack MJ, Maurer MS, Green P, Allen LA, et al. Frailty assessment in the cardiovascular care of older adults. *J Am Coll Cardiol*. 2014;63:747-62.
- 40.-Anker MS, von Haehling S, Springer J, Banach M, Anker SD. Highlights of mechanistic and therapeutic cachexia and sarcopenia research 2010 to 2012 and their relevance for cardiology. *Arch Med Sci*. 2013;9:166-71.
- 41.-Uchmanowicz I, Wleklik M, Gobbens RJ. Frailty syndrome and self-care ability in elderly patients with heart failure. *Clin Interv Aging*. 2015;10:871-7.
- 42.-Theou O, Cann L, Blodgett J, Wallace LM, Brothers TD, Rockwood K. Modifications to the frailty phenotype criteria: Systematic review of the current literature and investigation of 262 frailty phenotypes in the Survey of Health, Ageing, and Retirement in Europe. *Ageing Res Rev*. 2015;21:78-94.
- 43.-Guralnik JM, Ferrucci L, Simonsick EM, Salive ME, Wallace RB. Lower-extremity function in persons over the age of 70 years as a predictor of subsequent disability. *N Engl J Med*. 1995;332:556-61.
- 44.-Guralnik JM, Simonsick EM, Ferrucci L, Glynn RJ, Berkman LF, Blazer DG, et al. A short physical performance battery assessing lower extremity function: association with self-reported disability and prediction of mortality and nursing home admission. *J Gerontol*. 1994;49:M85-94.
- 45.-Minnecci C, Mello AM, Mossello E, Baldasseroni S, Macchi L, Cipolletti S, et al. Comparative study of four physical performance measures as predictors of death, incident disability, and falls in unselected older persons: the insufficienza Cardiaca negli Anziani Residenti a Dicomano Study. *J Am Geriatr Soc*. 2015;63:136-41.
- 46.-Studenski S, Perera S, Patel K, Rosano C, Faulkner K, Inzitari M, et al. Gait speed and survival in older adults. *JAMA*. 2011;305:50-8.
- 47.-Lo AX, Donnelly JP, McGwin G Jr, Bittner V, Ahmed A, Brown CJ. Impact of gait speed and instrumental activities of daily living on all-cause mortality in adults ≥ 65 years with heart failure. *Am J Cardiol*. 2015;115:797-801.
- 48.-Mathias S, Nayak US, Isaacs B. Balance in elderly patients: the "get-up and go" test. *Arch Phys Med Rehabil* 1986; 67: 387-9.

- 49.-Hwang R, Morris NR, Mandrusiak A, Mudge A, Suna J, Adsett J, et al. Timed Up and Go Test: A Reliable and Valid Test in Patients With Chronic Heart Failure. *J Card Fail*. 2015. In press. doi: 10.1016/j.cardfail.2015.09.018.
- 50.-Turner G, Clegg A; British Geriatrics Society; Age UK; Royal College of General Practitioners. Best practice guidelines for the management of frailty: a British Geriatrics Society, Age UK and Royal College of General Practitioners report. *Age Ageing*. 2014;43:744-7.
- 51.-Cruz-Jentoft AJ, Baeyens JP, Bauer JM, Boirie Y, Cederholm T, Landi F, et al. Sarcopenia: European consensus on definition and diagnosis: Report of the European Working Group on Sarcopenia in Older People. *Age Ageing*. 2010;39:412-23.
- 52.-Points to consider on frailty: Evaluation instruments for baseline characterisation of clinical trial populations. European Medicines Agency. Committee for Medicinal Products for Human Use, 2015.
- 53.-Volpato S, Cavalieri M, Guerra G, et al. Performance-based functional assessment in older hospitalized patients: feasibility and clinical correlates. *J Gerontol A Biol Sci Med Sci* 2008;63:1393-8.
- 54.-Chiarantini D, Volpato S, Sioulis F, Bartalucci F, Del Bianco L, Mangani I, et al. Lower extremity performance measures predict long-term prognosis in older patients hospitalized for heart failure. *J Card Fail*. 2010;16:390-5.
- 55.-Volpato S, Cavalieri M, Sioulis F, Guerra G, Maraldi C, Zuliani G, et al. Predictive value of the Short Physical Performance Battery following hospitalization in older patients. *J Gerontol A Biol Sci Med Sci*. 2011;66:89-96.
- 56.-Ellis G, Whitehead MA, Robinson D, O'Neill D, Langhorne P. Comprehensive geriatric assessment for older adults admitted to hospital: meta-analysis of randomised controlled trials. *BMJ* 2011; 343: d6553.
- 57.-Pilotto A, Addante F, Franceschi M, Leandro G, Rengo G, D'Ambrosio P, et al. Multidimensional Prognostic Index based on a comprehensive geriatric assessment predicts short-term mortality in older patients with heart failure. *Circ Heart Fail*. 2010;3:14-20.
- 58.-Sánchez E, Vidán MT, Serra JA, Fernández-Avilés F, Bueno H. Prevalence of geriatric syndromes and impact on clinical and functional outcomes in older patients with acute cardiac diseases. *Heart*. 2011;97:1602-6.
- 59.-Rodríguez-Pascual C, Vilches-Moraga A, Paredes-Galán E, Ferrero-Marinez AI, Torrente-Carballido M, Rodríguez-Artalejo F. Comprehensive geriatric assessment and hospital mortality among older adults with decompensated heart failure. *Am Heart J*. 2012;164:756-62.
- 60.-Rodríguez-Pascual C, Paredes-Galan E, Vilches-Moraga A, Ferrero-Martinez AI, Torrente-Carballido M, Rodríguez-Artalejo F. Comprehensive geriatric assessment and 2-year mortality in elderly patients hospitalized for heart failure. *Circ Cardiovasc Qual Outcomes*. 2014;7:251-8.

- 61.-Martín-Sánchez FJ, Fernández Alonso C, Gil Gregorio P. Key points in healthcare of frail elders in the Emergency Department. *Med Clin (Barc)*. 2013;140:24-9.
- 62.-Rolfson DB, Majumdar SR, Tsuyuki RT, Tahir A, Rockwood K. Validity and reliability of the Edmonton Frail Scale. *Age Ageing* 2006;35:526-29.
- 63.-Hilmer SN, Perera V, Mitchell S, Murnion BP, Dent J, Bajorek B, et al. The assessment of frailty in older people in acute care. *Australas J Ageing*. 2009;28:182-8.
- 64.-Hii TB, Lainchbury JG, Bridgman PG. Frailty in acute cardiology: comparison of a quick clinical assessment against a validated frailty assessment tool. *Heart Lung Circ*. 2015;24:551-6.
- 65.-Calvani R, Marini F, Cesari M, Tosato M, Anker SD, von Haehling S, et al. Biomarkers for physical frailty and sarcopenia: state of the science and future developments. *J Cachexia Sarcopenia Muscle*. 2015;6:278-86.
- 66.-Charlson ME, Pompei P, Ales KL, et al. A new method of classifying prognostic comorbidity in longitudinal studies: development and validation. *J Chronic Dis*. 1987;40: 373-383.
- 67.-Mahoney FI, Barthel D. Functional evaluation: the Barthel Index. *Maryland State Med Journal* 1965;14:56-61.
- 68.-Martín-Sánchez FJ, Gil V, Llorens P, Herrero P, Jacob J, Fernández C, et al. Barthel Index-Enhanced Feedback for Effective Cardiac Treatment (BI-EFFECT) Study: contribution of the Barthel Index to the Heart Failure Risk Scoring System model in elderly adults with acute heart failure in the emergency department. *J Am Geriatr Soc*. 2012;60:493-8.
- 69.-Morley JE, Vellas B, van Kan GA, Anker SD, Bauer JM, Bernabei R, et al. Frailty consensus: a call to action. *J Am Med Dir Assoc*. 2013;14:392-7.
- 70.-Rizzi MA, Torres Bonafonte OH, Alquezar A, Herrera Mateo S, Piñera P, Puig M, et al. Prognostic value and risk factors of delirium in emergency patients with decompensated heart failure. *J Am Med Dir Assoc*. 2015;16:799.e1-6.
- 71.-Inouye SK, van Dyck CH, Alessi CA, Balkin S, Siegal AP, Horwitz RI. Clarifying confusion: the confusion assessment method. A new method for detection of delirium. *Ann Intern Med*. 1990;113:941-8.
- 72.-Ely EW, Inouye SK, Bernard GR, Gordon S, Francis J, May L, et al. Delirium in mechanically ventilated patients: validity and reliability of the confusion assessment method for the intensive care unit (CAM-ICU). *JAMA*. 2001;286:2703-10.
- 73.-Van de Meeberg EK, Festen S, Kwant M, Georg RR, Izaks GJ, Ter Maaten JC. Improved detection of delirium, implementation and validation of the CAM-ICU in elderly Emergency Department patients. *Eur J Emerg Med*. 2016. In press.

- 74.-Cameron JD, Gallagher R, Pressler SJ, McLennan SN, Ski CF, Tofler G, et al. Sensitivity and Specificity of a Five-Minute Cognitive Screening Test in Patients With Heart Failure. *J Card Fail.* 2016;22:99-107.
- 75.-Cameron J, Worrall-Carter L, Page K, Stewart S, Ski CF. Screening for mild cognitive impairment in patients with heart failure: Montreal cognitive assessment versus mini mental state exam. *Eur J Cardiovasc Nurs.* 2013;12:252-60.
- 76.-Cherubini A, Oristrell J, Pla X, Ruggiero C, Ferretti R, Diestre G, et al. The persistent exclusion of older patients from ongoing clinical trials regarding heart failure. *Arch Inter Med* 2011;171:550-6.
- 77.-Miró Ò, Gil V, Müller C, Mebazaa A, Bueno H, Martín-Sánchez FJ, et al. How does a clinical trial fit into the real world? The RELAX-AHF study population into the EAHFE registry. *Clin Res Cardiol.* 2015;104:850-60.
- 78.-Boxer RS, Kenny AM, Schmotzer BJ, Vest M, Fiutem JJ, Piña IL. A randomized controlled trial of high dose vitamin D3 in patients with heart failure. *JACC Heart Fail.* 2013;1:84-90.
- 79.-Ng TP, Feng L, Nyunt MS, Feng L, Niti M, Tan BY, et al. Nutritional, Physical, Cognitive, and Combination Interventions and Frailty Reversal Among Older Adults: A Randomized Controlled Trial. *Am J Med.* 2015;128:1225-36.
- 80.-Ewe SH, Ajmone Marsan N, Pepi M, et al. Impact of left ventricular systolic function on clinical and echocardiographic outcomes following transcatheter aortic valve implantation for severe aortic stenosis. *Am Heart J* 2010;160:1113-20.
- 81.-Singh M, Rihal CS, Lennon RJ, Spertus JA, Nair KS, Roger VL. Influence of frailty and health status on outcomes in patients with coronary disease undergoing percutaneous revascularization. *Circ Cardiovasc Qual Outcomes* 2011;4:496-502.
- 82.-Green P, Woglom AE, Genereux P, et al. The impact of frailty status on survival after transcatheter aortic valve replacement in older adults with severe aortic stenosis: a single-center experience. *J Am Coll Cardiol Intv* 2012;5:974-81.
- 83.-Sepelari A, Beggs T, Hassan A, Rigatto C, Shaw-Daigle C, Tangri N, et al. The impact of frailty on outcomes after cardiac surgery: a systematic review. *J Thorac Cardiovasc Surg.* 2014;148:3110-7.
- 84.-Dominguez-Rodriguez A, Abreu-Gonzalez P, Jimenez-Sosa A, Gonzalez J, Caballero-Estevez N, Martín-Casañas FV, et al. The impact of frailty in older patients with non-ischaemic cardiomyopathy after implantation of cardiac resynchronization therapy defibrillator. *Europace.* 2015;17:598-602.
- 85.-Afilalo J, Eisenberg MJ, Morin JF, Bergman H, Monette J, Noiseux N, et al. Gait speed as an incremental predictor of mortality and major morbidity in elderly patients undergoing cardiac surgery. *J Am Coll Cardiol* 2010;56:1668-76.

- 86.-Afilalo J, Mottillo S, Eisenberg MJ, Alexander KP, Noiseux N, Perrault LP, et al. Addition of frailty and disability to cardiac surgery risk scores identifies elderly patients at high risk of mortality or major morbidity. *Circ Cardiovasc Qual Outcomes* 2012;5:222-8.
- 87.-Robinson TN, Eiseman B, Wallace JI, Church SD, McFann KK, Pfister SM, et al. Redefining geriatric preoperative assessment using frailty, disability and co-morbidity. *Ann Surg* 2009;250:449-55.
- 88.-Güder G, Rutten FH. Comorbidity of heart failure and chronic obstructive pulmonary disease: more than coincidence. *Curr Heart Fail Rep.* 2014;11:337-46.
- 89.-Hill SA, Booth RA, Santaguida PL, Don-Wauchope A, Brown JA, Oremus M, et al. Use of BNP and NT-proBNP for the diagnosis of heart failure in the emergency department: a systematic review of the evidence. *Heart Fail Rev.* 2014;19:421-38.
- 90.-Groenveld HF, Januzzi JL, Damman K, van Wijngaarden J, Hillege HL, van Veldhuisen DJ, et al. Anemia and mortality in heart failure patients a systematic review and meta-analysis. *J Am Coll Cardiol.* 2008;52:818-27.
- 91.-Mebazaa A, Gayat E, Lassus J, Meas T, Mueller C, Maggioni A, et al. Association between elevated blood glucose and outcome in acute heart failure: results from an international observational cohort. *J Am Coll Cardiol.* 2013;61:820-9.
- 92.-Damman K, Valente MA, Voors AA, O'Connor CM, van Veldhuisen DJ, Hillege HL. Renal impairment, worsening renal function, and outcome in patients with heart failure: an updated meta-analysis. *Eur Heart J.* 2014;35:455-69.
- 93.-Lee CS, Chien CV, Bidwell JT, Gelow JM, Denfeld QE, Masterson Creber R, et al. Comorbidity profiles and inpatient outcomes during hospitalization for heart failure: an analysis of the U.S. Nationwide inpatient sample. *BMC Cardiovasc Disord.* 2014;14:73.
- 94.-Cruz-Jentoft AJ, Boland B, Rexach L. Drug therapy optimization at the end of life. *Drugs Aging.* 2012;29:511-21.
- 95.-Holmes HM, Sachs GA, Shega JW, Hougham GW, Cox Hayley D, Dale W. Integrating palliative medicine into the care of persons with advanced dementia: identifying appropriate medication use. *J Am Geriatr Soc.* 2008;56:1306-11.
- 96.-Miró O, Escoda R, Martín-Sánchez FJ, Herrero P, Jacob J, Alquézar A, et al. Patients' perception of quality of emergency department care for acute heart failure: the CALPERICA study. *Emergencias.* 2015;27:161-8.

TABLES AND FIGURES.**Table 1.-Fried Criteria.**

| | Original frailty phenotype (as proposed by Fried et al. 2001) | Self-Reported Deficit |
|--------------------------|--|---|
| Exhaustion | 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 A moderate amount of the time (3–4 days) or most of the time = 1; rarely or none of the time (<1 day) or some or a little of the time (1–2 days) = 0 | Self-report of fatigue or felt unusually tired or weak in the past month. |
| Weight loss | In the last year, have you lost more than 5kg unintentionally (i.e., not due to dieting or exercise)? yes=1, no=0 | Self-report weight loss > 5kg unintentionally in the past year. |
| Physical activity | 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) -Men: Kcals/week < 383; -Women: Kcals/week < 270 | Self-report frequency and duration of physical activities. |
| Grip strength | 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) -Men: BMI ≤24kg and strength <29kg; BMI 24.1–26 and strength <30; Men: BMI 26.1-28kg and strength <30kg; BMI24.1–26 and strength <30; BMI >28 and strength <32 kg -Women: BMI ≤23kg and strength <17kg; BMI 23.1–26 and strength <17.3; Men: BMI 26.1-29kg and strength <18kg; BMI >29 and strength <21kg | Self-report of difficulty standing up from a chair. |
| Walking time | 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) -Men: height ≤173cm and speed ≤0.6531 m/s; height>173cm and speed ≤0.762 m/s -Women:height ≤159cm and speed ≤0.6531m/s; height >159cm and speed ≤0.762 m/s | Self-report of any difficulty for walking 100m. |

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.

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

IADL: instrumental activities of daily living

Table 3.-Brief Geriatric Assessment based on Comprehensive Geriatric Assessment.

| Domain | Tool |
|----------------|---|
| Cognitive | Montreal Cognitive Assessment (MOCA) |
| Depression | 5-item Geriatric Depression Scale (5-GDS) |
| Functional | Lawton index (LI) (8 instrumental activities of daily living) Barthel index (BI) (8 basic activities of daily living and 2 of mobility). |
| Nutrition | Mini-Nutritional Status – Short Form (MNA-SF) Serum albumin |
| Polypharmacy | START and STOPP Criteria |
| Comorbidity | Charlson Comorbidity Index |
| Hearing | Whispering test |
| Visual | Snellen test |
| Socio-economic | Gijon's social-familial evaluation scale (SFES) |

SUPPLEMENTARY MATERIAL.

Table S1.-Identification of Senior At Risk Tool.

| Identification of Seniors At Risk (ISAR) Tool | |
|---|---|
| <p>Sensorial</p> <p>In general, do you see well?</p> <ul style="list-style-type: none"> • No = 1 • Yes = 0 | <p>Drugs</p> <p>Do you take more than three different medications every day?</p> <ul style="list-style-type: none"> • No = 0 • Yes = 1 |
| <p>Mental</p> <p>In general, do you have serious problems with your memory?</p> <ul style="list-style-type: none"> • No = 0 • Yes = 1 | <p>Use of hospital services</p> <p>Have you been hospitalized for one or more nights during the past 6 months (excluding a stay in ED)?</p> <ul style="list-style-type: none"> • No = 0 • Yes = 1 |
| Functional | |
| <p>Before the illness that brought you to the ED, did you need someone to help you on a regular basis?</p> <ul style="list-style-type: none"> • No = 0 • Yes = 1 | <p>Since the illness that brought you to ED, have you needed more help than usual to take care of yourself?</p> <ul style="list-style-type: none"> • No = 0 • Yes = 1 |
| <p>Total Score:</p> | |

*Answer in bold = 1; ED = Emergency Department.

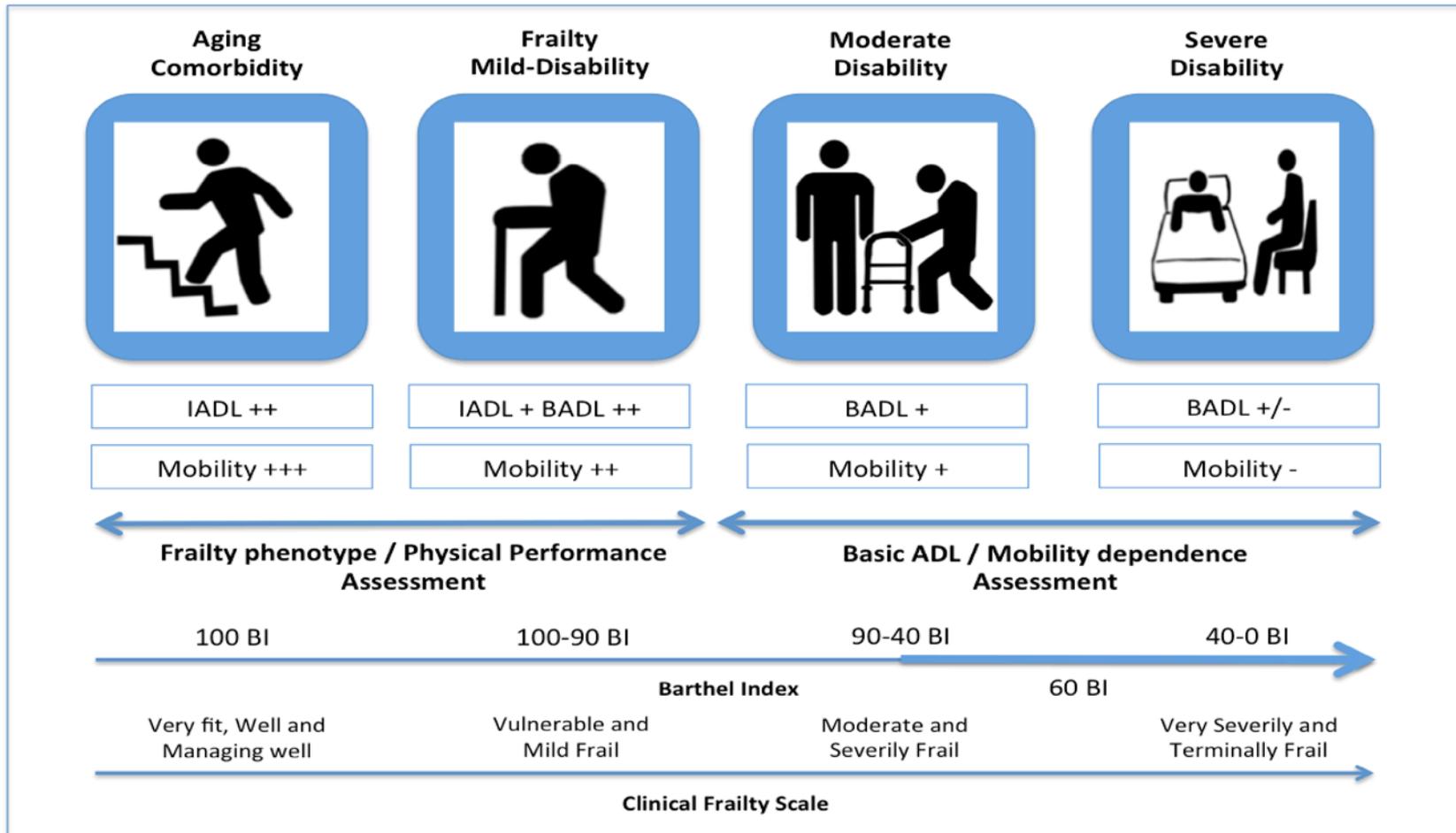
Table S2.-Charlson Cormobidity Index.

| Charlson Comorbidity Index | |
|--|---|
| Cardiovascular system | Neurologic disorders |
| Myocardial infarct • Yes = 1 Congestive heart failure • Yes = 1 Peripheral vascular disease • Yes = 1 | Dementia • Yes = 1 Cerebrovascular disease • Without hemiplegia = 1 • Hemiplegia = 2 |
| Respiratory system | Rheumatologic disorders |
| Chronic pulmonary disease • Yes = 1 | Connective tissue disease • Yes = 1 |
| Kidney and urinary tract | Infectious diseases |
| Moderate or severe renal disease • Yes = 2 | AIDS • Yes = 6 |
| Gastrointestinal system | Oncology and haematology |
| Ulcer disease • Yes = 1 Liver disease • Mild = 1 • Moderate or severe = 3 | Leukemia • Yes = 2 Lymphoma • Yes = 2 Malignant solid tumor • Non-metastatic = 2 • Metastatic = 6 |
| Endocrinology | Total Score: |
| Diabetes • Without end-organ damage = 1 • With end-organ damage = 2 | |

Table S3.-Barthel Index.

| Basic activities of daily living | |
|---|---|
| <p>Bathing (0-5)</p> <p>0 = dependent 5 = independent (or in shower)</p> | <p>Grooming (0-5)</p> <p>0 = needs help with personal care 5 = independent face / hair / teeth / shaving</p> |
| <p>Dressing (0-10)</p> <p>0 = dependent 5 = needs help but can do about half unaided 10 = independent (including buttons, zips,...)</p> | <p>Toilet use (0-10)</p> <p>0 = dependent 5 = needs some help but can do something alone 10 = independent (on and off, dressing, wiping)</p> |
| <p>Transfer from bed to chair (0-15)</p> <p>0 = dependent – no sitting balance 5 = mayor help (physical, one or two people) 10 = minor help (verbal o physical) 15 = independent</p> | <p>Feeding (0-10)</p> <p>0 = dependent 5 = needs help (cutting, spreading butter,...) 10 = independent</p> |
| <p>Continence Bowels (0-10)</p> <p>0 = incontinent 5 = occasional accident 10 = continent</p> | <p>Continence Bladder (0-10)</p> <p>0 = incontinent 5 = occasional accident 10 = continent</p> |
| Mobility | |
| <p>Mobility (0-15)</p> <p>0 = immobile 5 = wheelchair independent 10 = needs help of one person 15 = independent (may use any aid)</p> | <p>Stairs (0-10)</p> <p>0 = unable 5 = needs help 10 = independent to go up and down</p> |
| <p>Total Score: 0- 100</p> | |

Figure 1.-Practical approach to assess the vulnerability in older patients with acute heart failure.

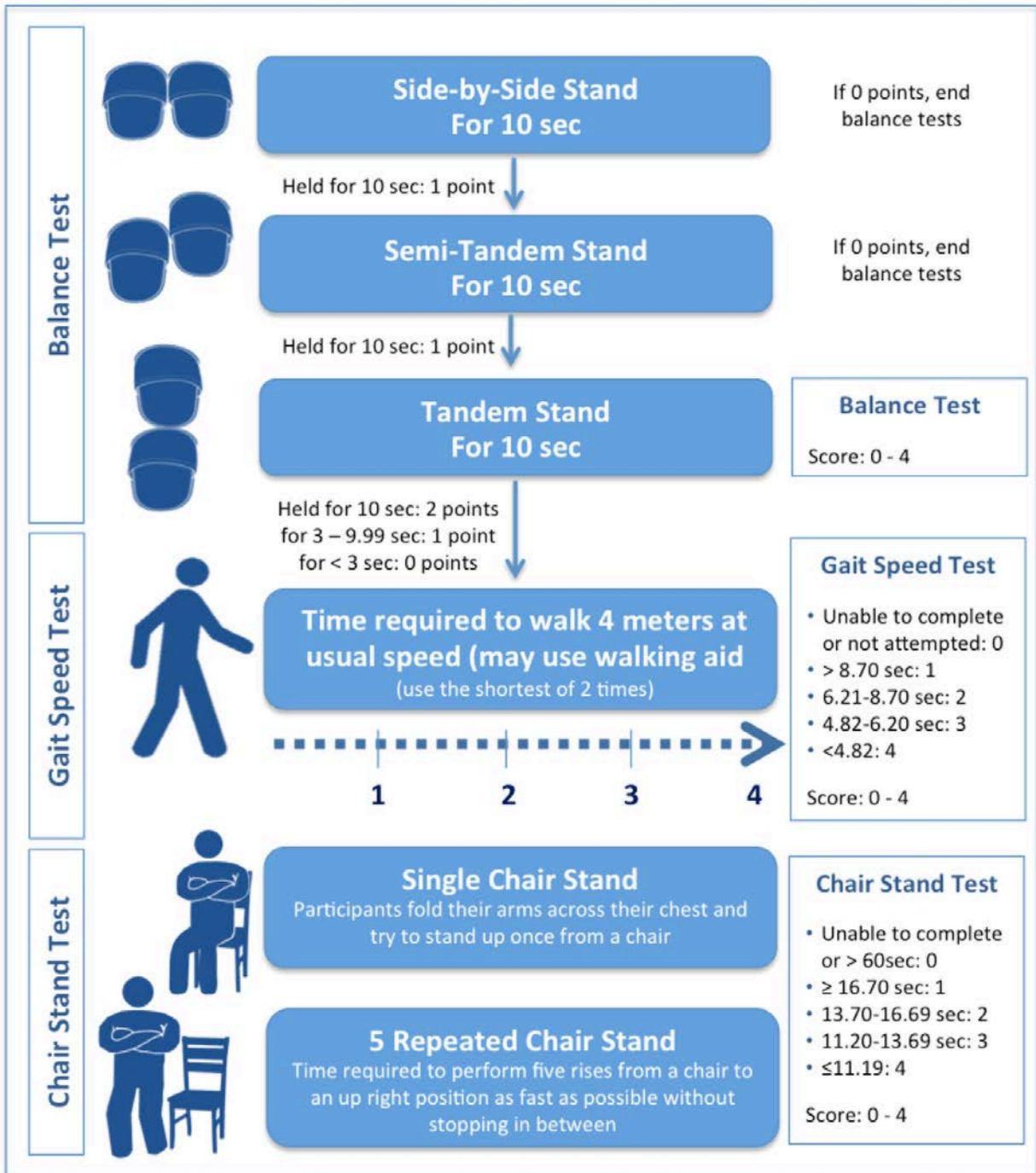


Instrumental activities of daily living (IADL): ability to use telephone, mode of transportation, ability to handle finances, responsibility for own medications, shopping, food preparation, housekeeping, laundry.

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 wheel chair; bedridden.

Figure 2.-Short Physical Performance Battery.



*See the videos in the supplementary electronic material.

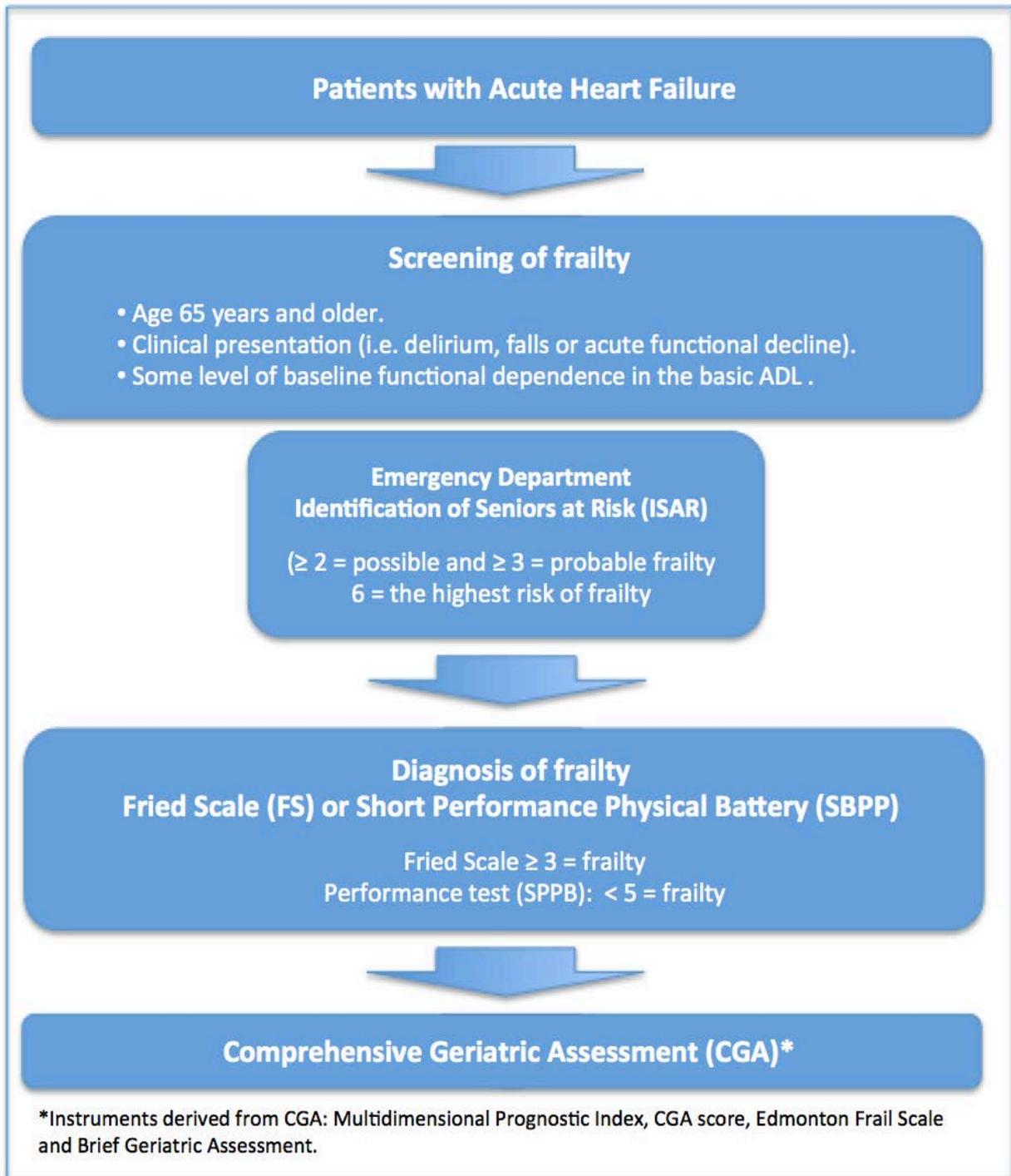
Figure 3.-Approach to assess the frailty in older patients with acute heart failure.

Figure 4.-Management of older patients with acute heart failure based on frailty.