# World Journal of Gastrointestinal Surgery

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# World Journal of Gastrointestinal Surgery

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#### **ABOUT COVER**

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#### **AIMS AND SCOPE**

The primary aim of World Journal of Gastrointestinal Surgery (WJGS, World J Gastrointest Surg) is to provide scholars and readers from various fields of gastrointestinal surgery with a platform to publish high-quality basic and clinical research articles and communicate their research findings online.

WJGS mainly publishes articles reporting research results and findings obtained in the field of gastrointestinal surgery and covering a wide range of topics including biliary tract surgical procedures, biliopancreatic diversion, colectomy, esophagectomy, esophagostomy, pancreas transplantation, and pancreatectomy, etc.

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**Retrospective Study** 

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ORIGINAL ARTICLE

# Comparison of different preoperative objective nutritional indices for evaluating 30-d mortality and complications after liver transplantation

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### Abstract

#### BACKGROUND

The nutritional status is closely related to the prognosis of liver transplant recipients, but few studies have reported the role of preoperative objective nutritional indices in predicting liver transplant outcomes.

#### AIM

To compare the predictive value of various preoperative objective nutritional indicators for determining 30-d mortality and complications following liver transplantation (LT).

#### **METHODS**

A retrospective analysis was conducted on 162 recipients who underwent LT at our institution from December 2019 to June 2022.

#### RESULTS

This study identified several independent risk factors associated with 30-d mortality, including blood loss, the prognostic nutritional index (PNI), the nutritional risk index (NRI), and the control nutritional status. The 30-d mortality rate was 8.6%. Blood loss, the NRI, and the PNI were found to be independent risk factors for the occurrence of severe postoperative complications. The NRI achieved the highest prediction values for 30-d mortality [area under the curve (AUC) = 0.861, P < 0.001] and severe complications (AUC = 0.643, P = 0.011). Compared to those in the high NRI group, the low patients in the NRI group had lower preoperative body mass index and prealbumin and albumin levels, as well as higher alanine aminotransferase and total bilirubin levels, Model for End-stage Liver Disease



scores and prothrombin time (P < 0.05). Furthermore, the group with a low NRI exhibited significantly greater incidences of intraabdominal bleeding, primary graft nonfunction, and mortality.

#### **CONCLUSION**

The NRI has good predictive value for 30-d mortality and severe complications following LT. The NRI could be an effective tool for transplant surgeons to evaluate perioperative nutritional risk and develop relevant nutritional therapy.

Key Words: Liver transplantation; Nutritional indicator; Complications; Prognosis; Nutrition assessment

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**Core Tip:** The preoperative nutritional status of liver transplant patients is closely related to prognosis. In this study, we analyzed clinical data from 162 patients to compare the value of different objective nutritional indices in predicting 30-d mortality and complications following liver transplantation. This provides insights for the preoperative assessment of liver transplant prognosis.

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#### INTRODUCTION

Liver transplantation (LT) is considered to be the most effective and definitive treatment option for patients suffering from end-stage liver disease. These conditions provide these patients with the opportunity not only to survive but also to extend their lifespan significantly. However, the occurrence of posttransplant complications remains prevalent and can greatly influence postoperative prognosis. This can largely be attributed to the compromised preoperative state of liver transplant recipients and the intricate nature of the surgical procedure. Recently, there has been increasing recognition of the critical roles played by preoperative nutrition and immune status in modulating surgical outcomes.

The serum prealbumin concentration, which can objectively reflect nutritional status and is almost unaffected by external supplementation, is an accurate biomarker for assessing the severity of liver disease. It can also be used for preoperative nutritional assessment and risk stratification[1-4]. The controlling nutritional status (CONUT), prognostic nutritional status index (PNI), and nutritional risk index (NRI) are widely used objective indicators for evaluating nutritional status. These indicators are associated not only with cancer-related complications but also with the long-term prognosis of cancer patients[5-10]. However, few studies have investigated the role of these nutritional indices in predicting liver transplant outcomes. Therefore, this study delves into this matter.

#### MATERIALS AND METHODS

#### Patient characteristics

This study was approved by the Ethics Committee of the People's Hospital of Guangxi Zhuang Autonomous Region. The inclusion criteria were as follows: (1) First-time liver transplant recipients aged 18-65 years; (2) Organ donation from deceased citizens; and (3) Complete clinical data. The exclusion criteria were: (1) Multiple organ transplants; (2) Severe pneumonia or severe cardiovascular and cerebrovascular diseases before surgery; (3) Receiving marginal livers[11]; or (4) Incomplete follow-up data. This study was approved by the ethics committee of the People's Hospital of Guangxi Zhuang Autonomous Region (KY-ZC-2023-056). All patients provided written informed consent for data analysis before transplantation.

#### Study design

Before performing a LT, patient demographic information, which include age, sex, body mass index (BMI), and relevant medical history, such as hypertension, diabetes, and hepatitis B, were collected. Additionally, donor age, graft weight, and various laboratory values, such as prealbumin, albumin (ALB), lymphocyte count, alanine aminotransferase (ALT), aspartate aminotransferase, total bilirubin, and creatinine, were collected. The Model for End-Stage Liver Disease (MELD) score, total cholesterol level, type of donor liver, prothrombin time (PT), and platelet count are also important factors to consider. During LT, data such as operating time, anhepatic phase time, total ischemic time, intraoperative blood loss, and intraoperative urine output were collected. After LT, the incidences of pneumonia, abdominal infection, abdominal bleeding, graft rejection, primary graft nonfunction, early graft dysfunction, severe complications (Clavien-Dindo grade  $\geq$ 



3), bile leakage, biliary stricture and mortality within 30 d were recorded.

Complications above Grade III include various conditions such as portal vein stenosis, portal vein thrombosis, hepatic artery stenosis, hepatic artery thrombosis, bile leakage, bile duct stenosis, retransplantation, pleural effusion requiring thoracentesis, peritoneal effusion requiring peritoneal puncture, intra-abdominal hemorrhage, respiratory failure necessitating extracorporeal membrane oxygenation support, liver insufficiency requiring artificial external liver support, renal failure requiring hemodialysis treatment, intracranial hemorrhage, and mortality.

The CONUT score consists of three components: The serum ALB concentration, total cholesterol concentration, and lymphocyte count[12]. The PNI can be calculated using the formula: ALB (g/L) +  $5 \times$  lymphocyte count (× 10<sup>9</sup>/mL). The following equation was used to determine the NRI: (1.519 × ALB, g/L) + (41.7 × actual body weight/ideal body weight) [13]. The ideal weight for males and females can be calculated as follows: For males, 2.3 kg per foot is added to a base weight of 50 kg (if height > 5 feet, with 1 foot equal to 30.48 cm); for females, 1.65 kg per foot is added to a base weight of 48.67 kg (if height > 5 feet, with 1 foot equal to 30.48 cm)[14]. If the actual weight exceeds the ideal weight, set the ratio to one[15].

This study aimed to analyze the risk factors associated with severe postoperative complications (Clavien-Dindo grade  $\geq$  3) and 30-d mortality following LT. Moreover, the researchers compared the effectiveness of the CONUT score, NRI, PNI, and prealbumin concentration as predictors of postoperative complications and mortality after LT using receiver operating characteristic (ROC) curves. Based on the area under the curve (AUC), the most accurate predictive index was identified and utilized to stratify patients into low-risk and high-risk groups using an appropriate cutoff value. Furthermore, the study compared the differences in baseline characteristics and postoperative complications between the two groups.

#### Postoperative assessment

After transplantation surgery and before discharge, the functionality of the transplant was assessed through routine laboratory tests. Surgical complications are typically diagnosed by evaluating clinical symptoms and conducting diagnostic examinations. These postoperative complications were documented in the patients' medical records. The Clavien-Dindo classification system was employed to assess and classify these complications. In this study, complications classified as Clavien-Dindo III or higher were considered severe. We recorded all adverse reactions, including pneumonia, abdominal infection, abdominal bleeding, graft rejection, primary graft nonfunction, early graft dysfunction, death, biliary leakage, and biliary stricture, in patients after surgery.

#### Statistical analysis

The statistical analysis was performed using SPSS 23.0 software. Continuous variables are represented using the median, 25<sup>th</sup> percentile, and 75<sup>th</sup> percentile, while categorical variables are represented using the frequency. A binary logistic regression model was used for both univariate and multivariate analyses of the entire sample. In the univariate analysis, indicators with a significance level of P < 0.05 were included in the multivariate analysis. However, given the existence of multicollinearity between the serum ALB concentration and the NRI, PNI, and CONUT score, the total serum ALB concentration was not incorporated into the multivariate analysis. The diagnostic results of multicollinearity, following the exclusion of ALB, indicated that the values for the variance inflation factor were less than 5. Consequently, no collinearity issues were observed within the model. The predictive values, optimal thresholds, sensitivities, and specificities for complications and mortality were calculated using ROC curves and AUC. MedCalc 10.2 software was used for the Z test, and the Delong test was used to compare the AUC of the different scoring systems. A P value of less than 0.05 was considered to indicate statistical significance.

#### RESULTS

#### Patient characteristics

A total of 162 patients were included in the study, including 133 males and 29 females. Fourteen patients (8.6%) died within a 30-d period following LT. The median age of the patients was 53.0 (45.0-57.0) years. The preoperative BMI was recorded as 23.0 (21.1-25.1). Preoperative hypertension was observed in 18 patients, diabetes was present in 22 patients, and 118 patients tested positive for hepatitis B surface antigen (Table 1).

#### Univariate and multivariate analyses of severe complications and mortality

The factors correlated with the 30-d mortality rate are outlined in Table 2. Univariate analysis revealed that the following factors were significantly correlated with 30-d mortality: BMI, operation time, blood loss, intraoperative urine volume, prealbumin concentration, NRI, CONUT, PNI, ALT, total bilirubin, preoperative MELD score, and PT. The multivariate analysis confirmed that blood loss [odds ratio (OR) = 1.001, 95% CI: 1.000-1.002, P = 0.034], the NRI (OR = 0.665, 95% CI: 0.446-0.991, *P* = 0.045), the CONUT (OR = 2.088, 95%CI: 1.016-4.291, *P* = 0.045), and the PNI (OR = 0.920, 95%CI: 0.848-0.997, P = 0.042) were risk factors for the 30-d mortality rate (Table 2).

Factors associated with severe complications (Clavien-Dindo grade  $\geq$  3) included operation time, blood loss, intraoperative urine volume, NRI, PNI, ALB, total bilirubin, preoperative MELD score, and PT. However, the results of the multivariate analysis showed that blood loss (OR = 1.003, 95%CI: 1.001-1.005, P = 0.004), the NRI (OR = 0.942, 95%CI: 0.901-0.986, *P* = 0.011), and the PNI (OR = 0.994, 95%CI: 0.989-0.999, *P* = 0.013) were risk factors associated with severe complications (Clavien-Dindo grade  $\geq$  3; Table 3).



Table 1 Patient characteristics			
Characteristics	Total (n = 162)		
Age, yr	53.0 (45.0-57.0)		
Male/female	133/29		
BMI	23.0 (21.1-25.1)		
Hypertension, yes/no	18/144		
Diabetes, yes/no	22/140		
HBsAg-positive, yes/no	118/44		
Operation time (min)	535.0 (440.0-600.0)		
Anhepatic phase (min)	58.0 (47.3-66.0)		
Donor age, yr	45.0 (36.0-55.0)		
Total ischemia time (min)	305.0 (250.5-372.6)		
Graft weight (kg)	1.4 (1.3-1.7)		
Split LT/whole LT	20/142		
Blood loss (mL)	1750.0 (975.0-3925.0)		
Intraoperative urine volume (mL)	2650.0 (1600.0-4000.0)		
Prealbumin (mg/L)	95.0 (89.7-101.8)		
NRI	95.1 (89.7-101.8)		
CONUT	6.0 (4.0-6.0)		
PNI	42.6 (38.7-46.4)		
ALB (g/L)	37.2 (33.6-40.7)		
Lymphocyte count (× $10^9$ /mL)	0.9 (0.6-1.5)		
Alanine aminotransferase (U/L)	31.0 (19.0-53.3)		
Aspartate aminotransferase (U/L)	46.5 (31.0-90.1)		
Total bilirubin (µmol/L)	30.8 (15.2-107.2)		
Creatinine (umol/L)	71.0 (58.8-85.3)		
Preoperative MELD score	12.0 (8.0-22.0)		
Total cholesterol (mmol/L)	3.4 (2.2-4.6)		
Prothrombin time (s)	16.4 (14.1-20.8)		
Platelet (× 10 <sup>9</sup> /mL)	67.0 (44.0-150.3)		
Death, yes/no	14/148		

The data are presented as the median (25th-75th percentile) or n. BMI: Body mass index; HBsAg: Hepatitis B surface antigen; CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index; MELD: Model for end-stage liver disease; LT: Liver transplantation; ALB: Albumin.

#### The value of different preoperative objective nutritional indicators for predicting severe complications and mortality

ROC curve analysis revealed that the NRI, CONUT score, PNI, and prealbumin concentration were significantly associated with 30-d mortality (P < 0.05). Among these, the NRI had the highest AUC value (0.861) for prediction (Tables 4 and 5, Figure 1). When the predictive ability of various indicators for severe complications was compared, the NRI, PNI, and prealbumin concentration all showed good predictive value for severe complications (P < 0.05). Among them, the NRI exhibited the highest predictive ability (AUC = 0.643; Tables 5 and 6, Figure 2).

#### Comparison of clinical characteristics and postoperative complications between the high NRI group and low NRI group patients

In terms of clinical characteristics, the high NRI group exhibited a greater BMI, improved liver function, and a lower preoperative MELD score than did the low NRI group. In terms of prognosis, the high NRI group had a significantly lower incidence of postoperative intra-abdominal bleeding, primary graft dysfunction, and 30-d mortality than did the low NRI group (P < 0.05). These findings are summarized in Tables 7 and 8.



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Table 2 Univariate and multivariate analyses of predictors of 30-d mortality						
Variables	Univariable OR (95%CI)	P value	Multivariable OR (95%CI)	P value		
Age	1.054 (0.996-1.114)	0.066				
Male	0.781 (0.204-2.999)	0.719				
BMI	0.772 (0.616-0.967)	0.024	0.720 (0.336-1.542)	0.397		
Hypertension	2.418 (0.606-9.648)	0.211				
Diabetes	1.067 (0.222-5.124)	0.936				
HBsAg-positive	0.644 (0.203-2.040)	0.454				
Operation time	1.005 (1.002-1.009)	0.008	1.004 (0.996-1.011)	0.367		
Anhepatic phase	1.039 (1.010-1.069)	0.091				
Donor age	1.004 (0.995-1.020)	0.475				
Total ischemia time	1.000 (0.996-1.005)	0.984				
Graft weight	1.002 (0.999-1.004)	0.253				
Split LT	0.938 (0.195-4.503)	0.936				
Blood loss	1.003 (1.001-1.004)	< 0.001	1.001 (1.000-1.002)	0.034		
Intraoperative urine volume	0.999 (0.999-1.000)	0.004	0.999 (0.998-1.000)	0.295		
Prealbumin	0.988 (0.977-0.999)	0.040	0.975 (0.929-1.023)	0.310		
NRI	0.258 (0.082-0.811)	0.020	0.665 (0.446-0.991)	0.045		
CONUT	5.756 (1.695-19.540)	0.005	2.088 (1.016-4.291)	0.045		
PNI	0.160 (0.051-0.500)	0.002	0.920 (0.848-0.997)	0.042		
ALB	0.798 (0.706-0.903)	< 0.001				
Lymphocyte count	0.723 (0.301-1.736)	0.468				
Alanine aminotransferase	1.002 (1.000-1.004)	0.045	1.002 (0.993-1.011)	0.639		
Aspartate aminotransferase	1.001 (1.000-1.002)	0.231				
Total bilirubin	1.007 (1.003-1.012)	0.001	1.004 (0.988-1.021)	0.606		
Creatinine	1.003 (0.999-1.007)	0.173				
Preoperative MELD score	1.099 (1.042-1.158)	< 0.001	1.003 (0.517-1.946)	0.994		
Total cholesterol	0.694 (0.452-1.065)	0.095				
Prothrombin time	1.114 (1.042-1.191)	0.001	0.773 (0.309-1.931)	0.773		
Platelet	1.002 (0.997-1.007)	0.432				

BMI: Body mass index; HBsAg: Hepatitis B surface antigen; CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index; MELD: Model for end-stage liver disease; OR: Odds ratio; LT: Liver transplantation; ALB: Albumin.

#### DISCUSSION

Early posttransplant mortality is the main factor affecting the overall effectiveness of LT, with most recipients dying within 1 mo after LT. In the current situation of severe shortage of donor livers and an increasing number of patients awaiting for LT, there is an urgent need for ideal risk prediction models to evaluate posttransplantation effectiveness and further determine the patients who are most likely to benefit from LT.

The MELD score is extensively applied in clinical practice and successfully predicts the likelihood of mortality in patients awaiting LT, as well as the risk of mortality after the transplant procedure[16,17]. However, the MELD score itself has limitations, as research has shown that it does not predict perioperative outcomes well in liver cancer patients without cirrhosis[18,19]. In recent years, scholars have shown greater interest in the relationship between nutritional status and post-LT complications. The serum prealbumin concentration serves as a reliable marker of liver synthesis capacity and nutritional status, making it a useful tool for predicting long-term survival in liver cancer patients undergoing liver resection[20]. Recent research has shown that prealbumin also demonstrates significant superiority in predicting complications after LT (AUC = 0.754)[1]. The COUNT score, PNI, and NRI are commonly used inflammatory nutritional indices in clinical practice. The CONUT score is a measure of the immune-nutritional status of patients and has been

Table 3 Factors that predict a Clavien-Dindo grade ≥ 3						
Variables	Univariable OR (95%CI)	P value	Multivariable OR (95%CI)	P value		
Age	0.997 (0.968-1.028)	0.852				
Male	0.518 (0.222-1.212)	0.129				
BMI	0.918 (0.819-1.028)	0.138				
Hypertension	1.073 (0.359-3.210)	0.900				
Diabetes	1.044 (0.380-2.868)	0.934				
HBsAg-positive	0.813 (0.377-1.754)	0.598				
Operation time	1.004 (1.001-1.006)	0.004	1.003 (1.000-1.006)	0.078		
Anhepatic phase	1.019 (0.999-1.041)	0.069				
Donor age	1.010 (0.998-1.022)	0.113				
Total ischemia time	1.001 (0.999-1.003)	0.350				
Graft Weight	1.065 (0.978-1.158)	0.146				
Split LT	1.515 (0.477-4.812)	0.582				
Blood loss	1.004 (1.002-1.005)	< 0.001	1.003 (1.001-1.005)	0.004		
Intraoperative urine volume	0.998 (0.996-1.000)	0.042	0.999 (0.995-1.002)	0.382		
Prealbumin	0.995 (0.990-1.001)	0.089				
NRI	0.945 (0.904-0.988)	0.013	0.942 (0.901-0.986)	0.011		
CONUT	1.037 (0.984-1.094)	0.169				
PNI	0.856 (0.738-0.994)	0.041	0.994 (0.989-0.999)	0.013		
ALB	0.910 (0.848-0.977)	0.009				
Lymphocyte count	1.113 (0.814-1.522)	0.502				
Alanine aminotransferase	1.002 (0.999-1.004)	0.138				
Aspartate aminotransferase	1.000 (1.000-1.001)	0.314				
Total bilirubin	1.004 (1.002-1.006)	< 0.001	1.005 (0.999-1.010)	0.079		
Creatinine	1.001 (0.997-1.004)	0.685				
Preoperative MELD score	1.057 (1.020-1.097)	0.003	0.894 (0.763-1.047)	0.165		
Total cholesterol	0.886 (0.721-1.088)	0.886				
Prothrombin time	1.075 (1.019-1.134)	0.009	1.075 (0.923-1.252)	0.354		
Platelet	1.001 (0.998-1.005)	0.422				

BMI: Body mass index; HBsAg: Hepatitis B surface antigen; CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index; MELD: Model for end-stage liver disease; LT: Liver transplantation; ALB: Albumin.

Table 4 Values of different preoperative objective nutritional indicators for predicting 30-d mortality						
	AUC	Sensitivity	Specificity	95%CI	Optimal threshold value	<i>P</i> value
CONUT	0.724	0.58	0.80	0.646-0.794	6	0.015
NRI	0.861	0.70	0.83	0.765-0.958	88	< 0.001
PNI	0.781	0.64	0.80	0.682-0.829	38	0.001
Prealbumin	0.666	0.76	0.60	0.589-0.754	79	0.003

AUC: Area under the curve; CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index.

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Table 5 Comparisons of the area under the curve of various nutritional indicators for predicting severe complications and 30-d mortality

POC	Severe complications		30-d mortality		
RUC	Z value	<i>P</i> value	Z value	<i>P</i> value	
CONUT vs NRI	1.851	0.064	1.550	0.121	
CONUT vs PNI	1.945	0.051	0.832	0.405	
CONUT vs Prealbumin	0.818	0.413	0.490	0.623	
NRI vs PNI	0.749	0.454	1.061	0.288	
NRI vs Prealbumin	0.582	0.560	2.337	0.019	
PNI vs Prealbumin	0.176	0.860	1.062	0.288	

CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index; ROC: Receiver operating characteristic.

Table 6 Values of different preoperative objective nutritional indicators for predicting severe complications						
	AUC	Sensitivity	Specificity	95%CI	Optimal threshold value	P value
CONUT	0.547	0.17	0.96	0.463-0.627	8	0.410
NRI	0.643	0.50	0.72	0.555-0.712	91	0.011
PNI	0.615	0.23	0.94	0.522-0.678	34	0.047
Prealbumin	0.603	0.63	0.61	0.533-0.695	82	0.027

AUC: Area under the curve; CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index.



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Figure 1 Receiver operating characteristic curve for the cutoff values of multiple preoperative objective nutritional indicators for predicting postoperative death. CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index; AUC: Area under the curve.

reported to independently predict the prognosis of various malignant tumors[21]. Among liver transplant patients, those with intermediate to high CONUT scores have a higher incidence of postoperative Clavien-Dindo grade III/IV complications and infections than do those with high CONUT scores[22]. The PNI has a certain role in predicting post-LT renal injury and postliver cancer recurrence[23,24]. The NRI was first introduced in 2005 as an objective nutritional assessment tool that accurately predicts the mortality rate of elderly patients in internal medicine[25]. Subsequent multicenter studies have demonstrated that patients with an NRI of 98 or lower have a 1.5-fold greater risk of postoperative complications following abdominal surgery than patients with an NRI above 98. However, to date, the relationship between the NRI and post-LT complications has not been thoroughly elucidated[26]. Therefore, we further investigated the relationship between nutritional indices and the prognosis of LT patients.

Table 7 Comparation of the clinical characteristics among different nutritional risk index groups					
Characteristics	Low NRI ( <i>n</i> = 30)	High NRI ( <i>n</i> = 132)	P value		
Age, yr	53.0 (44.0-56.0)	52.5 (46.0-58.0)	0.587		
Male/female	26/4	107/25	0.602		
BMI	21.5 (19.0-23.4)	23.3 (21.4-25.4)	0.012		
Hypertension, yes/no	4/26	14/118	0.747		
Diabetes, yes/no	5/25	17/115	0.563		
HBsAg-positive, yes/no	21/9	97/35	0.820		
Operation time (min)	540.0 (452.5-650.3)	520.0 (440.0-600.0)	0.344		
Anhepatic phase (min)	58.0 (48.5-65.0)	57.0 (47.0-66.0)	0.719		
Donor age, yr	46.5 (39.0-59.6)	43.0 (35.1-53.0)	0.651		
Total ischemia time (min)	329.5 (271.4-395.0)	286.5(234.7-356.2)	0.323		
Graft weight (kg)	1.3 (1.2-1.6)	1.5 (1.4-1.8)	0.409		
Split LT/whole LT	2/28	18/114	0.373		
Blood loss (mL)	2000.0 (850.0-5000.0)	1650.0 (925.0-3500.0)	0.305		
Intraoperative urine volume (mL)	2600.0 (1650.0-3225.0)	2800.0 (1600.0-4000.0)	0.636		
Prealbumin (mg/L)	56.0 (33.0-82.0)	109.5 (54.5-172.0)	< 0.001		
NRI	83.8 (81.2-85.5)	98.8 (93.1-103.5)	< 0.001		
CONUT	8.0 (7.0-9.8)	5.0 (4.0-6.0)	< 0.001		
PNI	34.9 (31.7-38.9)	43.7 (40.3-47.6)	< 0.001		
ALB (g/L)	29.8 (27.4-31.2)	39.0 (35.6-41.4)	< 0.001		
Lymphocyte count (× 10 <sup>9</sup> /mL)	0.9 (0.5-1.6)	0.9 (0.6-1.4)	0.978		
Alanine aminotransferase (U/L)	41.5 (19.8-62.5)	29.5 (19.0-45.8)	0.154		
Aspartate aminotransferase (U/L)	64.5 (38.8-126.3)	42.0 (29.3-77.5)	0.008		
Total bilirubin (µmol/L)	53.0 (20.4-250.4)	28.4 (14.8-87.8)	0.047		
Creatinine (umol/L)	73.0 (57.0-86.3)	70.0 (59.3-85.0)	0.848		
Preoperative MELD score	15.0 (11.8-24.8)	11.0 (7.3-20.8)	0.028		
Total cholesterol (mmol/L)	2.7 (2.0-4.4)	3.6 (2.3-4.6)	0.050		
Prothrombin time (s)	17.8 (16.2-23.4)	15.9 (14.1-20.0)	0.037		
Platelet (× $10^9/mL$ )	63.5 (32.5-143.0)	67.5 (47.0-152.8)	0.386		

BMI: Body mass index; HBsAg: Hepatitis B surface antigen; CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index; MELD: Model for end-stage liver disease; LT: Liver transplantation; ALB: Albumin.

This retrospective analysis revealed that the NRI, PNI, and prealbumin have certain value for predicting 30-d mortality and severe complications in liver transplant recipients, with the NRI having the highest AUC value. The CONUT score can predict 30-d mortality in liver transplant recipients but cannot predict severe postoperative complications. In the multifactorial logistic regression analysis, blood loss, NRI, PNI, and CONUT were independent predictors of 30-d mortality, while blood loss, NRI, and PNI were independent predictors of severe postoperative complications. Based on the optimal cutoff value of the NRI, patients with an NRI > 88 had better preoperative liver function; lower rates of intraabdominal bleeding (6.1% *vs* 20.0\%, *P* = 0.025) and primary graft nonfunction (1.5% *vs* 10.0\%, *P* = 0.044); and lower mortality rates (6.1% *vs* 20.0\%, *P* = 0.025) than patients with an NRI < 88.

The serum prealbumin concentration has good predictive ability for 30-d mortality and severe complications after LT, consistent with previous findings[1]. The variation in AUC values may be attributed to varying definitions of severe complications. Serum prealbumin is a carrier protein entirely produced by liver cells, and its main physiological function is to transport thyroid hormones and vitamin A, enhancing the body's immune function by promoting lymphocyte maturation [27]. Moreover, due to its short half-life and small amount of interference factors, prealbumin can sensitively reflect liver synthesis function and has high sensitivity and specificity for detecting hepatocyte damage[28,29]. Therefore, the serum prealbumin concentration can be a potential indicator for predicting poor early outcomes after LT.

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Table 8 Comparison of postoperative complications between the low nutritional risk index group and the high nutritional risk index aroun n (%)

	Total ( <i>n</i> = 162)	Low NRI ( <i>n</i> = 30)	High NRI ( <i>n</i> = 132)	P value
Pneumonia	37 (22.8)	8 (26.7)	29 (22.0)	0.631
Intra-abdominal infection	20 (12.3)	4 (13.3)	16 (12.1)	0.767
Intra-abdominal bleeding	14 (8.6)	6 (20.0)	8 (6.1)	0.025
Graft rejection	6 (3.7)	2 (6.7)	4 (3.0)	0.308
Primary graft nonfunction	5 (3.1)	3 (10.0)	2 (1.5)	0.044
Early graft dysfunction	4 (2.5)	0 (0.0)	4 (3.0)	1.000
Mortality	14 (8.6)	6 (20.0)	8 (6.1)	0.025
Clavien-Dindo grade $\geq 3$	43 (26.5)	12 (40)	31 (23.5)	0.071
Biliary leakage	3 (1.9)	1 (3.3)	2 (1.5)	0.461
Biliary stricture	4 (2.5)	2 (6.7)	2 (1.5)	0.156

NRI: Nutritional risk index.



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#### Figure 2 Receiver operating characteristic curve for the cutoff values of multiple preoperative objective nutritional indicators for predicting severe complications. CONUT: Control nutritional status; PNI: Prognostic nutritional index; NRI: Nutritional risk index; AUC: Area under the curve.

We found that the AUC for predicting 30-d mortality was the highest for the NRI, followed by the PNI, CONUT, and prealbumin concentration. Similarly, the AUC for predicting severe complications was the highest for the NRI, followed by the PNI and prealbumin concentration. Although both the NRI and the PNI incorporate the measurement of ALB, the NRI also reflects the degree of weight loss in patients. Malnutrition is prevalent among patients with end-stage liver disease, and the incidence of malnutrition in individuals with decompensated cirrhosis and liver failure ranges from 50% to 90% [30]. Surgical intervention exacerbates liver injury, reduces ALB synthesis, impairs immune function and body repair capacity, increases the likelihood of postoperative complications, and adversely affects survival prognosis in malnourished patients. Recent studies have elucidated the association between sarcopenia and the prognosis of liver transplant recipients[22,31]. These findings indicate that diminished muscle mass is linked to unfavorable outcomes following LT and is a predictive factor for short-term survival. Furthermore, low muscle mass has an equally significant impact on the prognosis of patients with malignancies. In patients with nonmetastatic breast cancer, the overall mortality rate is significantly greater in individuals with sarcopenia (hazard ratio, 1.41; 95%CI, 1.18-1.69)[32]. Similarly, among patients diagnosed with colorectal cancer, those exhibiting sarcopenia have a notably elevated overall mortality rate (hazard ratio, 1.27; 95% CI, 1.09-1.48) compared to that of their counterparts without sarcopenia[33]. This finding suggested a strong association between wasting and unfavorable prognosis. However, the existing evidence is insufficient to establish a definitive link between lymphocyte count and nutritional status. Consequently, the predictive efficacy of the NRI surpasses that of the PNI, highlighting its potential in clinical prognostication. The CONUT score incorporates cholesterol as an indicator. Cholesterol is primarily synthesized in the liver, and its levels indirectly reflect liver synthetic function. Changes in liver function due to cellular damage can lead to alterations in cholesterol levels. Although a smallscale study suggested an association between low cholesterol levels and unsuccessful liver transplant, there is currently

insufficient evidence to support the role of cholesterol in the prognosis following LT[34]. Therefore, these findings may explain why the predictive efficacy of the CONUT score is lower than that of the NRI and PNI.

This study aimed to compare the role of multiple objective nutritional indicators in predicting the prognosis of LT patients, thereby facilitating a comprehensive preoperative nutritional assessment, early identification of malnutrition, timely and appropriate nutritional support for enhancing surgical safety, and reducing the incidence of postoperative complications. This study has several limitations, including the following: (1) The sample size was not large enough; (2) This was a retrospective analysis, and further prospective analysis is needed to clarify the predictive value of different scoring systems for post-LT outcomes; and (3) We analyzed only a portion of the nutritional indicators and did not include all nutritional indicators in our analysis. Despite these limitations, our results still demonstrate the superiority of the NRI as a nutritional indicator for predicting post-LT 30-d mortality and severe complications.

#### CONCLUSION

This study identified several independent risk factors associated with 30-d mortality, including blood loss, the PNI, the NRI, and the CONUT. The 30-d mortality rate was 8.6%. Blood loss, the NRI, and the PNI were found to be independent risk factors for the occurrence of severe postoperative complications. The NRI achieved the highest predictive values for 30-d mortality (AUC = 0.861, P < 0.001) and severe complications (AUC = 0.643, P = 0.011). Compared to those in the high NRI group, the patients in the low NRI group had lower preoperative BMIs; prealbumin, and ALT levels; and higher ALT, total bilirubin, MELD score, and PT (P < 0.05). Furthermore, the low NRI group exhibited significantly greater incidences of intraabdominal bleeding, primary graft nonfunction, and mortality. In conclusion, the NRI can serve as an effective tool for transplant surgeons to assess perioperative nutritional risk in patients and formulate relevant nutritional interventions.

### ARTICLE HIGHLIGHTS

#### Research background

Nutritional status is closely associated with the prognosis of liver transplantation (LT) patients.

#### Research motivation

However, few studies have thoroughly investigated the relationship between the preoperative nutritional status of liver transplant recipients and postoperative prognosis. In clinical practice, there is a lack of a simple and effective tool for assessing the nutritional risk of patients during the perioperative period and for predicting the outcomes of LT.

#### Research objectives

The objective of this study was to compare the value of different preoperative objective nutritional indicators for predicting the 30-d mortality and the incidence of complications following LT.

#### Research methods

This study conducted a retrospective analysis of clinical data from 162 patients who underwent LT. The present study compared the ability of the serum prealbumin concentration, the controlling nutritional status (CONUT) score, the prognostic nutritional index (PNI), and the nutritional risk index (NRI) to predict the 30-d mortality rate and the incidence of severe complications after LT. This study also aimed to analyze the risk factors associated with the 30-d mortality rate and incidence of severe complications after LT. The area under the receiver operating characteristic curve was used to select the index with the best predictive ability. Patients were then divided into low-risk and high-risk groups based on the optimal cutoff value, and the differences in postoperative complications and mortality rates between the two groups were compared.

#### Research results

This study identified several independent risk factors associated with 30-d mortality, including blood loss, the PNI, the NRI, and the CONUT. The 30-d mortality rate was 8.6%. Blood loss, the NRI, and the PNI were found to be independent risk factors for the occurrence of severe postoperative complications. The NRI achieved the highest prediction values for 30-d mortality [area under the curve (AUC) = 0.861, P < 0.001] and severe complications (AUC = 0.643, P = 0.011). Compared to those in the high NRI group, the patients in the low NRI group had lower preoperative body mass index and prealbumin and albumin levels, as well as higher alanine aminotransferase and total bilirubin levels, Model for Endstage Liver Disease scores and prothrombin time (P < 0.05). Furthermore, the group with a low NRI exhibited significantly greater incidences of intraabdominal bleeding, primary graft nonfunction, and mortality.

#### **Research conclusions**

The NRI has good predictive value for 30-d mortality and severe complications following LT. The NRI could be an effective tool for transplant surgeons to evaluate the perioperative nutritional risk and provide relevant nutritional therapy.



#### Research perspectives

The purpose of this study was to investigate the predictive value of different objective nutritional indicators before surgery for the outcome of LT.

#### FOOTNOTES

Author contributions: Li C conceived the study, collected and analyzed the data and drafted the manuscript; Chen HC assisted with the data collection; Lai YH revised the manuscript; and all the authors read and approved the final version to be published.

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