Is total lymphocyte count related to nutritional markers in hospitalized older adults?

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ABSTRACT - Background - Older patients are commonly malnourished during hospital stay, and a high prevalence of malnutrition is found in hospitalized patients aged more than 65 years. Objective – To investigate whether total lymphocyte count is related to other nutritional markers in hospitalized older adults. Methods - Hospitalized older adults (N=131) were recruited for a cross-sectional study. Their nutritional status was assessed by the Nutritional Risk Screening (NRS), anthropometry, and total lymphocyte count. The statistical analyses included the chi-square test, Fisher's exact test, and Mann-Whitney test. Spearman's linear correlation coefficient determined whether total lymphocyte count was correlated with the nutritional markers. Multiple linear regression determined the parameters associated with lymphocyte count. The significance level was set at 5%. Results - According to the NRS, 41.2% of the patients were at nutritional risk, and 36% had mild or moderate depletion according to total lymphocyte count. Total lymphocyte count was weakly correlated with mid-upper arm circumference (r=0.20507); triceps skinfold thickness (r=0.29036), and length of hospital stay (r=-0.21518). Total lymphocyte count in different NRS categories differed significantly: older adults who were not at nutritional risk had higher mean and median total lymphocyte count (P=0.0245). Multiple regression analysis showed that higher lymphocyte counts were associated with higher triceps skinfold thicknesses and no nutritional risk according to the NRS. Conclusion – Total lymphocyte count was correlated with mid-upper arm circumference, triceps skinfold thickness, and nutritional risk according to the NRS. In multiple regression the combined factors that remained associated with lymphocyte count were NRS and triceps skinfold thickness. Therefore, total lymphocyte count may be considered a nutritional marker. Other studies should confirm these findings.

HEADINGS – Elderly nutrition. Lymphocyte count. Nutrition assessment.

INTRODUCTION

Older patients are commonly malnourished during hospital stay(8), and a high prevalence of malnutrition is found in hospitalized patients aged more than 65 years⁽¹³⁾. Therefore, the use of nutritional screening instruments⁽²⁴⁾ is critical given the prevalence of malnutrition and/or nutritional risk in hospitalized older adults(8,24).

In a recent prospective study, Calvo et al., 2012, found that 77% of hospitalized older adults were at risk of malnutrition or malnourished according to the mini-nutritional assessment, and that low serum albumin, cholesterol, and vitamins A and D were associated with malnutrition or risk of malnutrition⁽³⁾. Another recent study found better agreement between the calf circumference and body mass index of older adults⁽⁹⁾. Nutritional risk has been found to increase progressively in older adults compared with younger adults⁽¹⁹⁾. There is no gold standard for assessing the nutritional status of older adults, thus the use of many nutritional indicators is necessary to diagnose their nutritional status more accurately and establish an optimal nutritional intervention strategy(16,17).

This study aimed to investigate whether total lymphocyte count (TLC) is associated with other nutritional markers in hospitalized older adults.

METHODS

Characteristics and type of study, cases, and ethics

This cross-sectional study was conducted in a surgery unit of a general university hospital. A total of 131 older patients were recruited after approval of the local Research Ethics Committee.

Inclusion and exclusion criteria

The study inclusion criteria were age ≥60 years⁽¹⁶⁾, no terminal disease, nutritional assessment in the first 48 hours of hospital stay, and complete nutritional status information recorded the in medical records of the institution. Patients who had been hospitalized exclusively for clinical investigations or tests, for chemotherapy, radiotherapy, or hemodialysis, patients with inflammatory, immunological, and allergic diseases, infections, fever, sepsis, taking corticosteroids, or patients with incomplete nutritional status information in the medical records were excluded.

Data collection and study variables

The following variables were collected from the medical records: gender, age, length of hospital stay, death, Nutritional Risk Screening (NRS), and anthropometric data, namely body mass index (BMI), calf circumference (CC), mid-upper arm circumference (MUAC), triceps skinfold thickness (TST), mid-upper arm muscle circumference (MUAMC), and total lymphocyte count (TLC).

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Nutritional Risk Screening (NRS)

Nutritional risk was assessed by the NRS, a method proposed and validated by the European Society for Clinical Nutrition and Metabolism (ESPEN)^(10,11). The NRS takes into account recent weight loss, body mass index, low food intake, and disease severity, and classifies patients according to the final score: score \geq 3 indicates nutritional risk and score < 3 indicates no nutritional risk^(10,11).

Anthropometry

The anthropometric variables were measured using the standard procedures and classified according to the cut-off points established by the pertinent literature. BMI was classified as recommended by Lipschitz (1994)⁽¹⁴⁾, who defines the following cut-off points for older adults: underweight when BMI≤22; normal weight when 22<BMI<27, and overweight when BMI≥27. Calf circumference was classified as recommended by the World Health Organization (WHO)⁽²³⁾. The body composition indicators mid-upper arm circumference, triceps skinfold thickness, and mid-upper arm muscle circumference were classified as recommended by Frisancho (1990)⁽⁷⁾ and Burr & Phillips (1984)⁽²⁾.

Total lymphocyte count (TLC)

TLC was classified as recommended by Blackburn et al. (1) as follows: mild depletion when 1200 \(\text{TLC} \) \(\text{2000 cells/mm}^3 \);

moderate depletion when 800≤TLC≤1199 cells/mm³; and severe depletion when TLC<800 cells/mm³.

Statistical analysis

The patients were characterized by descriptive analysis with tables of frequency for the categorical variables and measures of position and dispersion for the continuous variables. The chi-square test or Fisher's exact test when necessary verified associations or compared proportions. The Mann-Whitney test compared continuous or ordinal measures between the groups. Finally, Spearman's linear correlation coefficient investigated the correlation between TLC and the nutritional indicators⁽⁴⁾. Multiple linear regression determined the parameters associated with lymphocyte count^(18,20). The significance level was set at 5% for all tests.

RESULTS

Table 1 shows the general description of the study variables. The patients had a mean age of 68.7 ± 6.9 years, most were male (65.65%), 41.22% of the sample was at nutritional risk according to the NRS, and 33.33% had low BMI. Calf circumference and TLC were low in 26.71% and 20.61% of the sample, respectively. The mean length of hospital stay was 10.4 ± 9.3 days. Table 1 also shows the means for the anthropometric variables and TLC.

TABLE 1. General description of the categorical and numerical variables (N=131)

Categorical variables		N	%
Gender	Female	45	34.35
	Male	86	65.65
Nutritional Risk Screening	At risk	54	41.22
	No risk	77	58.78
Total lymphocyte count	$<800 \text{ cel/mm}^3$	27	20.61
	800-1199 cel/mm ³	21	16.03
	≥1200 cel/mm ³	83	63.36
Body mass index	Overweight	33	25.58
	Normal weight	53	41.09
	Underweight	43	33.33
Calf circumference	<31 cm	35	26.72
	≥31 cm	96	73.28
Death	No	122	93.13
	Yes	9	6.87
Numerical variables	N	$X \pm SD$	Median
Age (years)	131	68.7 ± 6.9	67.0
Length of hospital stay (days)	131	10.4 ± 9.3	8.0
Body mass index (kg/m²)	129	24.3 ± 4.7	23.9
Mid-upper arm circumference (cm)	131	28.2 ± 4.1	27.5
Triceps skinfold thickness (mm)	131	17.8 ± 9.3	16.0
Mid-upper arm muscle circumference (mm)	131	228.5 ± 36.9	226.0
Calf circumference (cm)	131	33.3 ± 4.0	33.0
Total lymphocyte count (cell/mm³)	131	1528.9 ± 786.4	1520.0

Table 2 shows Spearman's linear correlation coefficient analyses, which found that TLC was weakly correlated with mid-upper arm circumference (r=0.20507), triceps skinfold thickness (r=0.29036), and length of hospital stay (r=-0.21518). Length of hospital stay decreased as TLC increased. However, TLC increased with mid-upper arm circumference and triceps skinfold thickness. TLC was not correlated with calf circumference.

TABLE 2. Correlation between total lymphocyte count and the study indicators

Indicators	Coefficient (r)	P-value	N
TLC vs length of hospital stay	-0.21518	0.0136	131
TLC vs age	-0.08249	0.3489	131
TLC vs body mass index	0.13500	0.1272	129
TLC vs mid-upper arm circumference	0.20507	0.0188	131
TLC vs triceps skinfold thickness	0.29036	0.0008	131
TLC vs mid-upper arm muscle circumference	-0.02702	0.7594	131
TLC vs calf circumference	0.08708	0.3227	131

The TLC of patients at nutritional risk also differed significantly from the TLC of patients not at nutritional risk according to the NRS. Older adults not at nutritional risk had higher mean and median TLC (P=0.0245). On the other hand, the TLC of patients in different categories of calf circumference and body mass index did not differ significantly (P=0.6716 and P=0.1077, respectively) (data not tabulated).

Individuals in different calf circumference categories (<31 cm and \ge 31 cm) had significantly different nutritional risk (P<0.0001) and body mass index (P<0.0001). Seventy-one percent of patients with calf circumference <31 cm were at nutritional risk according to the NRS (data not tabulated). The most common diseases found in the study sample were neoplasms (65.65%), digestive tract diseases (27.5%), and renal diseases (6.9%).

Nutritional risk (NRS) and anthropometric parameters (BMI, MUAC, TST, MUAMC, and CC) were assessed together by a multiple linear regression model with stepwise selection. The response variable (lymphocytes) was rank-transformed because it did not have normal distribution. Table 3 shows the factors that, together, remained associated with lymphocyte count, namely nutritional risk and TST. Higher TST and absence of nutritional risk according to the NRS were associated with higher lymphocyte counts.

TABLE 3. Results of multiple linear regression analysis with stepwise selection

Variables	Category	Beta *	EP **	P-value	$R^{2***} = 0.0934$
NRS	No risk vs At risk	10.088	6.71	0.0207	
TST		0.985	0.36	0.0065	

NRS: Nutritional Risk Screening; TST: Triceps Skinfold Thickness. * Beta: estimate or angular coefficient (slope) value in the regression line. ** EP: standard error beta. *** R^2 : determination coefficient. The response variable was rank-transformed because of the absence of normal distribution.

DISCUSSION

It is important to bear in mind that this study involved hospitalized patients from a surgery unit. Dudrick, 2011⁽⁵⁾, found that older surgical patients have higher morbidity and mortality, and higher rates of malnutrition and functional disorders than younger surgical patients.

The initial intent of the present study was to investigate whether TLC was related to anthropometric indicators and nutritional risk according to the NRS. None of the study variables differed between patients of the three TLC categories (based on the cut-off points). However, the numerical TLC values of patients in different categories of nutritional risk (P=0.0245), mid-upper arm circumference (P=0.0188), and triceps skinfold thickness (P=0.0008) differed significantly. The objective of the present study was based on the difficulty of assessing bedridden patients or patients who cannot be weighed due to locomotion problems. Hence, the study investigated the possibility of using TLC along with other nutritional indicators to assess the patients' nutritional status.

Other authors have already investigated the use of TLC and its relationship with the nutritional status of older adults. A study in Japan⁽¹²⁾ investigated whether TLC could be associated with other nutritional markers, such as anthropometry, the mini nutritional assessment, serum albumin, and cholesterol. Kuzuya et al.⁽¹²⁾ found that TLC (based on the cut-off point for depletion) was not associated with serum albumin, cholesterol, mini nutritional assessment, and anthropometry. These authors also found a negative correlation between TLC and age, but no correlation between TLC and nutritional markers⁽¹²⁾. Other studies⁽¹⁵⁾ also investigated TLC and serum albumin as markers of malnutrition in older adults aged more than sixty years with hip fractures, and found that both TLC and serum albumin behave as accurate and inexpensive clinical markers of malnutrition in such patients⁽¹⁵⁾.

The mini nutritional assessment is another widely used indicator for detecting malnutrition in older patients. This instrument has been considered by many authors as a useful and effective detector of nutritional risk^(13,19). Other instruments, such as the Malnutrition Universal Screening Tool (MUST) have also been considered effective and valid⁽¹⁶⁾. Another recent study⁽⁹⁾ reported that calf circumference was a better predictor of emerging care need in older adults than body mass index. The authors concluded that more studies are needed on anthropometry, especially calf circumference in older adult care⁽⁹⁾.

A study⁽²²⁾ of older adults investigated the relationship between anthropometric indicators and mortality, and found that mid-upper arm circumference was a better indicator than body mass index in older patients⁽²²⁾. Mid-upper arm circumference was more strongly associated with mortality than body mass index or calf circumference⁽²²⁾. Another study⁽²¹⁾ found that mid-upper arm circumference and calf circumference can predict the health status and risk of mortality of institutionalized older adults more effectively.

Given changes in nutritional status, the differences between the various indicators used for nutritional diagnosis^(3,6,9,16,17,21,24), and the peculiarities of older adults, new studies are needed on the nutritional status of hospitalized older surgical patients, and on continuous and routine nutritional status monitoring.

CONCLUSION

Although the different cut-off point-based categories of TLC were not associated with body mass index and calf circumference, TLC was significantly correlated with mid-upper arm circumference, triceps skinfold thickness, and nutritional risk according to the NRS in hospitalized older surgical patients. In multiple regression, the combined factors that remained associated with lymphocyte count were nutritional risk and triceps skinfold thickness. Therefore, TLC may be considered a nutritional marker. Other studies should confirm these findings.

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Authors' contributions

Leandro-Merhi VA conceived and designed the study, performed the statistical analyses, and wrote and reviewed the manuscript. Bráz VN collected and tabulated data and participated in manuscript writing. Aquino JLB participated in manuscript writing and review.

Leandro-Merhi VA, Bráz VN, Aquino JLB. A contagem de linfócitos totais apresenta alguma relação com marcadores nutricionais em idosos hospitalizados? Arq Gastroenterol. 2017,54(1):79-82.

RESUMO – Contexto – Durante a hospitalização é comum que pacientes idosos se encontrem desnutridos, sendo observada alta prevalência de pacientes idosos hospitalizados acima de 65 anos de idade. Objetivo – Investigar se a contagem de linfócitos totais apresenta alguma associação com outros marcadores nutricionais em pacientes idosos hospitalizados. Métodos – Pacientes idosos hospitalizados (N=131) foram recrutados em um estudo transversal. Investigou-se o estado nutricional por meio do *Nutritional Risk Screening* (NRS), antropometria e a contagem de linfócitos totais. A análise estatística foi realizada com os testes Qui-quadrado, Fischer e Mann-Whitney. Aplicou-se o coeficiente de correlação linear de Spearman para se investigar a presença de correlação entre a CLT e os marcadores nutricionais e para avaliação dos parâmetros associados à contagem de linfócitos foi utilizada a análise de regressão linear múltipla, com nível de significância de 5%. Resultados – O NRS mostrou 41,2% de pacientes com risco nutricional e 36% dos pacientes encontravam-se com depleção leve ou moderada na contagem de linfócitos totais. Houve correlação significativa, de fraca intensidade, entre a contagem de linfócitos totais e a circunferência de braço (r=0,20507); a prega cutânea triciptal (r=0,29036) e o tempo de internação (r=-0,21518). Verificou-se ainda diferença significativa na comparação da contagem de linfócitos totais com as categorias de risco nutricional pelo NRS; a contagem de linfócitos totais foi maior, em média ou mediana, entre os idosos que não apresentaram risco nutricional pelo NRS (*P*=0,0245). A análise de regressão múltipla mostrou que a maior contagem de linfócitos foi associada a valores maiores de prega cutânea triciptal e à classificação de sem risco nutricional pelo NRS. Conclusão – A contagem de linfócitos totais apresentou associação com a circunferência de braço, com a prega cutânea triciptal e com o NRS, podendo ser considerada como um marcador nutricional. Na regressão, os fatores que, em conjunto, ma

DESCRITORES – Nutrição do Idoso. Contagem de linfócitos. Avaliação nutricional.

REFERENCES

- Blackburn GL, Bristian BR, Maini BS, Schlamm HT, Smith MF. Nutritional and metabolic assessment of the hospitalized patient. The Journal of Parenteral and Enteral Nutrition. 1977;1:11-22.
- Burr ML, Phillips MK. Anthropometric norms in the elderly. Br Journal of Nutrition. 1984;51:165-9.
- Calvo I, Olivar J, Martínez E, Rico A, Díaz J, Gimena M. Mini Nutritional Assessment as a nutritional screening tool for hospitalized older adults; rationales and feasibility. Nutrición Hospitalaria. 2012;27:1619-25.
- Conover WJ. Practical Nonparametric Statistics. 3 ed. John Wiley & Sons Inc. Nova Iorque. 1999.
- Dudrick SJ. Nutrition management of geriatric surgical patients. Surg Clin North Am. 2011;91:877-96.
- Félix LN, Souza EMT. Avaliação nutricional de idosos em uma instituição por diferentes instrumentos. Revista de Nutrição. 2009;22:571-80.
- Frisancho AR. Anthropometric standards for the assessment of growth and nutritional status. Michigan: The University of Michigan Press; 1990.
- Holyday M, Daniells S, Bare M, Caplan GA, Petocz P, Bolin T. Malnutrition screening and early nutrition intervention in hospitalized patients in acute aged care: a randomised controlled trial. The Journal of Nutrition Health & Aging. 2012;16:562-8.
- Hsu WC, Tsai AC, Wang JY. Calf circumference is more effective than body mass index in predicting emerging care-need of older adults – results of a national cohort study. Clinical Nutrition. 2016;35:735-40.
- Kondrup J, Allison SP, Elia M, Vellas B, Plauth M. ESPEN guidelines for nutrition screening 2002. Clinical Nutrition. 2003;22:415-21.
- Kondrup J, Rasmussen HH, Hamberg O, Stanga Z, ESPEN Working Group. Nutritional risk screening (NRS 2002): a new method based on an analysis of controlled clinical trials. Clinical Nutrition. 2003;22:321-36.
- Kuzuya M, Kanda S, Koike T, Suzuki Y, Iguchi A. Lack of correlation between total lymphocyte count and nutritional status in the elderly. Clinical Nutrition. 2005;24:427-22
- Lara-Pulido A, Guevara-Cruz M, Malnutrition and associated factors in elderly hospitalized. Nutrición Hospitalaria. 2012;27:652-5.

- Lipschitz DA. Screening for nutritional status in the elderly. Prim Care. 1994;22:55-67.
- O'Daly BJ, Walsh JC, Quinlan JF, Falk GA, Stapleton R, Quinlan WR, O'Rourke SK. Serum albumin and total lymphocyte count as predictors of outcome in hip fractures. Clinical Nutrition. 2010;29:89-93.
- Poulia KA, Yannakoulia M, Karageorgou D, Gamaletsou M, Panagiotakos DB, Sipsas NV, Zampelas A. Evaluation of the efficacy of six nutritional screening tools to predict malnutrition in the elderly. Clinical Nutrition. 2012;31:378-5.
- Santos CA, Rosa COB, Ribeiro AQ, Ribeiro RCL. Patient-generated subjective global assessment and classic anthropometry: comparison between the methods in detection of malnutrition among elderly with câncer. Nutrición Hospitalaria. 2015;31:384-92.
- SAS System for Windows (Statistical Analysis System), version 9.4. SAS Institute Inc, 2002-12, Cary, NC, USA.
- Shpata V, Ohri I, Nurka T, Prendushi X. The prevalence and consequences of malnutrition risk in elderly albanian intensive care unit patients. Clinical Interventions in Aging. 2015;10:481-6.
- Tabachnick BG, Fidell LS. Using Multivariate Statistics. 4 ed. Allyn&Bacon. Needham Heights. MA. USA. 2001.
- Tsai ACH, Lai MC, Chang TL. Mid-arm and calf circumferences (MAC and CC) are better than body mass index (BMI) in predicting health status and mortality risk in institutionalized elderly Taiwanese. Archives of Gerontology and Geriatrics. 2012;54:443-7.
- Wijnhoven HAH, van Bokhorst-de van der Schueren MAE, Heymans MW, de Vet HCW, Kruizenga HM, Twisk JW, Visser M. Low Mid-Upper Arm Circumference, Calf Circumference, and Body Mass Index and Mortality in Older Persons. J Gerontol A Biol Sci Med Sci. 2010;65A:1107-14.
- World Health Organization. Physical status: the use and interpretation of anthropometry: report of a WHO Expert Committee. Geneva: WHO; 1995.
- Wu ML, Courtney MD, Shortridge-Baggett LM, Finlayson K, Isenring EA. Validity of the malnutrition screening tool for older adults at high risk of hospital readmission. J Gerontol Nurs. 2012;38:38-45.