

# Assessment of the Nutritional Status of the Egyptian Patient with End Stage Liver Disease Prior to Liver Transplantation

Waheed A. Monsef<sup>1</sup>, Ibrahim Mostafa<sup>2</sup>, Doaa Zaky<sup>1\*</sup>

<sup>1</sup>Tropical Medicine Department, Ain Shams University, Cairo, Egypt

<sup>2</sup>Theodor Bilharz Research Institute, Gastroenterology and Hepatology Department, Cairo, Egypt

Email: [\\*drdoazakaria@gmail.com](mailto:drdoazakaria@gmail.com)

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## Abstract

**Background and Aim:** Patients with advanced liver disease have several risk factors to develop nutritional deficiencies. Accurate nutritional assessment is a real challenge because many of the traditionally measured parameters of nutritional status vary with severity of liver disease independently of nutritional status. The objective of this study was to assess the Egyptian patients with end stage liver disease and to compare different tools used to assess their nutritional status. **Patients and Methods:** 60 patients were nutritionally assessed by SGA, RFH-SGA anthropometry, handgrip dynamometry and biochemical tests. Clinical variables were cross analyzed with the nutritional assessment methods. **Results:** Malnutrition ranged from 7% by BMI and 100% by SGA. Agreement among all the methods was low compared with the SGA. Correlation between Malnutrition prevalence and the severity of liver disease was verified using Child-Pugh score more than MELD score. **Conclusion:** Malnutrition is highly prevalent among the Egyptian patients with end stage liver disease prior to liver transplantation. Although the diagnosis of nutritional status is not easy among this category of patients, it varied according to the method used. Nutritional support should be an important part of the preoperative care of liver transplantation patients.

## Keywords

Liver Transplantation; Nutritionalstatus; SGA; RFH-SGA and MELD

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## 1. Introduction

Hepatic cirrhosis (HC) is characterized by chronic and irreversible change in liver parenchyma, leading to alte-

\*Corresponding author.

rations in both liver structure and in the functional capacity of hepatocytes. Together, these changes result in progressive liver function loss, thus affecting nutritional status and body homeostasis in patients with HC to varying degrees. Protein caloric malnutrition (PCM) is highly prevalent in patients with terminal liver disease [1]. Roongpi-sunthinpong *et al.* suggest that PCM is dependent on its severity regardless of disease cause. However, malnutrition is associated with high morbidity and mortality, influencing the short and long term patient survival [2].

Impairment of cellular and humeral immune response in patients with HC may contribute to increased risk for infections [3], decreased quality of life and survival, increased length of hospitalizations and, therefore, increased hospital costs [4].

Malnutrition has been estimated to be present in 65% to 100% of patients with chronic hepatic diseases [5]-[8]. It has also been associated with increased ed morbidity and mortality in patients undergoing liver transplantation [4] [7] [9] [10], and the costs of transplant are significantly higher [11]. Thus, the nutritional status of these patients should be assessed early and routinely. However, commonly used methods, such as subjective global assessment, anthropometrics and bioimpedance analysis, are influenced or potentially influenced by the presence of liver disease per se [12] [13].

The subjective global assessment proposed by Detsky *et al.* [14] has been considered a good approach for subjective assessment of patients with liver disease [15], but water retention complicates its applicability since the estimated weight loss, which is part of this assessment, becomes impaired [16].

Royal Free Hospital-Subjective Global Assessment (RFH-SGA), a more advanced version which combines both subjective and objective parameters for nutritional assessment including measures of BMI calculated from estimated dry weight, MAMC, and details of dietary intake [17].

Although no consensus on parameter has the greatest value in assessing the nutritional state in CLDs [18], anthropometry can be comparable in accuracy to more sophisticated major tools [19]. The SFT has been identified as the most reliable measure that detects the effect of CLDs on nutritional status especially in the presence of ascites and/or edema [20]. SGA, anthropometric measurements and the HGS (handgrip strength) are more commonly used in routine nutritional assessment. However, there is no gold standard method of easy application and low cost, without subjective data and not influenced by the professional who performs it [21].

Thus, it is a challenge to adequately nutritionally assess these individuals, as none of the methods can be considered a gold standard [22].

Given the high prevalence of malnutrition and its relationship with morbidity and mortality in patients with liver cirrhosis as well as the absence of a gold-standard method for nutritional evaluation of these patients, we conducted the present study to assess the Egyptian patient with chronic liver disease patients on the waiting list for liver transplantation using different nutritional assessment tools.

## 2. Patients and Methods

Sixty patients with liver cirrhosis of different etiologies treated in hepatology department in Ain Shams University hospitals and Wady El-Nile Hospital, were evaluated between June 2012 and June 2013, after approval by the Ethics Committee of Ain Shams University. Patients older than 20 yrs. who were on the waiting list for liver transplantation at the time of the study were included. All subjects provided written informed consent

All patients that were included in the liver transplant waiting list presented major complications related to liver disease, with higher Meld and Child score at the time of inclusion.

### 2.1. Clinical Evaluation

Disease severity was classified according to the criteria proposed by Child-Pugh *et al.* [23], and MELD—the Model for End stage Liver Disease MELD [24]. Complications of cirrhosis, such as presence of ascites and/or edema at the time of evaluation and previous episode of encephalopathy, were also investigated. Due to the variation of the score of Child and Meld while on transplant waiting list, the values reported here in represent the values obtained in one given moment at the nutritional evaluation.

### 2.2. Nutritional Evaluation

Nutritional assessment was always undertaken by one observer as following:

1) **Nutritional assessment tools:**

- The subjective global assessment (SGA): The technique of uses clinical information collected during history taking and physical examination to determine nutritional status without recourse to objective measurements. This method of assessment has been successfully used to assess nutritional status in general medical and surgical patients, it shows good to excellent inter observer reproducibility and good convergent validity. The subjective global nutritional assessment (SGA) was carried out according to Detsky [14].
- Royal Free Hospital-Subjective Global Assessment (RFH-SGA): which include both subjective and objective parameters like BMI calculated from estimated dry weight, MAMC, and details of dietary intake. Intakes were categorized as adequate if they met estimated requirements, inadequate if they failed to meet estimated requirements but exceeded 500 kcal/day, or negligible if they provided fewer than 500 kcal/day. The three variables were incorporated into a semi structured, algorithmic construct, which allocates patients to one of three nutritional categories [17].

## 2) Anthropometric measurement:

- BW and BMI Before liver transplantation body weight (kg) was always measured after treating ascites and/or water retention with diuretics and/or paracentesis. Dry weight was estimated also considering the amount of the remaining fluid in ascites and edema evaluated clinically and by ultrasonography [25].
- Mid-arm circumference (MAC, cm) was measured at the midpoint between the tip of the acromion and the olecranon process on the non-dominant side of the body by using a flexible tape measure [26].
- Triceps skinfold (TSF) measurement (mm) was determined according to Durmin and Wormersley [27]. All the measurements were taken on the non-dominant side of the body, with the patients standing in a relaxed position, using a Harpenden skinfold caliper (John Bull British Indicators Ltd., St. Albans, UK).
- Mid-arm muscle circumference (MAMC), was calculated using the MAC and the TSF according to standard equations.  $MAMC = MAC (cm) - [0.314 - TST (cm)]$  [28].
- Functional methods: handgrip strength was measured by Hand Power Strength Measurement Grip Dynamometer which is considered a sensitive marker of body cell mass depletion. Three measurements were made on each arm and an average taken from all measurements.

## 3) Laboratory determinants:

- Biochemical evaluation included hemoglobin, hematocrit, total lymphocyte count and serum albumin. Hemoglobin and hematocrit were compared with reference values for males (14 - 18 g/dl and 40% - 57%, respectively) and females (12 - 16 g/dl and 37% - 47%, respectively). Total lymphocyte count values lower and  $1500/mm^3$  for TLC were considered indicative of malnutrition [29]. The cutoff value for albumin was  $<3.5$  g/dl for depletion diagnosis.

## 4) Statistics:

All data were statistically analyzed through statistical package for social science (SPSS) software version 13.0. Descriptive Data were expressed as number and percentage for categorized data, mean and SD for parametric data and mean, SD and median for laboratory variables. Analytical Data: Paired test: for comparing between 2 dependent means. Mann Whitney U test: for two-group comparisons in nonparametric data. The kappa test was applied to evaluate the extent of agreement between different methods used. For association of liver disease severity, nutritional indicators, Pearson's correlation was used.

Significance Was Evaluated as Follows:

P-value:  $> 0.05$  (non-significant).

P-value:  $< 0.05$  (significant).

P-value:  $< 0.01$  (highly significant).

## 3. Results

A total of 60 patients, 44 men and 22 women, Regarding age and sex, the age group 50 - 60 years was the most frequent age group presenting in 43.3%. As regard occupation, nearly 30% of patients were managerial (high income) & all were male. As regard residence, nearly half of them were from rural areas (56.7%). The main cause of end-stage liver disease was viral hepatitis with 39 cases; 65.0%, all of the patients had claimed abstinence from any alcoholic drinks, As regard child classification, all of them had liver decompensation, those with Child C score values were more than four folds those with Child B score values (49 and 11 respectively). The mean MELD value was  $17.42 \pm 4.2$  (6.0 - 25.0). Patients' characteristics are shown in [Table 1](#).

**Table 1.** Descriptive data of all studied patients.

Parameter	Male (No = 44)		Female (No = 16)		Total	
	No	%	No	%	No	%
<b>Age group</b>						
• <50 years	12	27.3	4	25.0	16	26.7
• ≥ 50: < 60 years	21	47.7	5	31.3	26	43.3
• ≥ 60 years	11	25.0	7	43.7	18	30.0
<b>Occupation</b>						
<b>Managerial</b> (high income)	18	41.0	0	0.0	18	30.0
<b>University graduated</b> (high Intermediate income)	10	23.0	0	0.0	10	16.7
<b>Employee</b> (low Intermediate income)	6	14.0	1	6.0	7	11.7
<b>Else</b>	10	23.0	1	6.0	11	18.3
<b>House-wife</b>	0	0.0	14	88.0	14	23.3
<b>Residence</b>						
<b>Urban</b>	19	43.2	7	43.8	26	43.3
<b>Rural</b>	25	56.8	9	56.3	34	56.7
<b>Indication for liver transplantation</b>						
<b>Focal lesion</b>	5	11.4	1	6.3	6	10.0
<b>Viral</b>	24	54.5	15	93.7	39	65.0
<b>Viral &amp; HFL</b>	13	29.5	0	0.0	13	21.7
<b>Viral &amp; bilharzial</b>	2	4.5	0.0	0.0	2	3.3
<b>Childpugh classification</b>						
<b>A</b>	0	0.0	0.0	0.0	0.0	0.0
<b>B</b>	8	18.2	3	18.8	11	18.3
<b>C</b>	36	81.8	13	81.2	49	81.7
<b>MELD Score</b>	18.2 ± 3.43		15.22 ± 4.31		17.42 ± 4.2	

81.7% of the enrolled patients confirmed they had decreased appetite, the majority of the patients had ascites and/or edema at the time of assessment (93.3%), and almost half of the patients had had a previous episode of encephalopathy (53.3%). The clinical data of patients according to liver disease are depicted in **Table 2**.

Laboratory assessment of the studied patients revealed that the percentage below and above cut off more frequent in Hb, PT, INR, PTT, Albumin, Na and Ca presenting in 81.7%, 88.3%, 81.7%, 85%, 88.3%, 61.7% and 80% respectively. The rest of lab. showed Median, As regard percentage below and above cut off more frequent in PLT, AST, *T. bilirubine*, *D. bilirubine*, urea and BUN presenting in 75%, 76.7%, 70%, 95%, 96.7% and 58.3% respectively, laboratory assessments of the patients are reported in **Table 3**.

Malnutrition ranged from 7% to 100% according to the different methods investigated. No significant differences were observed in the rates of malnutrition when comparing SGA, dynamometry, albumin and TLC (**Figure 1**). According to SGA, malnutrition was present in 100% of the patients, and of these, 24 patients (40%) were moderately and 36 (60%) were severely malnourished. The mean BMI was  $21.2 \pm 5.3 \text{ kg/m}^2$ , dynamometry was  $30.8 \pm 15.4 \text{ kgf}$  for males and  $22.9 \pm 13.4$  for females, MAC was  $20.6 \pm 6.9 \text{ cm}^2$ .

As for TSF, the median was  $8.3 \pm 1.8$  and TLC median was  $1164.0/\text{mm}^3$  (108 – 4416). The percentage of malnutrition according to sex, using the different tools is shown in **Figure 1**.

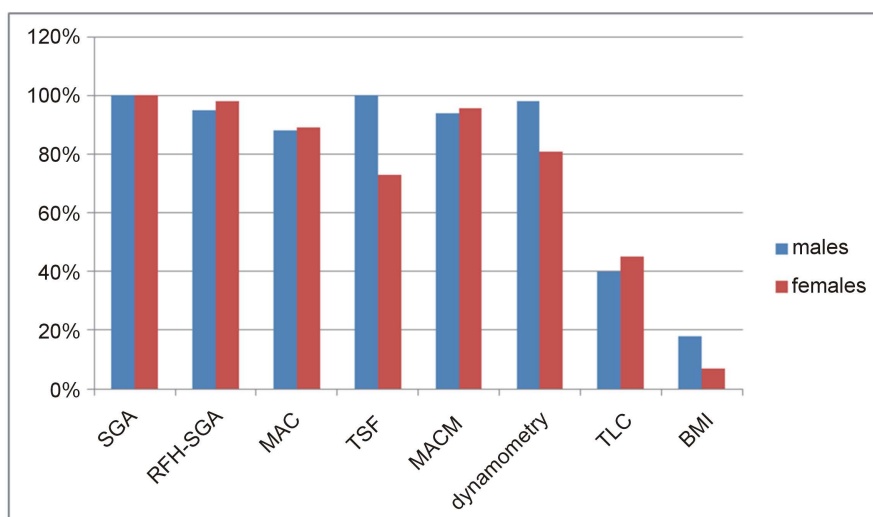
Agreement among all the methods, as performed by the kappa test, was low, except for RFH-SGA it showed perfect agreement with SGA (**Table 4**).

Kappa test,  $k < 0$  = no agreement;  $0 < k < 0.19$  = poor agreement;  $0.2 < k < 0.39$  = fair agreement;  $0.4 < k < 0.59$  = moderate agreement;  $0.6 < k < 0.79$  = substantial agreement;  $0.8 < k < 1.0$  = almost perfect agreement.

RFH-SGA, Royal Free Hospital -subjective global assessment; TLC, total lymphocyte count; MAC, midarm circumference; AMA, arm muscle area; TSF, triceps skinfold thickness; BMI, body mass index

Nutritional indicators showed negative correlations for both Child-Pugh method and Meld, but with higher correlations for the former. Among the indicators, MAMC ( $r = -0.32$ ,  $p = 0.01$ ), and dynamometry ( $r = -0.32$ ,  $p = 0.013$ ) showed a significant correlation for the severity criteria evaluated by Child-Pugh (**Table 5**, suggesting that the lower the values for nutritional indicators, worse is the severity of liver disease).

The correlation between laboratory and anthropometric variables & SGA showed that there is no correlation with most laboratory variables (p value > 0.05) except with Hb, there are negative correlation with SGA ( p value <0.05). As regard anthropometric variables, there is negative correlation between most anthropometric variables represent in (Total BW, dry BW, BMI total BW, BMI dry BW, MAC, TSF, MAMC and hand dynamometry.) & SGA (p value < 0.01) **Table 6**.



**Figure 1.** Percentage of malnutrition among male and female patients on the waiting list for liver transplantation according to different nutritional assessment tools SGA, subjective global assessment; RFH-SGA, Royal Free Hospital subjective global assessment; TLC, total lymphocyte count; MAC, midarm circumference; AMA, arm muscle area; TSF, triceps skinfold thickness; SSF, subscapular skinfold thickness; BMI, body mass index.

**Table 2.** Clinical examination of the patients.

Symptoms & signs	Male (No = 44)		Female (No = 16)		Total								
	No	%	No	%	No	%							
<b>Decrease of Appetite</b>	36	81.8	13	81.3	49	81.7							
<b>Dyspepsia</b>	35	79.5	13	81.3	48	80.0							
<b>Nausea</b>	25	56.8	11	68.8	36	60.0							
<b>Vomiting</b>	3	6.8	3	18.8	6	10.0							
<b>Diarrhea</b>	2	4.5	1	6.3	3	5.0							
<b>Constipation</b>	43	97.7	15	93.8	58	96.7							
<b>Fever</b>	8	18.2	4	25.0	12	20.0							
<b>Jaundice</b>	25	56.8	8	50.0	33	55.0							
<b>Hepatic encephalopathy</b>	25	56.8	7	43.7	32	53.3							
<b>LL.edema</b>													
<b>Nil</b>	3	6.8	1	6.3	4	6.7							
<b>Mild/Minimal</b>	8	18.2	3	18.8	11	18.3							
<b>Moderate</b>	19	43.2	7	43.7	26	43.3							
<b>Severe</b>	14	31.8	5	31.2	19	31.7							
<b>Ascites</b>													
<b>Clinical</b>	<b>U/S</b>												
<b>Nil</b>	<b>Nil</b>	4	4	9.1	9.1	0	0	0.0	0.0	4	4	6.7	6.7
<b>Mild/Minimal</b>	<b>Mild/Minimal</b>	6	6	13.6	13.6	3	4	18.8	25.0	9	10	15.0	16.7
<b>Moderate</b>	<b>Moderate</b>	18	19	40.9	43.2	8	7	50.0	43.7	26	26	43.3	43.3
<b>Sever</b>	<b>Sever</b>	16	15	36.4	34.1	5	5	31.2	31.3	21	20	35.0	33.3

**Table 3.** (a): Descriptives of laboratory data of studied patients at Start of Study presented as Mean (+/- SD) (normally distributed). (b): Descriptives of laboratory data of studied patients at Start of Study presented as Median (Interquartile Range) (Abnormally distributed).

(a)

Laboratory data	Normal values	Male (No = 44)		Female (No = 16)		Total	
		Mean (SD)	% below or above cut off	Mean (SD)	% below or above cut off	Mean (SD)	% below or above cut off
<b>Hb</b>	Male: 13.5 - 17.5 grams/dL Female: 12.0 - 15.5 grams/dL	10.8 ± (2.6)	81.8	10.1 ± (2.7)	81.3	10.6 ± (2.6)	81.7
<b>PT</b>	12 - 13 seconds	18.4 ± (5.6)	90.9	19.0 ± (6.2)	81.3	18.6 ± (5.7)	88.3
<b>INR</b>	0.8 - 1.2	1.7 ± (0.6)	79.5	1.8 ± (0.7)	87.5	1.7 ± (0.6)	81.7
<b>PTT</b>	25 - 35 seconds	41.4 ± (7.3)	84.1	41.7 ± (6.6)	87.5	41.5 ± (7.1)	85.0
<b>Albumin</b>	3.5 - 4.8 U/L	2.7 ± (0.7)	86.4	2.6 ± (0.7)	93.8	2.7 ± (0.7)	88.3
<b>Cholesterol</b>	120 - 200 mg/dl	106.0 ± (36.7)	2.3	110.3 ± (34.3)	0.0	107.1 ± (35.9)	1.7
<b>TG</b>	50 - 250 mg/dl	72.1 ± (28.1)	0.0	95.2 ± (46.3)	0.0	78.2 ± (35.0)	0.0
<b>Na</b>	137 - 145 mEq/L	133.8 ± (5.6)	61.4	132.5 ± (9.2)	62.5	133.5 ± (6.7)	61.7
<b>K</b>	3.6 - 5.0 mEq/L	4.1 ± (0.5)	20.5	4.1 ± (0.9)	25.0	4.1 ± (0.7)	21.7
<b>Ca</b>	8.9 - 10.4 mg/dl	8.2 ± (0.8)	79.5	8.0 ± (1.1)	81.3	8.1 ± (0.9)	80.0
<b>Cl</b>	98 - 110 mEq/L	104.5 ± (6.1)	18.2	105.3 ± (4.1)	6.3	104.7 ± (5.6)	15.0
<b>FBS</b>	65 - 110 mg/dL	122.6 ± (57.7)	31.8	108.4 ± (40.4)	31.3	118.8 ± (53.7)	31.7
<b>PPG</b>	70 - 145 mg/dL	159.9 ± (81.0)	34.1	143.1 ± (55.9)	31.3	155.4 ± (75.1)	33.3

(b)

Lab.	Normal values	Male			Female			Total		
		Median	25 - 75	% below or above cut off	Median	25 - 75	% below or above cut off	Median	25 - 75	% below or above cut off
<b>PLT</b>	150 - 450 billion/L	82.5	65.0 - 130.75	77.3	98.0	65.8 - 271.8	68.8	85.0	65.0 - 153.75	75.0
<b>Lymph. count</b>	1.00 - 4.00 × 10 <sup>9</sup> /L	1.0	0.67 - 1.53	45.2	1.2	0.78 - 1.83	25.0	1.1	0.7 - 1.61	39.7
<b>AST</b>	5 - 35 U/L	57.0	41.25 - 78.0	81.8	47.5	29.3 - 85.75	62.5	52.0	28.25 - 78.0	76.7
<b>ALT</b>	7 - 56 U/L	29.5	17.25 - 49.5	18.2	28.5	14.3 - 45.5	18.8	29.5	16.25 - 47.75	18.3
<b>T.bil</b>	0.2 - 1.3 mg/dl	2.2	1.3 - 4.0	70.5	2.25	1.0 - 4.3	68.8	2.25	1.3 - 4.0	70.0
<b>D.bil</b>	<0.3 mg/dl	1.2	0.7 - 2.3	97.7	1.1	0.43 - 2.6	87.5	1.18	0.7 - 2.33	95.0
<b>ALP</b>	38 - 126 U/L	107.0	75.5 - 129.75	29.5	86.5	57.8 - 109.8	12.5	103.0	70.25 - 127.75	25.0
<b>Urea</b>	10 - 20 mg/dl	29.5	20.25 - 43.0	97.7	27.5	22.8 - 64.25	93.8	29.0	21.18 - 53.75	96.7
<b>BUN</b>	7 - 21 mg/dl	23.5	16.0 - 43.75	61.4	21.0	14.0 - 28.5	50.0	23.0	16.0 - 40.0	58.3
<b>Creatine</b>	0.5 - 1.4 mg/dl	0.8	0.63 - 1.35	22.7	0.75	0.61 - 1.1	18.8	0.8	0.61 - 1.18	21.7

**Table 4.** Kappa values of different nutritional tools compared with SGA.

	RFH-SGA	BMI	MAC	MAMC	TSF	Dynamometry	TLC
<b>Kappa</b>	0.81	-0.01	0.26	0.24	0.19	0.83	0.18
<b>Level of agreement</b>	perfect	No	Fair	Fair	poor	substantial	poor

**Table 5.** Pearson's correlation between nutritional indicators and liver disease severity.

		BMI	MAC	MAMC	TS	Dynamometry	TLC
<b>MELD</b>	R-value	-0.0039	-0.1143	-0.13872	-0.28618	-0.18248	-0.14836
	P-value	0.6572	0.4022	0.18032	0.0628	0.2415	0.3424
<b>CHILD</b>	R-value	-0.1039	-0.1043	-0.3872	-0.28618	-0.32785	-0.33717
	P-value	0.5072	0.5022	0.0103	0.0628	0.01339	0.0270

**Table 6.** Laboratory & anthropometric variables correlation with SGA among studied patients at start of study.

Variables	Correlation with SGA	
	R	P value*
Hb	-0.360	0.005
PLT	0.149	0.149
Lymph.count	-0.40	0.768
PT	-0.148	0.259
INR	-0.196	0.133
PTT	-0.084	0.523
AST	0.004	0.978
ALT	-0.029	0.826
Albumine	-0.133	0.313
T.bil	-0.063	0.634
D.bil	-0.011	0.936
ALP	-0.212	0.104
Cholesterole	-0.033	0.802
TG	-0.079	0.549
Urea	+0.198	0.129
BUN	+0.082	0.533
Creatinine	+0.130	0.323
Na	+0.004	0.974
K	-0.034	0.795
Ca	-0.188	0.151
Cl	+0.071	0.592
FBS	-0.039	0.767
PPG	-0.025	0.850
Total.Wt	-0.349	0.006
Dry.Wt	-0.597	0.000
Ht	+0.271	0.076
BMI.T.Wt	-0.526	0.000
BMLDry.Wt	-0.778	0.000
MAC	-0.357	0.005
TSF	-0.233	0.033
MAMC	-0.363	0.004
Hand.dynamometry	-0.036	0.034

\* = Ranked Spearman correlation test.

#### 4. Discussion

For many years, malnutrition has been related to worse clinical outcomes and higher incidence of complications such as ascites, hepatic encephalopathy and infections, hepatorenal syndrome and diabetes mellitus [6]. It is considered a risk factor for morbidity and mortality in short and long term before and after transplantation [30] [31] as well as in abdominal surgery [30] [31]. A recent study by Sam and Nguyen 23 showed the association of PCM with longer hospitalization (8.7 days versus 5.7 days,  $p < 0.0001$ ), higher in-hospital mortality and a higher rate of readmissions [30].

The etiology of malnutrition in patients on the waiting list for liver transplantation is multifactorial. It has been associated with decreased food intake as a consequence of restrictive diets, nausea, vomiting and early satiety. All these are directly associated with: ascites; previous encephalopathy episodes; malabsorption and maldigestion because of gastrointestinal tract and hepatocyte dysfunction; changes in synthesis, metabolism and storage of nutrients; side effects of drug therapy [32] and hypermetabolism [33]. All these derangements make it difficult to assess the nutritional status of patients with liver disease. Thus, quite often malnutrition is under-diagnosed [12].

In this study, we have been able to show how difficult it can be to diagnose the nutritional status of these patients, because the prevalence of malnutrition ranged from 7% (by BMI) to 100% (by SGA) according to the different methods used [15]. Furthermore our results showed a low agreement rate among methods, as previously described by other authors [34]. We compared the different used nutritional tools with the SGA, the RFH-SGA showed perfect agreement ( $\kappa = 0.81$ ), both MAC and MAMC showed fair agreement, only the BMI showed no agreement at all.

SGA is a practical method for nutritional diagnosis. Its applicability to cirrhotic patients and to liver transplant candidates is valid, since liver disease changes the majority of objective indicators. The choice of SGA as the reference method was made because there is evidence that clinical evaluation is better than anthropometric or biochemical assessments [33]. SGA is recommended by ESPEN as a practical bedside method in assessing undernourished patients [35].

The study by Vulcano *et al.*, they found that according to SGA, 46.5% of the patients on the waiting list for liver transplantation showed a certain degree of malnutrition, and when weighed against the Child-Pugh criteria, for each unit of increase there is a 2.93 fold chance that the patient is malnourished. The other nutritional indicators did not show such strong association with the severity of liver disease [36]. Roongpisunthinpong *et al.* [2] also reported that SGA correlates with liver disease severity, thus indicating the usefulness of SGA in patients with chronic liver disease [2].

Adopting the SGA as a tool to diagnose malnutrition, revealed that our patients were 100% malnourished at the time of transplantation, this percentage is higher than that found by other authors [36], this may be explained by the late stage of liver disease in our study (all patients were Child class B & C, with average MELD score  $17.42 \pm 4.2$ ) However, SGA has limitations, such as the fact that some of the parameters evaluated (weight change, functional capacity and presence of ascites and edema) may be because of chronic liver failure and its treatment and may not be related to malnutrition.

Campillo *et al.* [37] proposed different BMI cut-off values for patients with liver cirrhosis depending on the presence and severity of ascites. Although we used the clinically estimated dry weight to assess the BMI of the patients, the lowest rates of malnutrition among all tools was that diagnosed by BMI in spite of exclusion of the factor of the ascites and lower limb edema. As reported in other study [38] malnutrition was also under diagnosed when Body Mass Index (BMI) was used, regardless of gender. However, malnutrition was more severe when AC and AMA were measured for females and males respectively. Further studies are needed to investigate if there is any relation between this finding and the high prevalence of obesity in Egypt.

In this study, the loss of muscle mass assessed by MAMC but not the decrease of fat reserve assessed by TSF was associated with severity of disease by Child-Pugh criteria, In study by Merli *et al.* [39], the reduction in muscle mass in Child-Pugh classes A and B assessed by MAMC was associated with decreased survival, but the depletion of fat did not influence survival in any Child-Pugh class. Nonetheless, techniques that use two compartments provide inadequate estimation of body composition in patients with cirrhosis, when compared with models that use four body compartments [17]. Alberino *et al.* [1] also refer a negative correlation between AMA and the Child-Pugh criteria, but not with TSF.

Dynamometry was a more sensible method to diagnose malnutrition in our patients. moreover, this parameter was associated with Child-Pugh classes, but not the MELD, Alvares-da-Silva and Reverbel da Silveira [34] recommended this method to assess nutritional status in the early phases of liver disease. However, the authors imply that dynamometry can lead to increased numbers of false positive diagnoses.

The use of biochemical tests for nutritional assessment in individuals with liver disease is questioned, because it may represent liver dysfunction and does not necessarily represent changes in nutritional status. Some authors use albumin as a nutritional index [2] [40], but Piquet *et al.* [41] found that the decrease in albumin in patients with cirrhosis is related to liver function and not to nutritional indices. In this study, we couldn't test the albumin as a nutritional marker as all included patients were Child B&C which means below normal albumin level by default.

Low lymphocyte count was found in the majority of chronic liver disease patients, probably influenced by the presence of infection, electrolyte imbalance, renal failure, metabolic stress, inflammatory bowel diseases or immunosuppression, conditions observed in patients with liver failure and thus impairing its use as a nutritional indicator [36]. TLC showed negative correlation with liver disease severity by Child and MELD, however showed poor agreement in diagnosis of malnutrition by SGA.

An inverse correlation between malnutrition (by SGA) and Hemoglobin level. Although hemoglobin is inde-



pendent of hepatic metabolism, they may be altered in patients with severe portal hypertension and hypersplenism as well as in the presence of gastrointestinal hemorrhages. It is noteworthy that laboratory parameters are affected by liver disease severity as well as by the etiological factors of the disease. Hence, they are more strongly correlated with disease severity than with malnutrition per se. Further studies are needed to explore the correlation between hemoglobin and the nutritional status of the liver disease patients and the possible causes regardless the liver condition.

Unlike the Child-Pugh score, the MELD score is not associated with the prevalence of malnutrition in regard to the severity of liver disease. The fact that only albumin and TLC were associated with MELD indicates that this parameter is related to the degree of liver failure rather than to malnutrition. In Egypt, since liver transplantation priority is based on MELD scores. We might thus assume that those who are transplanted first are not necessarily the more severely malnourished. On the other hand, mortality while on the waiting list might be associated with malnutrition the longer they remain on it. However, to confirm this theory, further studies are needed.

Similar studies proved the same outcome in which nutritional indicators showed higher correlation with the Child-Pugh criteria than with Meld [36]. It is noteworthy that, in the previous version of the Child-Pugh criteria, malnutrition level was considered as an indicator; however, the absence of an accurate tool for nutritional diagnosis has led to its exclusion. On the other hand, Meld estimation, so far, does not take nutritional status into account. Possibly, according to Stickel *et al.* [42], this model fails in predicting survival in 15-20% of recipients of liver transplants for not considering the nutritional status.

The methods utilized for the assessment of nutritional status in patients with liver cirrhosis in this study are simple and inexpensive, thus easily utilized in clinical practice. There are, of course, more sophisticated methods for the analysis of body composition, such as bioelectrical impedance analysis, dual-energy X-ray absorptiometry, K measurement or deuterium concentration, which are only used in special centers, require complex equipment and are certainly not as cost effective as the methods previously reported [43].

## 5. Conclusion

Malnutrition is highly prevalent among the Egyptian patients with end stage liver disease prior to liver transplantation. The diagnosis of nutritional status is not easy among this category of patients; it varied according to the method used. The agreement between the different methods is low. Body mass index was less suitable for malnutrition detection. The nutritional indicators that best correlated with the severity criteria proposed by Child-Pugh were, arm muscle circumference, arm muscle area and global assessment of nutritional status.

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## Abbreviation

SGA: Subjective global assessment

RFH-SGA: Royal free hospital-subjective global assessment

Meld: Model for end stage liver disease