

Review

Screening Tools for Sarcopenia

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Abstract. *The elderly people are characterized by multiple comorbidities, dementia, and are at risk of developing sarcopenia and frailty. Sarcopenia is defined by loss of muscle mass and muscle strength or physical decline. Sarcopenia is a main component of physical frailty. Screening tools for sarcopenia that can be easily determined in daily practice are useful and include the SARC-F screening tool. SARC-F is a questionnaire consisting of five questions: Strength (S), Assistance walking (A), Rising from a chair (R), Climbing stairs (C), and Falls (F) on a scale of 0 to 2. The recommended cutoff value is ≥ 4 points. The SARC-F has been shown to correlate well with clinical outcomes in the elderly and various underlying diseases, while it is also true that the SARC-F has its shortcomings such as low sensitivity for sarcopenia. In this review, we mainly outline the SARC-F and mention other screening tools for sarcopenia.*

The aging of Japan's population continues to accelerate (1). According to the Japanese government report, the aging rate (65 years old or older) was 15.1% of the total population in 1996, but it increased significantly to 28.4% in 2019 (2). In addition, the estimated aging rate as of January 1997 is projected to be 27.4% in 2025, but as of 2019, it has already

exceeded the projection. The number of people aged 65 or older will continue to increase, and is estimated to reach 33.3% in 2036 and 38.4% in 2065. In order to improve the quality of life (QOL) of the increasing number of elderly people, efforts have been made in the fields of health, medical care, and welfare to extend healthy life expectancy (the period of time during which there are no limitations in daily life), but there are still many issues to be addressed. The elderly people are characterized by multiple comorbidities, dementia, and risk of sarcopenia and frailty (3). In 2019, the Ministry of Health, Labour and Welfare (MHLW) in Japan published the "Guidelines for Health Services Based on the Characteristics of the Elderly, Version 2", and announced the strengthening of measures against frailty, especially for elderly people aged 75 or older. The guidelines newly added the implementation of "health checkups for frailty" with the aim of integrating health guidance and care prevention. Specifically, the report has recommended the use of a "questionnaire for the elderly aged 75 or older" focusing on frailty in health checkups and other examinations conducted by municipalities since fiscal 2020. This questionnaire is expected to help prevent and raise awareness of frailty among many elderly people. Early intervention through such efforts will lead to a prolongation in healthy life expectancy. In addition, sarcopenia is a condition that forms the basis of physical frailty (4). Sarcopenia leads to decreased physical activity, and decreased physical activity leads to sarcopenia. Both are involved in frailty, forming the so-called "frailty cycle" (4). On the other hand, the number of people aged 60 or older in the world was estimated to be 600 million in 2000, and is projected to increase to 1.2 billion by 2025 and to 2 billion by 2050. Even if the conservative estimation of the prevalence of sarcopenia, the sarcopenic population is expected to exceed 200 million in the next 40 years (5).

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Therefore, the “enclosure” of patients at risk for sarcopenia is clinically important, and a screening tool that can be easily used in daily practice is useful.

Changes in the Definition of Sarcopenia and the SARC-F as a Screening Tool

More than 30 years have passed since Rosenberg proposed that aging-related skeletal muscle mass loss be called sarcopenia, from the Greek words “sarco” meaning muscle and “penia” meaning loss (6). Later, Baumgartner used the skeletal muscle mass index (SMI), which is the appendicular skeletal muscle mass of the extremities divided by the square of the height (m), as in the calculation of body mass index (BMI), and defined sarcopenia as a SMI lower than two standard deviations from the mean value (7). However, the significance of defining sarcopenia solely in terms of loss of skeletal muscle mass was reviewed, because it was reported that the loss of muscle strength was more significant than the loss of skeletal muscle mass in the elderly, and that dynapenia, which is an aging-related loss of muscle strength, was more reflective of functional disability and mortality (8-10). In 2010, the consensus on sarcopenia by the European Working Group on Sarcopenia in Older People (EWGSOP) has developed the concept of sarcopenia to include not only a decrease in skeletal muscle mass but also a decrease in muscle strength and physical performance such as grip strength and walking speed (11). Later, the International Working Group on Sarcopenia (IWGS) in 2011 (12) and the Asian working group for sarcopenia (AWGS) in 2014 (13) provided definitions of sarcopenia. In 2016, the Japanese Society of Hepatology (JSH) published criteria for secondary sarcopenia specific to liver disease (14). This criterion has attracted attention as the world’s only criterion for secondary sarcopenia, and is characterized by the omission of age restriction from the viewpoint of disease-related sarcopenia, the omission of gait speed, which is complicated to measure in daily practice, and the reference values of skeletal muscle mass on computed tomography (14). Subsequently, in Japan, the Japanese Association on Sarcopenia and Frailty, the Japan Geriatrics Society, and the National Center for Geriatrics and Gerontology took the lead in creating clinical practice guidelines for sarcopenia in 2017 (15). Recently, the definition of sarcopenia has been updated by EWGSOP second edition (EWGSOP2) (16) and AWGS second edition (AWGS2) (3), and practical guidelines for screening, diagnosis, and management of sarcopenia have been published. EWGSOP2 involves 1) detecting low muscle strength and low skeletal muscle mass and quality to confirm sarcopenia as these are important characteristics of sarcopenia, and identifying sarcopenia severity by physical function decline (gait speed, *etc.*) and 2) providing clear cutoff values that identify sarcopenia (16). However,

AWGS2 recommends early intervention of nutritional and exercise therapy for people at risk of sarcopenia, using only grip strength or chair stand test to assess sarcopenia as an alternative when it is difficult to measure muscle mass (3). In AWGS2, the threshold for grip strength for males has been raised by 2 kg to 28 kg. The threshold for grip strength for women in AWGS2 remains at 18 kg. In addition, the standard value for walking speed has been changed from less than 0.8 m/s to less than 1 m/s (3).

EWGSOP2 also proposes a new algorithm for the detection of sarcopenia. This algorithm uses a pathway for defining sarcopenia using Find-Assess-Confirm-Severity (F-A-C-S), instead of the conventional method of determining sarcopenia by the loss of skeletal muscle mass in addition to decreased walking speed or muscle strength decline. First, muscle strength (grip strength) is assessed if screening is positive or there is clinical suspicion; and if there is muscle weakness, sarcopenia is suspected and targeted for intervention. Next, skeletal muscle mass is assessed, and if a decrease is observed, sarcopenia is confirmed. In addition to a decrease in grip strength and skeletal muscle mass, a decrease in physical performance (gait speed, *etc.*) is classified as severe sarcopenia (16). AWGS2 also recommends the assessment of sarcopenia using a similar technique (3). The SARC-F is used for the screening of sarcopenia.

The selection of subjects for sarcopenia, either for clinical or research purposes, is extremely important. In the following, the SARC-F is mainly outlined, and other screening tools for sarcopenia are also mentioned.

The SARC-F and International Versatility

The SARC-F was presented by Morley as a screening tool for sarcopenia at the EU/US Committee on Sarcopenia in the Elderly at the International Conference on Sarcopenia Research (ICSR) held in Orlando in 2012 (17, 18). The SARC-F is a questionnaire consisting of five questions: Strength (S), Assistance walking (A), Rising from a chair (R), Climbing stairs (C), and Falls (F) on a scale of 0 to 2 from “not at all” to “very difficult”, and the total score (out of 10) is calculated (17). The recommended cutoff value is ≥ 4 points (17).

Both EWGSOP2 and AWGS2 recommend the use of the SARC-F for the screening of sarcopenia (3, 16). Similarly, the Society for Sarcopenia, Cachexia and Wasting Disorders also recommends the use of the SARC-F (19). On the other hand, during difficult times such as the COVID-19 pandemic, the health care system needs to adapt quickly and efficiently. It is essential that all patients have access to nutritional care as part of their primary health care services, even if social distance measures are taken because of the COVID-19 pandemic. Therefore, for example, in Croatia, a simple nutritional screening tool and practical guidance for nutritional care in

primary practice were proposed and a tele-medicine process suitable for health care providers has been introduced. This protocol includes the Malnutrition Universal Screening Tool (“MUST”), SARC-F and practical guidance on nutritional interventions for family physicians to identify nutritional risk and loss of muscle mass and function (20). The clinical importance of SARC-F has been acknowledged worldwide.

Sarcopenia Detection Power of the SARC-F, the SARC-calf and the SARC-EBM

The SARC-F. Several reports have examined the power of the SARC-F to detect sarcopenia as defined by the EWGSOP and AWGS criteria, using receiver operating characteristic (ROC) curves. Woo *et al.* (21) reported in Asians that ROC analysis for sarcopenia defined by EWGSOP, IWGS, and AWGS had high specificity (0.94-0.99) but low sensitivity (0.038-0.048). The data of Kim *et al.* (22) in Koreans showed similar results in terms of sensitivity and specificity, and the area under the ROC curves (AUCs) for sarcopenia by AWGS criteria were not so high, 0.618 for men and 0.695 for women. In a comparative study by Yang *et al.* between the SARC-F and other sarcopenia screening tools, the sensitivity and specificity were similar when using the recommended cutoff value of the SARC-F (4 points), but the AUC was considerably higher, 0.86 for men and 0.90 for women (23). However, Ida *et al.* reported data from Japanese patients with diabetes mellitus (24). Although the specificity was rather low (0.858 in men and 0.724 in women), the sensitivity was somewhat higher (0.146 in men and 0.333 in women) than that of other reports (24). Ida *et al.* also conducted a meta-analysis and reported a specificity of 0.90 and a sensitivity of 0.21 when data from 12,800 subjects in seven articles were combined (25). Therefore, a positive result on the SARC-F indicates a high rate of sarcopenia, while the sensitivity to the presence of sarcopenia is clearly low. For this reason, it is recommended to check muscle strength when there is clinical suspicion for sarcopenia in EWGSOP2 algorithm, regardless of whether SARC-F is positive or negative (16).

The SARC-calf. AWGS2 recommends screening with either calf circumference (CC), which is known to correlate well with limb skeletal muscle mass and BMI (cut-off value: <34 cm for men and <33 cm for women (26), the SARC-F questionnaire, or the SARC-calf, which combines the two, as a trigger for case finding (3). Recent studies have shown a strong correlation between CC and appendicular skeletal muscle mass [correlation coefficient (r): r=0.78 (men) and r=0.76 (women)] (27). Using the cut-off value for CC recommended by AWGS2, the AUC for sarcopenia was 0.819, with a good sensitivity/specificity of 0.804/0.718 (28). The cutoff value for SARC-calf is 11 or higher (max, 20

points). If CC value is less than the cutoff value, 10 points are added to the SARC-F score to evaluate. CC has been reported to have the highest detection of sarcopenia among arm circumference, arm muscle circumference, CC, arm muscle area, triceps skinfold thickness, waist circumference, and BMI (29). The SARC-calf has the role of compensating for the low sensitivity, which is a problem pointed out when the SARC-F is used alone. In an observational study of 384 patients aged 60 years or older, the sensitivity, specificity, and AUC for sarcopenia as determined by AWGS were 0.295, 0.981, and 0.89 for the SARC-F, compared with 0.607, 0.947, and 0.92 for the SARC-calf (30). In 309 patients with advanced cancer, the sensitivity, specificity, and AUC for sarcopenia as determined by AWGS were 0.321, 0.907, and 0.70 for the SARC-F vs. 0.666, 0.701, and 0.75 for the SARC-calf, respectively (31). In a recent meta-analysis, the sensitivity of SARC-calf for sarcopenia was reported to be 0.459-0.572 and specificity 0.877-0.913, while the sensitivity of SARC-F for sarcopenia was reported to be 0.289-0.553 and specificity 0.689-0.889 (32).

The SARC-EBM. Kurita *et al.* devised the “SARC-F+EBM” method, which adds “EBM” (Elderly and BMI) to the SARC-F and investigated the degree of improvement in the diagnostic performance of sarcopenia. In addition to the score from the SARC-F questionnaire, this method also takes into account whether the patient is 75 years or older and whether the BMI is less than 21 kg/m² (33). Elderly people requiring care with a low BMI show a decline in ADL, swallowing function, and cognitive function, and often consume foods that do not require chewing ability (34). When validated in 959 patients with musculoskeletal diseases, the sensitivity, specificity, and AUC of the SARC-F+EBM diagnostic method were 0.778, 0.696, and 0.824, respectively, for the diagnosis of sarcopenia. They reported that the sensitivity, specificity, and AUC of the SARC-F questionnaire alone were 0.417, 0.685, and 0.557, respectively, for the diagnosis of sarcopenia, and that the SARC-F+EBM diagnostic method was clearly superior in sensitivity (33).

Validity of the SARC-F. By correlating the SARC-F with other sarcopenia screening tools and health-related indicators, the validity of the SARC-F can be examined. For example, there is a significant correlation between the SARC-F and the Short portable sarcopenia measure (SPSM, see below), which is one of the sarcopenia screening tools (35). In order to confirm the validity of the Japanese version of the SARC-F, Ida *et al.* examined the correlations with the nursing care prevention checklist, instrumental activity of daily living, intellectual activity, social roles, and self-efficacy for falling, all of which showed significant correlations (24). All the results show the validity of the SARC-F. It is also attractive that the SARC-F requires only

a questionnaire and does not require measurement of CC as the SARC-calf does. The low sensitivity of the SARC-F for sarcopenia is problematic because the diagnosis is made by first narrowing down the candidates with a highly sensitive test and then confirming the diagnosis with a highly specific test. However, in light of the above reports, the SARC-F can be a useful tool for sarcopenia in daily practice.

With regard to the low sensitivity of the SARC-F to sarcopenia, this is because the questions are only related to muscle strength and physical performance, and do not include items that reflect skeletal muscle mass, such as weight and BMI. Kim *et al.* reported that there was no difference in SMI between the SARC-F \geq 4 group and the SARC-F<4 group in women (22). Although the SARC-F is a simple assessment tool, it implies that it is not sufficient for sarcopenia screening. However, the SARC-F is closely related to the future decline in physical functioning, thus the SARC-F has the same significance as the direct assessment of sarcopenia (36). In terms of predictive ability for future decline in physical function, the SARC-F and sarcopenia assessed by EWGSOP, AWGS, and IWGS criteria were almost equal regarding predictive ability (36). On the other hand, the correlation between survival and the SARC-F has also been reported. A meta-analysis of five observational studies reported that the pooled hazard ratio for the SARC-F questionnaire positivity and mortality was 1.87 ($p<0.0001$) (37). These findings suggest that the SARC-F is very important in daily practice.

The SARC-F, Physical Function and Frailty

There is a study called the ZEVIOUS study that examines the validity of classifying the level of nursing care in home health care patients based on subjective measures of daily function and sarcopenia. In this study, the relationship between the level of nursing care and self-reported daily living and physical functions in 181 patients receiving home health care was cross-sectionally analyzed (38). In the results, the level of nursing care varied, ranging from support level 1 (5.5%) to care-needs level 5 (10.5%), with care-needs level 2 being the most prevalent (24.9%). Lifestyle function was assessed with the World Health Organization Disability Assessment Schedule (WHODAS2.0), which reflects the International Classification of Functioning, Disability and Health (ICF), and physical function was assessed with the SARC-F. The higher the level of nursing care, the lower the level of self-reported life and physical functioning (38). Therefore, the level of nursing care and the SARC-F are significantly positively correlated. The SARC-F is also useful for enclosing frailty patients (39). In a study of 447 elderly patients by Bahat *et al.*, the sensitivity and specificity of SARC-F (cutoff \geq 1 point) for diagnosing frailty were 0.914 and 0.449, respectively. In the case of SARC-F (cutoff

\geq 2 points), the best balance of sensitivity and specificity was achieved in identifying frailty (sensitivity=0.741 vs. specificity=0.737, AUC=0.807) (39). In the case of SARC-F (cutoff \geq 4 points), the sensitivity and specificity were 0.462 and 0.926, respectively (39).

Clinical Significance of the SARC-F According to Diseases

Several reports have examined the significance of SARC-F in Parkinson's disease. In Parkinson's disease patients, the SARC-F has been reported to be a prognostic factor for disability (40) and falls (41). In a study of 60 Parkinson's disease patients, the sensitivity of SARC-F for sarcopenia using EWGSOP2 criteria was as low as 0.231, and the SARC-calf was more useful than the SARC-F (42). In 198 diabetic patients, the SARC-F has been reported to be useful in predicting hospitalization and development of disability (43). In diabetic patients, the SARC-F is also useful in predicting mild cognitive impairment (44), depression (45), overactive bladder (46), and sleep disorders (47). In a study of 127 patients undergoing peritoneal dialysis, the sensitivity and specificity of SARC-F for sarcopenia assessed by AWGS2 were 0.700 and 0.812, respectively (48). In a study of 717 patients with chronic liver diseases (CLDs), the sensitivity and specificity of SARC-F for sarcopenia assessed by the JSH were 0.65 and 0.68 when the cutoff value of SARC-F was set at 1, and the predictive ability was better than when the cutoff value of SARC-F was set at 4 (49). In a study of 94 patients with systemic sclerosis, the sensitivity and specificity for sarcopenia assessed by EWGSOP2 were 0.400 and 0.810 for the SARC-F, and 0.533 and 0.848 for the SARC-calf, respectively (50). In a study of 355 patients with chronic heart failure, the sensitivity and specificity of SARC-F for sarcopenia assessed by AWGS were 0.525 and 0.962, respectively, and in terms of predictive ability on Mini Sarcopenia Risk Assessment (MSRA, described later) for sarcopenia, the MSRA-7 had a sensitivity of 0.924 and a specificity of 0.516, and the MSRA-5 had a sensitivity of 0.939 and a specificity of 0.573. The AUCs for the SARC-F, MSRA-7, and MSRA-5 were 0.78, 0.74 and 0.77, and MSRA-5 and MSRA-7 were reported to be sensitive screening methods (51). The usefulness of SARC-F in diagnosing sarcopenia in cardiovascular diseases has also been reported (52, 53). In 596 cases of non-valvular atrial fibrillation, the SARC-F has been reported to be useful in predicting mortality (53). A study in 108 gastrointestinal cancers reported a low concordance between the SARC-F and SMI (kappa coefficient=0.2) (54). In a study of 77 cancer patients, a significant correlation between the SARC-F and the Charlson comorbidity index, a prognostic score, was noted (55). The cancer-related cachexia prediction model (cancer cachexia score, CSS) in advanced cancer incorporates the SARC-F, which consists of five items: weight loss, SARC-F, PS, appetite loss, and abnormal biochemistry

Table I. Screening tools for sarcopenia.

Modality	Assessment method	Notes
SARC-F	Five queries (0-2 points in each query)	Cutoff point: ≥ 4 points (max, 10 points)
CC	Measurement of CC	Cutoff point: 34 cm (men) and 33 cm (women)
SARC-calf	SARC-F and CC (CC: 10 or 0 points)	Cutoff point: ≥ 11 points (max, 20 points)
SARC-EBM	SARC-F, age, and BMI	Age: 75 years or older, BMI: < 21 kg/m ²
MSRA-7	Age, hospitalization history, activity level, three dietary items, and weight	Range=0-40, cutoff value: 30
MSRA-5	Age, hospitalization history, activity level, one dietary item, and weight	Range=0-60, cutoff value: 45
SPSM	BMI, grip strength, and timed chair stand	International versatility is low.
Ishii screening tool	Age, grip strength, and CC	Range: 0-145 (men)/0-150 (women), cutoff value: 105 (men)/120 (women)
Finger-circle test	A circle with the index finger and thumb around the thickest part of the calf	Three groups: "bigger", "just-fits", and "smaller"

CC: Calf circumference; BMI: body mass index; MSRA: mini sarcopenia risk assessment; SPSM: short portable sarcopenia measure.

(56). In a study of 187 patients with advanced cancer, the SARC-F was shown to be strongly correlated with cancer-related fatigue (57). In a study of 256 cancer patients aged 60 years or older, the SARC-F correlated well with impaired instrumental activities of daily living and frailty as well as reduced physical and mental health-related QOL (58). Recently, the relationship between the SARC-F and progression of COVID-19 infection has been reported. In a study of 114 patients with COVID-19 infection, COVID-19 patients with higher risk of sarcopenia as assessed by the SARC-F were demonstrated to be more likely to develop severe disease than those without (68% vs. 22%, $p < 0.001$) (59). Of the 407 patients hospitalized with COVID-19, 22% had rapid weight loss during hospitalization and 73% had a SARC-F score of 4 or higher (60). In 152 patients with acute cerebral infarction, the SARC-F was reported to be useful in predicting functional outcome 3 months after stroke onset (61). The SARC-F is also closely correlated with lower limb muscle weakness in acute stroke patients (62). In our study of 1,282 patients with gastrointestinal diseases using the SARC-F, the percentage of patients with SARC-F score of 4 or higher was 17.5% (57/326) in upper gastrointestinal diseases, 12.0% (43/357) in lower gastrointestinal diseases, 17.3% (72/416) in biliary and pancreatic diseases, and 13.7% (25/183) in liver diseases, and there was no significant difference among the four groups ($p = 0.1235$) (63). In our multivariate analysis of SARC-F 4 or higher, age, gender, albumin level, lymphocyte count, C reactive protein, and presence of advanced cancer were independent factors (63).

Other Screening Tools for Sarcopenia

MSRA. The MSRA is available in two versions: the MSRA-7 (range=0-40), which consists of seven items: age,

hospitalization history, activity level, three dietary items, and weight, and the MSRA-5 (range=0-60), which includes age, hospitalization history, activity level, weight, and one dietary item (64). In a study of 274 elderly Italian patients, a sensitivity of 0.804 and a specificity of 0.505 for sarcopenia with a cut-off value of 30 in MSRA-7, while a sensitivity of 0.804 and a specificity of 0.505 for sarcopenia with a cut-off value of 45 in MSRA-5, have been reported (64). In a study of 384 elderly people in China, MSRA-7 had a sensitivity of 0.869 and a specificity of 0.396, whereas the MSRA-5 had a sensitivity of 0.902 and a specificity of 0.706 (23).

SPSM. SPSM is a prediction equation that combines BMI, grip strength, and timed chair stand; it has been used in 998 individuals aged 49-65 years, and robustness has been confirmed by sensitivity analysis (65), but its international versatility is low.

Ishii screening tool. The Ishii screening tool is a method to determine the total score of age, grip strength, and CC [range=0-145 (men)/0-150 (women), cutoff value: 105 (men)/120 (women)] (66). It was developed based on the data of 1,971 elderly Japanese patients. Ishii screening tool has been reported to have high sensitivity and specificity (0.849 and 0.882 for men, 0.755 and 0.920 for women) for the diagnosis of sarcopenia in the elderly (66). In a study of 94 patients with systemic sclerosis, the sensitivity, specificity, and AUC of the Ishii screening tool for sarcopenia were 0.867, 0.734, and 0.862, respectively (50).

Finger-circle test. The finger-circle test is a method of making a circle with the index finger and thumb around the thickest part of the calf and dividing it into three groups: "cannot enclose (bigger)", "just-fits", and "a smaller gap

(smaller)” (67). Of 1,904 participants (mean age=72.8 years), 53% were classified as “bigger”, 33% were in “just fits” and 14% were in “smaller”. Relative to “bigger”, the results statistically associated with sarcopenia [“just fits”; odds ratio (OR)=2.4, “smaller”; OR=6.6]. The results also increased the risk of sarcopenia incidence (“just fits”; OR=2.1, “smaller”; OR=3.4). Furthermore, the “smaller” had 2.0- and 3.2-fold elevated risks of requiring long-term care insurance services and mortality, respectively (67). The usefulness of the finger-circle test has also been shown in patients with CLDs (68, 69). In our study of 202 patients with CLDs (56 liver cirrhosis cases), the rates of SMI reduction in “bigger”, “just-fits”, and “smaller” were 9.9% (14/142), 45.5% (15/33), and 77.8% (21/27; overall $p < 0.01$). The percentages of sarcopenia in “bigger”, “just-fits”, and “smaller” were 3.5% (5/142), 18.2% (6/33), and 33.3% (9/27; overall $p < 0.01$) (68). Screening tools for sarcopenia are listed in Table I.

Final Remarks

In this review article, screening tools for sarcopenia, with a focus on the SARC-F, were outlined. Although the SARC-F has the limitation of low sensitivity, it is an excellent assessment tool because it has high specificity for sarcopenia, and correlates well with other screening tools, prognosis and the level of nursing care. The SARC-F also plays a role in predicting prognosis in the COVID-19 pandemic. However, assessment tools that combine SARC-F with other factors (SARC-calf, SARC-F+EBM, etc.), Ishii screening test, finger-circle test, etc. are also useful. There are many reports on the usefulness of SARC-F not only in the elderly but also in patients with underlying diseases. Regardless of which screening tool is used, it is important to assess muscle mass and strength and physical function in patients suspicious of having sarcopenia. As the number of sarcopenia patients is expected to increase with the aging population, it is hoped that a simpler and more accurate evaluation tool will become available.

Conflicts of Interest

The Authors have no conflicts of interest to declare in relation to this study.

Authors' Contributions

Writing the article: H.N.; Editing and reviewing the article: A.A., S.F., M.G., T.O., T.T., T.M., K.K., S.Na., S.Ni., K.H.; Final approval: all Authors.

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