1	Validation of the Mini Nutritional Assessment-Short Form in a population of frail elders
2	without disability. Analysis of the Toulouse Frailty Platform population in 2013.
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30 Abstract

Objective: To assess the validity of the Mini Nutritional Assessment-Short Form (MNA-SF)
 in elderly patients from the Toulouse Frailty Platform.

Participants: Overall, 267 patients aged 65 and over, without severe cognitive impairment (i.e. Mini Mental Status Examination > 20 and CDR<1), no physical disability (i.e. Activities of Daily Living \geq 5) and no active cancer history (over the past 12 months) were included in 2013.

Measurements: Receiver operating characteristic (ROC) analyses were used to assess the predictive validity of the French version of the MNA-SF for good nutritional status (defined as a full MNA score≥24/30). Analyses were conducted in the overall sample and then in subgroups of frail and pre-frail subjects according to the frailty phenotype. Optimal cut-off points were determined to obtain the best sensitivity/specificity ratio and the highest number of correctly classified subjects.

Results: Among 267 patients, mean age=81.5±5.8; women=67.0%; 138 (51.7%) were frail,
98 (36.7%) were pre-frail and 31 (11.6%) were robust. Given their MNA-SF scores, 201
(75.3%) had a good nutritional status, 61 (22.8%) were at risk of malnutrition and 5 (1.9%)
were malnourished. In the overall sample, but also in subgroups of pre-frail or frail elders, the
areas under ROC curves were 0.954, 0.948 and 0.958 respectively. The 11 points cut-off
provided the best correct classification ratio (91.4%); sensitivity=94.0%, specificity=83.3%.

49 **Conclusion:** The MNA-SF appeared to be a validated and effective tool for malnutrition 50 screening in frail elders. Implementing this tool in clinical routine should contribute to 51 improving the screening of malnourished frail individuals.

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53 Key words: Malnutrition; Elderly; Mini Nutritional Assessment; Frailty

Introduction

In western aging societies, protein-energy malnutrition defined as an energy deficit 55 due to chronic deficiency of all macronutrients appears as a major Public Health concern that 56 affects 5 to 15% of community dwelling older subjects (1-3). Since a poor nutritional status is 57 associated with adverse clinical and economic outcomes such as increased mortality or 58 increased costs of hospitalization (4, 5), the need for formal screening procedures has 59 emerged. Promising perspectives of interventional procedures (e.g. dietary protein 60 supplementation) aiming at reversing the burden of malnutrition have also been suggested (6, 61 7). 62

Numerous tools have been validated for malnutrition screening in the elderly (8). 63 64 Among them, the Mini Nutritional Assessment® (MNA) (8) has demonstrated several strengths. For example, in community dwelling elderly, the MNA® can detect risk of 65 malnutrition while albumin and BMI are in the normal range and life style characteristics are 66 associated with nutritional risk (9). In outpatients and hospital patients, the MNA® is 67 predictive of outcome and cost of care (10). However, this test requires substantial time to 68 complete (up to 15 minutes) (8). Therefore, a shorter version of the MNA has been 69 elaborated: the MNA short-form (MNA-SF), which consists of 6 items and takes only 3 70 minutes to be completed, albeit keeping the usefulness and accuracy of the full version (3). 71 Subjects can be classified in 3 categories: normal nutritional status, at risk of malnutrition and 72 malnourished. In addition, the revised version of the MNA-SF proposed by Kaiser and 73 colleagues (11) allows to assess the calf circumference (instantly measured with a tape) when 74 the BMI is unavailable. Thus, this brief tool may represent a first-choice instrument for 75 clinicians looking for a quick and efficient malnutrition screening instrument, designed for 76 elderly patients. 77

Frailty is a state of extreme vulnerability, characterized by insufficient homeostatic 78 reserves to efficiently cope against stressors (2). This condition is also known to be a 79 "dynamic state", suggesting that frail people can transit to non-frail status with ad hoc 80 interventions (12). Despite the absence of consensual definition of frailty, the frailty 81 phenotype proposed by Fried and colleagues is considered as an operational delineation of 82 this condition (2). This tool consists of five criteria: exhaustion, involuntary weight loss, low 83 activity, slow walk and poor grip strength. Older adults meeting these criteria are at higher 84 risk of developing impairment of activities of daily living and show higher morbi-mortality 85 (13-15). In the former subjects, nutrition surely represents a cornerstone to maintain good 86 87 functional performances and prevent poor health outcomes. Screening malnourished (or at risk of malnutrition) frail elders enables to perform a geriatric assessment, review critically 88 their diet and offer them corrective measures and nutritional support (16). Precisely, the 89 90 MNA-SF was designed to assess malnutrition is different populations of vulnerable older adults such as hospitalized patients, nursing home residents or demented subjects (9). 91 However, to our knowledge this instrument has not been validated yet in a frail outpatients 92 population meeting Fried and colleagues' criteria. 93

Therefore, in the present study, we aimed at validating the screening accuracy of the French version of the MNA-SF compared with the full French MNA in outpatients from the geriatric Frailty Platform (structured as a Day Hospital) of Toulouse, France. The cut-point of 12 points and over was shown to be the most appropriate for nutritional screening in a heterogeneous population of hospitalized elders and community dwelling older adults (17). We hypothesized that this very cut-off is correct to screen frail elderly patients for malnutrition.

Methods

102 **Population**

All the outpatients who were admitted to the Toulouse Frailty Platform, France in 103 2013 were eligible for the present analyses. Participants were referred either by their general 104 practitioner, by hospital specialist consultants or by the oncogeriatrics consulting team. The 105 detailed methodology of the Frailty Platform has been published previously (18, 19). Briefly, 106 the main objective of this day hospital is to provide a comprehensive assessment of the 107 medical, functional, cognitive, nutritional and social resources of frail older individuals. 108 Although these patients usually meet frailty criteria, they do not present disability in activities 109 of daily living (ADL) or major cognitive impairment. Accordingly, personalized interventions 110 may be provided by a dietitian (e.g. nutritional counseling), a physical trainer (e.g. training 111 program for muscle reinforcement) or a neuropsychologist (e.g. thorough cognitive 112 113 evaluation).

Before July 1st, 2013 the MNA-SF scores of the patients were not recorded in our database. However, only data with both the MNA-SF and the full MNA available were considered in the present analyses. The inclusion criteria were: $age \ge 65$ years, no active cancer history over the past 12 months (since the fraily phenotype has been questionned in cancer patients) and an ADL score ≥ 5 (i.e. no physical disability). As the frailty phenotype has never been validated in demented subjects, subjects with an MMSE score <21 or a Clinical Dementia Rating score ≥ 1 have also been excluded.

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122 Variables

In the present study, the French version of the MNA-SF and the full MNA were used (20). Sociodemographic characteristics were obtained through questionnaire. A medical interview performed by a geriatrician provided the following: medical and surgical history, ongoing medication and physical examination. Height and weight were measured and Body
Mass Index (BMI) was calculated as the weight in kg divided by the square of the height in
meters. Daily self-care activities were assessed with the ADL (21) and the Instrumental ADL
(IADL) (22) scales.

Frailty assessment was performed by specialized nurses and was based on Fried's 130 frailty phenotype (2) i.e. involuntary weight loss, self-reported exhaustion, muscle weakness, 131 slow gait speed, and low physical activity. A 5 kg-weight loss over the past year (either 132 measured or reported by the patient) was considered to be significant. The exhaustion 133 criterion was met if the answer was "Much or most of the time" when asked, "How often in 134 the last week did you feel this way" to either of the following two statements: "I felt that 135 everything I did was an effort" and "I could not get going." The muscle weakness criterion 136 was met when the average of 3 handgrip strength measurements by a handheld dynamometer, 137 was less than or equal to the sex- and BMI specific cutoff points provided by Fried and 138 colleagues. Slow usual gait speed (assessed over a 4-meter distance, starting from a still 139 position) was defined either as a time of more than 6 seconds for men whose height is less 140 than or equal to 173 cm (or women ≤ 159 cm respectively); or as a time of more than 5 141 seconds for men whose height is more than 173 cm (or women > 159 cm respectively). We 142 defined as "sedentary" those participants who had performed no physical activity, spent most 143 of the time sitting, or rarely had a short walk (or other non-demanding physical activity) in the 144 past year according to a validated interviewer-administered questionnaire (23). The Fried 145 score was defined with these five items; patients with a 0 score were considered as robust, 146 those with a 1 or 2 score were considered as pre-frail, and those with a 3 to 5 score were frail. 147 The physical evaluation was completed with the Short Physical Performance Battery (24). 148

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150 Statistical analyses

Receiver operating characteristic (ROC) analyses were used to assess the predictive validity of the MNA-SF. The reference was a good nutritional status (defined as a score above 23.5/30 according to the full MNA). Analyses were conducted in the overall sample and after categorization of our participants in frail and pre-frail groups. Optimal cut-off points were determined to obtain the best sensitivity/specificity ratio and the highest number of correctly classified subjects. Analyses were performed using STATA v11.0 (Stata Corp., College Station, TX). 8 Results

Data of 412 subjects from the Toulouse Frailty Clinic were recorded in our 2013 database. Fifty nine of them were excluded because of their low cognitive performances (MMSE<21), 57 were because they had insufficient Activities of Daily Living performances (score of less than 5) and one because he was under 65 years old. Two MNA scores had missing items. Twenty six had active cancer history. Thus, 145 subjects were excluded, and our final sample comprised 267 individuals. The flow-chart for participants' selection is presented in Figure 1.

The main nutritional and physical characteristics of the study sample are presented in Table 1. Mean age was 81.5 (Standard deviation (SD) 5.8) years old; 67.0% were women; 57.9% had a normal ADL score (6/6) and 63.9% a 7 to 8 IADL score. According to the Fried frailty criteria, 11.3% of the participants were robust, 51.9% were pre-frail and 36.8% were frail. All of our malnourished participants were frail.

In the overall sample (Figure 1) as well as in pre-frail and frail considered separately 171 (Figure 2), the areas under ROC curves were 0.954 95% Confidence Interval (CI) 0.928-172 0.980, 0.948 95% CI 0.908-0.987 and 0.956 95% CI 0.906-0.996 respectively. The 11 points 173 cut-off allowed the best correct classification ratio (91.4%), with a sensitivity of 94.0% and a 174 specificity of 83.3%. With a 12 points cut-off the sensitivity was: 76.1, and the specificity: 175 95.5% (Table 2). After stratification on pre-frail and frail status, there was no significant 176 difference of areas under curve between the former and the latter individuals (0.95 vs. 0.96, p 177 = 0.78). 178

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IV. Discussion

180 In this study, the MNA-SF appeared to be an accurate tool for malnutrition screening in patients meeting frailty criteria. The best cut-point for good nutritional status was a score of 181 11 and above, allowing a sensitivity of 94.0% and specificity of 83.3%, hence a Youden's 182 index of 0.77. Given this threshold, 91.4% of our subjects were correctly classified. Pre-frail 183 and frail older adults share common characteristics and are significantly more affected by 184 nutritional issues than robust elders (25). In our sample, there was no significant difference of 185 correctly classified subjects using the MNA-SF between pre-frail and frail elders. Therefore, 186 this study confirms that the MNA-SF compares well with the full MNA and represents a valid 187 188 instrument for nutritional screening in a frail out-patients population.

Our findings were consistent with previous validation studies of the MNA-SF (3, 26). 189 However, the usual MNA-SF threshold for good nutritional status is a 12 out of 14 score. As a 190 191 result, subjects with a score below 12 need to complete the full MNA. Rubenstein and colleagues (3) had already acknowledged that the 11 cut-point provided a better 192 193 sensitivity/specificity ratio to indicate undernutrition. Yet, raising the threshold to 12 reduced the number of persons incorrectly identified as well-nourished despite a higher number of 194 false negative (i.e. people without malnutrition who will be referred to the dietician). Frail 195 elders represent primary targets for nutritional screening. Consistent with the foregoing study, 196 we assumed that failing to screen a malnourished elder would be of greater concern than 197 requiring additional evaluation for people mistakenly identified as malnourished. Thus, the 198 previously established 12 threshold appears as appropriate in our population of frail older 199 adults. 200

201 Malnutrition and frailty are two interrelated syndromes. Firstly, malnutrition is very 202 common among the elderly frail populations (25). Secondly, this condition may directly 203 impact the Fried's Frailty phenotype. Weight loss reflects both conditions. An imbalance

between energy intake and expenditure leads to muscle weakness that may in turn impact the 204 205 four remaining criteria: poor muscle strength, slowness, exhaustion and reduced functional activities (27). Diet quality and frailty have also been largely studied. Overall diet quality is 206 lower in frail than in robust older adults (28). High protein diet reduces the risk of incident 207 frailty (29). Conversely, low serum micronutrient concentrations were shown to be an 208 independent risk factor for frailty in community-dwelling women (30). Frailty also increases 209 muscle protein catabolism and enhances age-related loss of muscle mass resulting in 210 sarcopenia and impaired mobility (31). Of note, all of our malnourished participants (N=5) 211 were frail according to the frailty phenotype. 212

The identification of a pre-disability state (i.e., frailty) enables to detect older persons at risk of adverse health events who may still benefit from preventive interventions against disability. Our population was specifically recruited by general practitioners or specialists to meet these frailty criteria. The good accuracy of the MNA-SF in this population encourages expanding its use in these subjects so as to improve the screening of malnutrition.

This study also had limitations. Our sample was smaller than many studies focused on 218 the effectiveness of the MNA-SF (26, 32-34). We also excluded a substantial number of 219 participants (246) due to missing data. However, we achieved comparable results to the main 220 validation studies of this tool. We did not compare the results of the screening test with a 221 dietitian assessment but with another test (i.e. the full MNA) which was used as a surrogate 222 for the diagnosis of malnutrition. Nevertheless, the MNA was demonstrated to be both a 223 screening and assessment tool with a good internal consistency and inter-observer reliability 224 and validity (8, 9). The aim of the MNA-SF is definitely not to replace the full MNA, but to 225 refine the selection of subjects who should be tested with the full version. 226

V. Conclusion

The MNA-SF has already been suggested as an accurate screening tool in various 228 populations of elderly subjects. This study confirmed its usefulness among frail (and pre-frail) 229 older adults, with similar cut-points to indicate good nutritional status than in previous 230 studies. All the subjects with a MNA-SF score below 12 should undergo a full MNA to 231 establish whether they present under-nutrition or not. Malnourished frail elders are priority 232 targets for comprehensive assessment and multidimensional management, and in particular 233 for nutritional interventions. Therefore, The MNA-SF allows a quick and appropriate 234 screening of frail older adults and may indeed be advantageously part of the clinical routine of 235 general practitioners as well as hospital specialists. 236

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