

Original

Nutritional assessment of hospitalized HIV-infected patients by the phase angle z-score measurement

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Abstract

Background: This study analyzed the phase-angle (PA) values of hospitalized HIV-infected patients by comparing them with those reported for a healthy population and investigated their relation with nutritional parameters.

Methods: This is a cross-sectional study including 101 hospitalized patients diagnosed with HIV infection and evaluated by bioimpedance, anthropometry and biochemical tests. The phase angle values, weight loss percentage (%WL), body mass index (BMI), arm muscle circumference (AMC), tricipital skinfold (TSF), body fat percentage (%BF) and albumin were considered. In order to compare with values for the healthy population, the PA z-score of the patients under study was calculated. Spearman's correlation and the multiple linear regression model were used to identify nutritional parameters associated with the PA z-score.

Results: The patients showed a mean PA z-score of -2.6 ± 1.5 , and only 6.6% of them with a positive value. The PA z-score values correlated with %WL ($r = -0.51$; $p < 0.0001$), albumin ($r = 0.49$; $p < 0.0001$), BMI ($r = 0.58$; $p < 0.0001$), AMC ($r = 0.41$; $p < 0.0001$), TSF ($r = 0.47$; $p < 0.001$) and %BF ($r = 0.48$, $p < 0.0001$). In multiple analysis %WL ($p = 0.008$), albumin ($p = 0.01$), AMC ($p < 0.0001$) and %BF ($p = 0.0003$) remained associated with the score.

Conclusions: Low PA z-score values were observed, suggesting a worse clinical prognosis for the patients. The inclusion of the PA z-score as a nutritional indicator during care provision to HIV-infected patients is recommended.

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Key words: *Electrical bioimpedance. HIV. Nutritional status. Phase angle.*

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EVALUACIÓN NUTRICIONAL DE LOS PACIENTES HOSPITALIZADOS INFECTADOS POR EL VIH MEDIANTE LA MEDICIÓN DE LA PUNTUACIÓN Z DEL ÁNGULO DE FASE

Resumen

Antecedentes: Este estudio analizaba los valores del ángulo de fase (AF) de pacientes hospitalizados e infectados por el VIH al compararlos con los notificados para una población sana e investigaba su relación con los parámetros nutricionales.

Métodos: Se trataba de un estudio transversal que incluyó a 101 pacientes hospitalizados diagnosticados de infección por el VIH y en los que se evaluó la bioimpedancia, la antropometría y parámetros bioquímicos. Se consideraron los valores del ángulo de fase, el porcentaje de pérdida de peso (%PP), el índice de masa corporal (IMC), la circunferencia muscular del brazo (CMB), el pliegue tricipital (PT), el porcentaje de grasa corporal (%GC) y la albúmina. Para compararlos con los de una población sana, se calcularon las puntuaciones z del AF de los pacientes estudiados. Se emplearon la correlación de Spearman y el modelo de regresión lineal múltiple para identificar los parámetros nutricionales asociados con la puntuación z del AF.

Resultados: Los pacientes mostraron una puntuación z del AF de $-2,6 \pm 1,5$, el 23,7% de ellos tenían un valor superior a $-1,65$ y sólo el 6,6% un valor positivo. Los valores de las puntuaciones z del AF se correlacionaron con el %PP ($r = -0,51$; $p < 0,0001$), la albúmina ($r = 0,49$; $p < 0,0001$), el IMC ($r = 0,58$; $p < 0,0001$), la CMB ($r = 0,41$; $p < 0,0001$), el PT ($r = 0,47$; $p < 0,001$) y el %GC ($r = 0,48$, $p < 0,0001$). En el análisis múltiple, el %PP ($p = 0,008$), la albúmina ($p = 0,01$), la CMB ($p < 0,0001$) y el %GC ($p = 0,0003$) seguían asociándose con la puntuación z.

Conclusiones: Obtuvimos unos valores bajos de puntuación z del AF, lo que sugiere un peor pronóstico clínico en los pacientes. Recomendamos la inclusión de la puntuación z del AF como un indicador nutricional durante la provisión de la atención sanitaria a los pacientes infectados por el VIH.

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Palabras clave: *Bioimpedancia eléctrica. HIV. Estado nutricional. Ángulo de fase.*

Abbreviations

AC: Arm circumference.
AMC: Arm muscle circumference.
BIA: Bioimpedance.
BMI: Body mass index.
PA: Phase angle.
PW: Present weight.
TSF: Tricipital skinfold.
UW: Usual weight.
%BF: Body fat percentage.
%WL: Weight loss percentage.

Introduction

Nutritional assessment is a key tool to identify patients at nutritional risk and to more effectively treat HIV-infected patients.¹ Bioimpedance analysis (BIA) is widely used to evaluate body composition because it can be easily performed and it is a non-invasive low-cost method.² However, BIA does not directly measure body components, but it estimates resistance and reactance values, which are used in equations that calculate body components. Such equations have been developed for specific populations, and they are restricted to such populations.³

The phase angle (PA) is a parameter that is directly obtained from the relationship between reactance and resistance by a simple calculation, and it can be interpreted as a marker for muscle depletion and for nutritional status general compromising.^{4,6} Its values are influenced by chronic HIV infection and the existence of opportunistic diseases, and low values are associated with worse survival for HIV-infected patients, regardless of the immune deficiency level or viremia.^{7,8}

The interpretation of PA values in clinical practice has been limited to the consideration of only one cutoff point, without distinction between genders and patient ages. Values lower than 5.3° in HIV patients,⁷ to 6° in dialysis patients⁹ and to 4.5° in lung cancer patients¹⁰ are referred to, in the literature, as mortality predictors. The consideration of only one cutoff point for PA seems to be adequate for its interpretation as a prognostic factor, but not as a nutritional indicator.

This study analyzed PA values of hospitalized HIV-infected patients by comparing them with the distribution of values in a healthy population³ in terms of the z-score. The presence of a correlation between the PA z-score and nutritional parameters commonly used in clinical practice was also investigated.

Methodology

This cross-sectional study was approved by the local Research Ethics Committee, and participants provided written informed consent. The study was conducted in 2006 and 2007 and included 101 patients of both

genders who were older than 18 years and hospitalized with an HIV diagnosis confirmed by the ELISA (enzyme-linked immunosorbent assay) method, regardless of the disease stage. Pregnant women were excluded. Data concerning age, gender, antiretroviral therapy use and CD4 lymphocytes were collected.

Nutritional assessment, including BIA, anthropometry and biochemical testing, was performed up to two days after patients' hospitalization. BIA (equipment Model 101; 50 KHz, 800 μ A; RJL Systems, Detroit, MI) was used for PA estimation (arc-tangent reactance \times 180°/resistance \times π), which was expressed as the PA z-score (z-score = patient's observed value - median for the healthy population/standard deviation for the healthy population) of each evaluated patient. The values for the healthy population, according to gender and age range and as presented by Barbosa-Silva et al. (2005)³, were adopted as reference.

A Lange® adipometer, an extensible measuring tape and a digital scale with a stadiometer were used for anthropometric evaluation. Present weight (PW) and height as well as the tricipital (TSF), bicipital, suprascapular and suprailiac skinfolds and arm circumference (AC) on the patient's non-dominant side were measured. The patient was asked about his/her usual weight six month ago (UW). Then, the body mass index (BMI = PW/height²), body fat percentage (%BF) by the sum of the 4 skinfolds^{11,12}, arm muscle circumference (AMC = AC - TSF \times 0.314) and weight loss percentage in the last six months (%WL = (UW - PW)/UW \times 100) were calculated. In the laboratory analysis, albumin was dosed (bromocresol green method).

Data were expressed as percentage, mean \pm standard deviation or median and 1st and 3rd quartiles. Spearman's correlation and the multiple linear regression model were used to identify the nutritional parameters associated with the PA z-score. The model was adjusted for CD4 and included all the variables showing $p < 0.05$ in the correlation analysis. Statistical difference was considered when $p < 0.05$. Statistical analysis was performed by using software SAS 9.2 (Cary, NC, EUA).

Results

The patients were predominantly males, 40 years old on average and 89.4% of them showed CD4 values lower than 500 cell/mm³. A large number of patients (53.5%) did not use antiretroviral therapy (table I). 32.7% showed BMI lower than 18.5 kg/m², and 70.3% showed albumin lower than 3.5 g/dL. The patients showed a PA z-score mean of -2.6 ± 1.5 , with a median of -2.54 (-3.62 ; -1.6), minimum value of -6.0 and maximum value of 0.75 .

By comparing to normal PA values of a healthy population, the patients showed lower values, with only 6.6% of them showing a positive score value (fig. 1).

Table I
General characteristics of hospitalized HIV-infected patients

Variable	n = 101
Age (years)	39.8 ± 9.7
Males/ Females - n (%)	61 (60.4%)/40 (39.6%)
Patients using antiretroviral therapy - n (%)	47 (46.5%)
CD4 cell/mm ³	164 (61; 277)
BMI (kg/m ²)	20.9 ± 4.6
PA (°)	5.2 ± 1.3
PA z-score	-2.6 ± 1.5
Albumin (g/dL)	3.2 (2.8; 3.7)
AMC (cm)	21,7 ± 3,03
%BF (%)	23 (17,7; 33,0)

BMI = Body mass index; PA= Phase angle; AMC= Arm muscle circumference; %BF= Body fat percentage.

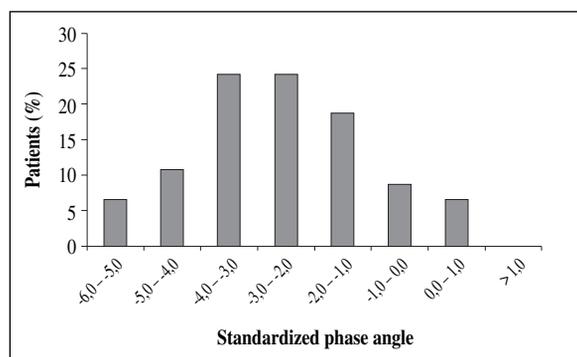


Fig. 1.—Distribution of the phase angle z-score of hospitalized HIV-infected patients by considering the normality values of a healthy population.

The PA z-score correlated with several nutritional parameters (table II), and such association was reinforced by multiple analysis for parameters %WL, albumin, AMC and %BF, regardless of the CD4 level, an indicator of the disease stage (table III).

Discussion

PA is considered as a cell integrity marker, and it has shown to be a prognostic factor in various clinical situations, but its use as a nutritional indicator is still incipient in clinical practice.

In several pathologies, an association between lower PA values and worse nutritional status has been observed.^{4,6} Maggiorie et al. (1996)⁴ identified PA association with AMC, albumin, protein intake and the subjective global assessment score (SGA) in hemodialysis-treated patients. When evaluating patients in the pre-operative period of gastrointestinal surgery, Barbosa-Silva et al. (2003)⁶ observed a significant decrease in PA values with malnutrition aggravation, as diagnosed by SGA, pointing to an association

Table II
Correlation of the phase angle z-score with nutritional parameters

Variable	r	p value
% WL (%)	-0.51	< 0.0001
Albumin (g/dL)	0.49	< 0.0001
BMI (kg/m ²)	0.58	< 0.0001
AMC (cm)	0.41	< 0.0001
TSF (mm)	0.47	< 0.0001
%BF (%)	0.48	< 0.0001

%WL = Weight loss percentage in relation to usual weight; BMI = Body mass index; AMC = Arm muscle circumference; TSF = Tricipital skinfold; %BF = Body fat percentage.

Table III
Multiple regression model for the phase angle z-score adjusted for CD4

Variable	Estimate	Standard error	p value
%WL (%)	-0.028	0.012	0.021
Albumin (g/dL)	0.626	0.232	0.009
AMC (cm)	0.179	0.065	0.007
%BF (%)	0.079	0.029	0.007
BMI (kg/m ²)	0.024	0.057	0.671
TSF (mm)	-0.025	0.035	0.479

%WL = Weight loss percentage in relation to usual weight; AMC = Arm muscle circumference; %BF = Body fat percentage; BMI = Body mass index; TSF = Tricipital skinfold.

between a low phase angle and subcutaneous fat tissue depletion, muscle mass depletion and body mass depletion, all parameters considered in SGA. However, in a study on patients before elective gastrointestinal surgery or hernia repair, the PA z-score presented weak agreement with BMI, SGA and Nutritional Risk Screening.¹³ Reinforcing the role played by PA as a nutritional status marker, in this study, a positive association was observed between that parameter and %BF, AMC and albumin, and a negative association was found with %WL even after adjustment for CD4 in hospitalized HIV-infected patients. Considering that this population is very susceptible to nutritional disorders, evidence of muscle or fat mass depletion suggests worse prognosis and, therefore, should be viewed with caution. Hence, the use of comparing the PA z-score in HIV-infected patients with that of a healthy population for detection of global nutritional deficit seems to be promising.

Considering the influence of malnutrition on HIV progression and on the prognosis of infected patients, higher PA values are associated with patients' longer survival,^{7,8} which has added to the use of PA as a nutritional parameter in this group of patients.

In the study by Schwenk et al. (2000),⁷ HIV-infected patients with PA smaller than 5.3° showed higher

mortality than those with larger values ($p < 0.001$), and relative mortality risk was 67% higher with a 1 degree reduction in PA. Additionally, the relationship of PA with CD4 lymphocytes and viral load reinforces its role as an indicator of disease status and its development.

The methodology for analysis of PA as a z-score adopted in this study was recently used by Paiva et al. (2011)¹⁴ in the prognostic assessment of patients with cancer. The authors observed that patients with PA z-score lower than -1.65 showed higher mortality (OR 3.12 CI: 2.03-4.79; $p < 0.00$) in relation to those with higher values.

Given the cross-sectional design in this study, population survival was not evaluated; however, 51.4% of the patients showed PA values lower than 5.3°, and 76.3% of them showed a lower PA z-score than -1.65. These data suggest a worse prognosis for this population.

In the nutritional evaluation by BIA, PA is obtained by a simple calculation, regardless of the pathology presented by the patient, since it is based on the relationship between reactance and resistance. These parameters are directly measured and not only estimated by the equipment. However, in order to determine free adipose and fat mass by BIA, formulas whose use is limited only to the population for which they were developed are used, and these equations are inadequate for other clinical situations, particularly in cases of alteration in hydration status.³

In the nutritional evaluation, considering the PA z-score is preferable to using its absolute values, since it allows for quantifying the distance for each value in relation to the distribution expected for a healthy population instead of using only one reference value as normality for the evaluation. Hence, negative PA z-score values suggest malnutrition, and the more negative such values are, the worse the nutritional status. This form of analysis facilitates the perception of the magnitude of nutritional compromising in these patients. Therefore, using PA for nutritional assessment in clinical practice seems to be extremely valuable due to its practicality and dependability, and it can be adopted as a nutritional parameter in the care for hospitalized HIV-infected patients.

Once the influence of gender and age on body composition, and consequently on the absolute values of PA¹⁵, is known, the availability of a reference standard which takes such factors into account, thus enabling PA expression as a z-score, makes it possible to compare individuals and populations with different characteristics (gender, age) as well as to monitor each individual's nutrition.

As to our knowledge, this is the first study that has quantified the dispersion of PA values in a population of HIV-positive individuals around values that are considered to be indicative of a healthy clinical condition and related the magnitude of such dispersion to parameters which are frequently used for that population's nutritional assessment.

A nutritional assessment routine for hospitalized patients must be established due to the important nutri-

tional alterations that occur in the course of HIV infection and to its impact on patients' clinical development.^{16,17} The PA z-score is a useful, practical, accessible and reproducible tool that can help with the assessment of hospitalized patients' nutritional status.

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