## **ORIGINAL ARTICLE**



# Nutritional status and risk of contrast-associated acute kidney injury in elderly patients undergoing percutaneous coronary intervention

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# Abstract

**Background** This study aimed to investigate the connection between malnutrition evaluated by the Controlling Nutritional Status (CONUT) score and the risk of contrast-associated acute kidney injury (CA-AKI) in elderly patients who underwent percutaneous coronary intervention (PCI).

**Methods** A total of 1308 patients aged over 75 years undergoing PCI was included. Based on the CONUT score, patients were assigned to normal (0–1), mild malnutrition (2–4), moderate-severe malnutrition group ( $\geq$ 5). The primary outcome was CA-AKI (an absolute increase in  $\geq$  0.3 mg/dL or  $\geq$  50% relative serum creatinine increase 48 h after contrast medium exposure).

**Results** Overall, the incidence of CA-AKI in normal, mild, moderate-severe malnutrition group was 10.8%, 11.0%, and 27.2%, respectively (p < 0.01). Compared with moderate-severe malnutrition group, the normal group and the mild malnutrition group showed significant lower risk of CA-AKI in models adjusting for risk factors for CA-AKI and variables in univariate analysis (odds ratio [OR] = 0.48, 95% confidence interval [CI]: 0.26–0.89, p = 0.02; OR = 0.46, 95%CI: 0.26–0.82, p = 0.009, respectively). Furthermore, the relationship were consistent across the subgroups classified by risk factors for CA-AKI except anemia. The risk of CA-AKI related with CONUT score was stronger in patients with anemia. (overall interaction p by CONUT score = 0.012).

Conclusion Moderate-severe malnutrition is associated with higher risk of CA-AKI in elderly patients undergoing PCI.

**Keywords** Malnutrition · Controlling Nutritional Status (CONUT) score · Contrast-associated acute kidney injury · Percutaneous coronary intervention · Elderly

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# Introduction

With the development of coronary angiography, more and more elderly patients with coronary heart disease receive percutaneous coronary intervention (PCI). However, elderly patients with coronary diseases who underwent percutaneous coronary intervention (PCI), were more likely to develop contrast-associated acute kidney injury (CA-AKI) than general population [1]. CA-AKI is a relatively common complication after intravascular contrast media administration, which significantly prolongs days for hospitalization, increases risk of mortality and morbidity [2]. Since therapeutic strategies for CA-AKI are limited, early screening of this high-risk population and implement preventive measurement are particularly important.

Malnutrition, which is high present in elderly patients [3, 4], is also a predisposing factor for AKI [5, 6]. The

Controlling Nutritional Status (CONUT) score, an objective and comprehensive tool for nutrition assessment, is calculated from the serum albumin value, the total cholesterol level, and the total lymphocyte count [7]. The prognostic value of the CONUT score has been proved in patients with coronary artery disease [8, 9]. However, previous studies emphasized on the outcomes of mortality and adverse cardiovascular events. The role of the CONUT score in CA-AKI, one of the adverse outcomes after PCI, has not been investigated in elderly patients undergoing PCI.

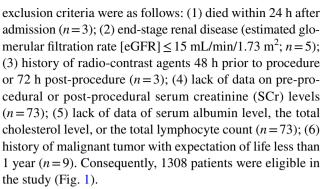
It has been demonstrated that each component of the CONUT score is related to kidney injury [5, 10, 11]. Therefore, we hypothesized that the CONUT score was associated with the incidence of CA-AKI in elderly patients undergoing PCI. We aim to evaluate the predictive value of the CONUT score for CA-AKI.

# Methods

# **Study population**

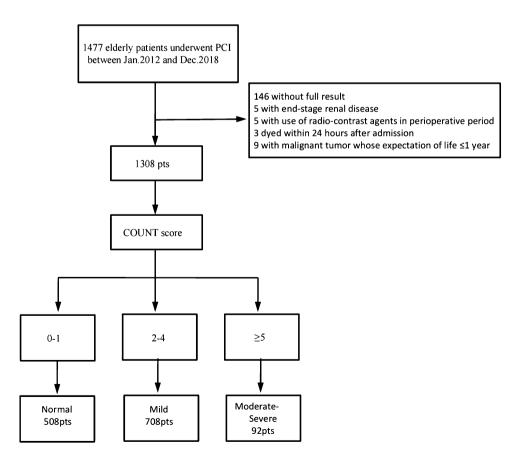
We conducted a retrospective cohort study at the Fujian Provincial Hospital, Fujian Cardiovascular Institute, between January 2012 and December 2018. A total of 1477 elderly patients aged  $\geq$  75 years undergoing PCI were enrolled. The

## Fig. 1 Study population



# Protocol

Data collected included basic characteristics [e.g. age, hypertension, diabetes mellitus (DM)] and procedure related data (e.g. number of diseased vessels). The serum albumin level, total cholesterol level, and the lymphocyte count was measured for each patient at admission. SCr was measured at admission and daily for 3 days after contrast exposure. We also measured white blood cell (WBC) count, hemo-globin (HGB), hematocrit (HCT) and other standard clinical parameters on the morning of the first or second day after admission. The eGFR was calculated using the modified modification of diet in renal disease equation [12]: eGFR =  $175 \times (SCr/88.4)^{-1.554} \times age^{-0.203} \times 0.742$  (if female) × 1.212



(if black). The use of medication was determined by the cardiologists according to clinical protocols based on guidelines. Left ventricular ejection fraction (LVEF) was evaluated using echo-cardiography during hospitalization. PCI was performed by experienced interventional cardiologists. All patients received nonionic contrast media (either Iopamiron or ultravist, both 370 mg/mL). In addition, all patients received 0.9% normal saline (NS) at a rate of 1 mL/kg/h for 12 h during perioperative period(or 0.5 mL/kg/h for 12 h if patients had overt heart failure) [13]. The protocol met the requirements of the Declaration of Helsinki and was approved by the ethics committee of the Fujian Provincial Hospital, China (ethics approval number: K2012-01-011).

## **Nutrition status evaluation**

Nutritional status was evaluated based on the CONUT score, which includes values of serum albumin, total cholesterol level, and total lymphocyte count, with a score ranging from 0 to 12 (Table S1) [7]. Higher scores indicates a worse nutritional status. Based on the CONUT score, patients were divided into normal (0–1), mild malnutrition (2–4), moderate-severe ( $\geq$ 5) malnutrition groups. The classification of nutritional status has also been applied in other studies [14, 15].

#### **Definitions and end points**

The primary end point was the development of CA-AKI, defined as an absolute increase in  $\geq 0.3$  mg/dL or  $\geq 50\%$  from the baseline SCr levels within 48 h after exposure to contrast media (CM) [16, 17]. Anemia was defined as HCT < 0.39 (for males) or, < 0.36 (for females). The diagnosis of myocardial infarction (MI) is detection of cardiac troponin I (cTnI) above the 99th percentile upper reference limit (URL) and with one of the following: symptoms of ischaemia; new or presumed new significant ST-segment-T wave (ST-T) changes or new left bundle branch block (LBBB); development of pathological Q waves in the ECG; identification of an intracoronary thrombus by angiography [18].

#### Statistical analyses

All statistical analyses were performed using R version 4.0.2. The baseline characteristics were compared among three groups divided by CONUT score. Normally distributed continuous variables were expressed as mean ± standard deviation (SD). The Student's *t*-test, Wilcoxon rank sum test was performed to determine the differences between CA-AKI and Non CA-AKI groups. And one way-analysis of variance was performed to determine the differences between groups classified by CONUT score. The categorical variables were

represented as percentages and analyzed using chi-square test or Fisher's exact test.

After testing for proportional odds assumptions, multivariate logistic analysis was used to examine the association of CONUT score 0-1 and CONUT score 2-4 (vs. CONUT score > 5) with CA-AKI in models adjusted as follows: model 1 adjusted for traditional risk factors for CA-AKI (age, anemia, DM, contrast media(CM) volume > 200 ml, eGFR < 60 mL/min/1.73m<sup>2</sup>); and model 2 adjusted for variables in model 1 plus the variables with p value < 0.05 in the univariate statistical results including AF, emergency PCI, MI, perioperative hypotension, WBC, blood glucose, hyperuricemia (HUA). Interactions between the primary end point and prespecified subgroups stratified by several CA-AKI risk factors were assessed using a likelihood ratio test for interaction. The p values for interaction were calculated in each subgroup. A 2-sided p value < 0.05 was considered as statistically significant.

## Result

## **Baseline characteristics**

A total of 1308 elderly patients were included in this study. Baseline characteristics are listed in Table 1. By CONUT calculations, 508 (38.8%) patients were not malnourished, 708 (54.1%) patients had mild malnutrition, 92 (7.0%) patients had moderate-severe malnutrition, respectively. Patients with moderate-severe malnutrition were more likely to have MI, emergency PCI, anemia, hyperuricemia. They also had lower level of WBC, lymphocyte count, serum albumin, HCT, total cholesterol, but higher level of glucose, higher percentage of eGFR < 60 mL/min/1.73 m<sup>2</sup> (all p < 0.05).

And baseline characteristics between CA-AKI group and non CA-AKI group are presented in Table 2. Based on the CONUT score, 25 (15.8%) patients in CA-AKI group and 67 (5.8%) patients in non CA-AKI group had moderatesevere malnutrition, respectively. Patients who developed CA-AKI were more likely to have MI, emergency PCI, DM, AF, HUA, as well as higher level of WBC, blood glucose, lower level of serum albumin and lymphocyte count. More patients in the CA-AKI group were more likely to be treated with contrast volume of  $\geq$  200 mL (all p < 0.05).

# **Risk factors of CA-AKI**

The incidence of CA-AKI was 10.83%, 11.02%, 27.17%, from the group of CONUT 0–1 to the group of CONUT  $\geq$  5, respectively (p < 0.001) (Table 3). After adjusting for traditional risk factors for CA-AKI, such as age, anemia, DM, CM > 200 ml, eGFR < 60 mL/min/1.73

#### Table 1 Baseline characteristics of patients in different groups classified by CONUT score

Variables	Normal	Mild	Moderate-severe	p Value	
	(CONUT 0–1) N = 508	(CONUT 2–4) N = 708	$(\text{CONUT} \ge 5)$ $N = 92$		
	11-300	11 - 700	11-72		
Demographics	79 90 2 24	70.00.2.55	90 66 1 19	< 0.001	
Age, years	78.89 3.34	79.09 3.55	80.66 4.48	< 0.001	
Sex, female, <i>n</i> (%)	194 (38.19%)	152 (21.47%)	20 (21.74%)	< 0.001	
Systolic blood pressure, mmHg	138.24 23.40	134.44 22.05	127.30 20.62	< 0.001	
Diastolic blood pressure, mmHg	72.27 12.43	71.55 23.58	67.94 12.71	0.155	
Hypertension, $n$ (%)	377 (74.2%)	552 (78.0%)	74 (80.4%)	0.211	
Diabetes, $n$ (%)	175 (34.5%)	260 (36.7%)	37 (40.2%)	0.498	
Smoker, $n$ (%)		0 (30.2%) 229 (36.2%) 27 (32		0.114	
Atrial fibrillation, $n$ (%)	62 (12.2%)	102 (14.4%)	12 (13.0%)	0.536	
Emergency PCI, <i>n</i> (%)	68 (13.4%)	109 (15.4%)	25 (27.2%)	0.003	
Myocardial infarction, <i>n</i> (%)	191 (37.6%)	304 (42.9%)	67 (72.8%)	< 0.001	
Perioperative hypotension, <i>n</i> (%)	45 (8.9%)	76 (10.7%)	18 (19.6%)	0.009	
Laboratory measurements					
Serum creatinine, µmol/L	84.46 28.86	89.69 30.60	108.09 85.05	< 0.001	
WBC, $\times 10^{9}$ /L	7.74 2.44	7.39 2.66	7.78 3.45	0.049	
lymphocyte, $\times 10^{9}$ /L	2.04 0.58	1.51 0.58	1.02 0.40	< 0.001	
HGB, g/L	132.1514.92	129.34 16.13	117.07 20.98	< 0.001	
НСТ	0.39 0.04	0.38 0.04	0.35 0.06	< 0.001	
ALB, g/L	40.73 3.33	39.46 4.04	33.27 4.36	< 0.001	
Cholesterol, mg/dL	189.88 39.51	43.70 32.91	121.97 27.93	< 0.001	
Glucose, mmol/L	7.12 2.97	7.23 3.08	9.02 4.82	< 0.001	
Uric acid, µmol/L	380.68 102.36	377.61 108.65	354.82 121.18	0.103	
eGFR, mL/min/1.73 m <sup>2</sup>	80.37 24.70	78.93 25.36	70.17 25.31	0.002	
eGFR < 60 mL/min/1.73 m <sup>2</sup> , $n$ (%)	100 (19.7%)	156 (22.0%)	31 (33.7%)	0.011	
Urine PH	6.27 0.71	6.29 0.74	6.25 0.72	0.885	
Anemia, $n$ (%)	181 (35.6%)	331 (46.8%)	65 (70.7%)	< 0.001	
Hyperuricemia, <i>n</i> (%)	203 (40.0%)	243 (34.3%)	26 (28.3%)	0.035	
Medical therapy during hospitalization					
Statin, <i>n</i> (%)	498 (98.0%)	699 (98.7%)	91 (98.9%)	0.637	
ACEI/ARB, n (%)	405 (79.7%)	586 (82.8%)	71 (77.2%)	0.242	
Antiplatelet, n (%)	498 (98.0%)	695 (98.2%)	89 (96.7%)	0.598	
Metformin, n (%)	73 (14.4%)	87 (12.3%)	12 (11.0%)	0.539	
Procedure characteristic					
Multi-vessel coronary artery disease, n (%)	403 (79.3%)	569 (80.4%)	78 (84.8%)	0.479	
Number of diseased vessels, $n$ (%)	$2.32 \pm 0.82$	$2.38 \pm 0.82$	$2.45 \pm 0.75$	0.254	
Number of stents, $n$ (%)	$1.59 \pm 0.75$	$1.68 \pm 0.80$	$1.65 \pm 0.74$	0.160	
Iso-osmolar contrast media use, n (%)	172 (33.9%)	267 (37.7%)	34 (37.0%)	0.381	
Volume of contrast media, mL	177.07 52.39	177.70 57.76	179.78 54.41	0.909	
Contrast volume > 200 mL, $n$ (%)	88 (17.3%)	128 (18.1%)	19 (20.7%)	0.741	

WBC white blood cell, HGB hemoglobin, HCT hematocrit, ALB albumin, eGFR estimated glomerular filtration rate, PCI percutaneous coronary intervention, ACEI angiotensin-converting enzyme inhibitor, ARB angiotensin receptor blocker

m<sup>2</sup>, multiple logistic regression analysis confirmed that the group of CONUT score 0–1 and the group of CONUT score 2–4 were associated with a lower risk of CA-AKI after PCI, compared with the group of CONUT score  $\geq$  5 (odds ratio[OR] 0.40, 95% confidence interval [CI] 0.23–0.72, p = 0.002; OR 0.40, 95%CI 0.23–0.70, p = 0.001) (Table 3). After adjusting for variables in model 1 plus the other variables including AF, emergency PCI, MI, perioperative hypotension, WBC, blood glucose, hyperuricemia(HUA), the group of CONUT score 0–1

Table 2Baseline characteristicsbetween non CA-AKI groupand CA-AKI group

	Non CA-AKI	CA-AKI	p Value	
	N=1150	N=158		
Demographics				
Age, years	79.0 53.49	79.70 4.06	0.121	
Sex, female, $n$ (%)	311 (27.0%)	55 (34.8%)		
Systolic blood pressure, mmHg	135.96 22.21	131.64 25.47	0.078	
Diastolic blood pressure, mmHg	71.66 19.93	71.03 13.88	0.723	
Hypertension, n (%)	877 (76.3%)	126 (79.8%)	0.384	
Emergency PCI, $n(\%)$	153 (13.3%)	49 (31.0%)	< 0.001	
Diabetes, n (%)	590 (51.3%)	109 (69.0%)	< 0.001	
Atrial fibrillation, n (%)	142 (12.4%)	34 (21.5%)	0.002	
Smoker, <i>n</i> (%)	352 (34.0%)	44 (30.4%)	0.430	
MI, <i>n</i> (%)	455 (39.6%)	107 (67.7%)	< 0.001	
Perioperative hypotension, $n$ (%)	96 (8.3%)	43 (27.2%)	< 0.001	
Medical therapy during hospitalization				
Statin, <i>n</i> (%)	1132 (98.4%)	156 (98.7%)	1.000	
Antiplatelet agents, $n$ (%)	1129 (98.3%)	153 (96.8%)	0.231	
ACEI/ARB, <i>n</i> (%)	931 (81.0%)	123 (77.9%)	0.413	
Metformin, n (%)	150 (13.0%)	21 (13.3%)	1.000	
Laboratory measurements				
Serum creatinine, µmol/L	88.09 29.88	95.27 69.28	0.597	
WBC, $\times 10^{9}$ /L	7.39 2.48	8.72 3.42	< 0.001	
Lymphocyte, $\times 10^{9}$ /L	1.70 0.64	1.54 0.69	0.005	
HGB, g/L	129.96 16.06	126.71 19.03	0.099	
НСТ	0.38 0.05	0.37 0.05	0.073	
ALB, g/L	39.78 4.15	37.57 4.21	< 0.001	
Cholesterol, mg/dL	159.85 43.50	161.98 38.06	0.187	
Glucose, mmol/L	7.17 3.07	8.38 4.01	< 0.001	
eGFR, mL/min/1.73 m <sup>2</sup>	78.93 23.75	78.47 34.06	0.247	
eGFR < 60 mL/min/1.73 m <sup>2</sup> , n (%)	242(21.0%)	45 (28.5%)	0.044	
Urine PH	6.29 0.72	6.22 0.80	0.090	
CONUT score			< 0.001	
0–1, <i>n</i> (%)	453 (39.4%)	55 (34.8%)		
2–4, <i>n</i> (%)	630 (54.8%)	78 (49.4%)		
≥5, <i>n</i> (%)	67 (5.8%)	25 (15.8%)		
Anemia, $n$ (%)	267 (23.2%)	48 (30.4%)	0.061	
Hyperuricemia, n(%)	395 (34.4%)	77 (48.7%)	0.001	
Procedure characteristic				
Contrast volume, mL	176.14 55.91	188.23 51.01	0.006	
Contrast volume $\geq$ 200 mL, <i>n</i> (%)	190 (16.5%)	45 (28.5%)	0.003	
Iso-osmolar contrast media use, $n$ (%)	416 (36.2%)	57 (36.1%)	1.000	
Number of stents, $n$ (%)	1.65 0.78	1.58 0.73	0.388	
Multi-vessel coronary artery disease, $n$ (%)	915 (79.57%)	135 (85.44%)	0.102	

*MI* myocardial infarction, *PCI* percutaneous coronary intervention, *ACEI* angiotensin-converting enzyme inhibitor, *ARB* angiotensin receptor blocker, *eGFR* estimated glomerular filtration rate, *ALB* albumin, *WBC* white blood cell, *HGB* hemoglobin, *HCT* hematocrit

and the group of CONUT score 2–4 remained significant lower risk of CA-AKI in elderly patients after PCI, compared with the group of CONUT score  $\geq$  5 (OR 0.48, 95%CI 0.26–0.89, p = 0.02; OR 0.46, 95%CI 0.26–0.82, p = 0.009) (Table 3). In other words, CONUT score  $\geq$  5 was independently associated with a higher risk of CA-AKI, compared with CONUT score 0–1 and CONUT score 2–4.

The effects of the CONUT score on the rate of CA-AKI were consistent across the prespecified subgroups (HUA,

	Participants, n	Events, n	Rate,%	Model 1 <sup>*</sup> OR (95%CI)	p Value	Model 2 <sup>†</sup> OR (95%CI)	p Value
CONUT score 0–1	508	55	10.83	0.40 (0.23-0.72)	0.002	0.48 (0.26–0.89)	0.02
CONUT score 2-4	708	78	11.02	0.40 (0.23-0.70)	0.001	0.46 (0.26-0.82)	0.009
CONUT score $\geq 5$	92	25	27.17	Reference	-	Reference	-

Table 3 Associations between CONUT score and CA-AKI

<sup>\*</sup>Model 1 adjusted for age, anemia, diabetes, contrast media volume > 200 ml, eGFR < 60 ml/(min  $\cdot 1.73 \text{ m}^2$ ). <sup>†</sup>Model 2 adjusted for variables in model 1 plus atrial fibrillation, emergency PCI, MI, WBC, glucose, hyperuricemia, perioperative hypotension. *CI* confidence interval, *OR* odds ratio

DM, CM, eGFR, AF, emergency PCI, MI) (Fig. 2). However, there was an modification by anemia: the risk of CA-AKI related with CONUT score was stronger in patients with anemia than in those without anemia (overall interaction *p* by CONUT score = 0.012) (Figs. 2, 3).

# Discussion

To our knowledge, this is the first study to demonstrate the relationship between the objective nutrition scoring tool "CONUT" and the incidence of CA-AKI in elderly patients undergoing PCI. Our results show that the CONUT score  $\geq$  5, recognized as moderate or severe malnutrition, is associated with an increased risk of CA-AKI in elderly patients undergoing PCI. Moreover, the association is stronger in elderly patients with anemia.

In our study, the incidence of CA-AKI in elderly patients undergoing PCI was up to 12.1%, which was almost consistent with the data available in a meta-analysis [19]. Age over 75 years was widely recognized as an independent risk factor of CA-AKI in patients after PCI [1]. It's well-known that comorbidities are high present in the admitted elderly patients [20], such as the age-related decrease in kidney function [21] and anemia [22], which were identified as risk factors of CA-AKI [1, 23]. Due to multiple chronic diseases, the elderly patients often take multiple oral drugs, which may worsen kidney function. Moreover, as poor vascular and heart condition, adequate hydration is difficult to achieve and thus nephrotoxic contrast cannot be discharged as soon as possible. Therefore, the elderly is more vulnerable. It's important to effectively further identify elderly patients at higher risk for CA-AKI and implement precise prevention.

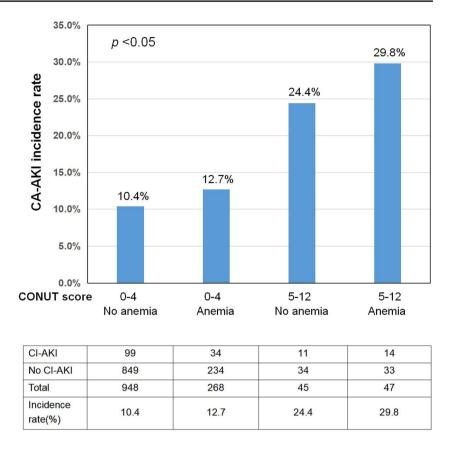
In addition to the above factors, malnutrition is high present in elderly patients [3], which is also associated with the development of kidney injury [24]. However, the best tool to identify patients at high risk of malnutrition is still in dispute. Single nutrition indicators are often affected by many factors [25]. Subjective comprehensive nutritional scoring systems like Subjective Global Assessment (SGA), are sophisticated, which need the assistance of specialized nutritionists. But there are some objective comprehensive nutritional scoring systems such as the CONUT score, Prognostic Nutritional Index (PNI), which only require simple blood biomarkers and make it convenient to apply in clinic. The CONUT score is calculated from the serum albumin value, the total cholesterol level, and the total lymphocyte count [7], which only included one more indicator than PNI. The CONUT score has been used for assessing the prognosis of heart failure [15], acute ischemic stroke [26], and a variety of malignant tumors [27, 28]. Recently, the prognostic value of the CONUT score has been validated and shown better than PNI in patients with coronary artery disease. In an observational, retrospective study of 3118 cohort, patients undergoing PCI were divided into four groups by their CONUT score (0-1 vs. 2-4 vs. 5-8 vs. 9-12). It revealed that patients with higher CONUT scores had higher rates of major adverse cardiac events (hazard ratio [HR]: 1.14; 95% CI: 1.07–1.22, p < 0.05 [8]. Basta et al. evaluated the CONUT and PNI score in 945 patients with ST-elevation myocardial infarction undergoing PCI and found that patients with severe CONUT but not patients with severe PNI index had the highest rate for all-cause death, with a log-rank of p < 0.001[9]. Roubín et al. made a similar conclusion that the CONUT score has a higher sensitivity than the PNI for all-cause death and major adverse cardiovascular events (MACEs) in patients with acute coronary syndrome [29]. Therefore, it was appropriate to apply the CONUT score in our study. However, these studies emphasized on the outcomes such as mortality and adverse cardiovascular events in patients after PCI. The role of the CONUT score in CA-AKI, which is one of the adverse outcomes of PCI, has not been investigated in elderly patients.

Our study first revealed that elderly patients with moderate-severe malnutrition (CONUT score  $\geq$  5) had higher risk of CA-AKI. Even after adjusting for multiple risk factors, moderate-severe malnutrition is also significantly associated with the development of CA-AKI. Our subgroup analysis also confirms the relationship between the CONUT score and the risk of CA-AKI, although there is an modification by anemia. This could probably be explained by the added risk of anemia to CA-AKI. It was recognized that anemia on admission is associated with an increased risk of CA-AKI [1]. The possible explanation may be the anemia-induced

				Interactio
Subgroup			OR(95%CI) P.value	P.value
All patients		CONUT: ≥ 5	(Ref.)	N/A
		CONUT: 2-4	► <b>■</b> 0.447(0.255-0.800) 0.006	
		CONUT: 0-1	► <b>■</b> 0.455(0.250-0.841) 0.011	
HUA	No	CONUT: ≥ 5	(Ref.)	0.377
		CONUT: 2-4	► <b>1</b> 0.506(0.247-1.080) 0.069	
		CONUT: 0-1	• <b>-</b>	
	Yes	CONUT: ≥ 5	(Ref.)	
		CONUT: 2-4	• 0.395(0.151-1.062) 0.060	
		CONUT: 0-1	► 0.306(0.113-0.843) 0.020	
DM	No	CONUT: ≥ 5	(Ref.)	0.422
		CONUT: 2-4	• 0.976(0.316-3.739) 0.969	
		CONUT: 0-1	<b>1.245(0.379-4.986)</b> 0.734	
	Yes	CONUT: ≥ 5	(Ref.)	
		CONUT: 2-4	0.353(0.180-0.698) 0.002	
		CONUT: 0-1	0.329(0.162-0.676) 0.002	
СМ	<200ml	CONUT: ≥ 5	(Ref.)	0.201
		CONUT: 2-4	0.589(0.298-1.215) 0.137	
		CONUT: 0-1	0.503(0.242-1.082) 0.071	
	>=200ml	CONUT: ≥ 5	(Ref.)	
		CONUT: 2-4	← ● 0.242(0.073-0.782) 0.018	
		CONUT: 0-1	← ■ 1 0.324(0.094-1.089) 0.069	
anemia	No	CONUT: ≥ 5	(Ref.)	0.012
anemia	NO	CONUT: 2-4		0.012
		CONUT: 0-1	0.432(0.189-1.043) 0.053	
	Yes	CONUT: ≥ 5		
	165	CONUT: 2-4	(Ref.) 0.256(0.105-0.618) 0.002	
		CONUT: 2-4 CONUT: 0-1		
	≥60			0.357
eGFR	200	CONUT: ≥ 5	(Ref.)	0.357
nl/min/1.73m2)		CONUT: 2-4		
	-00	CONUT: 0-1	0.406(0.196-0.867) 0.017	
	<60	CONUT: ≥ 5	(Ref.)	
		CONUT: 2-4		
. –		CONUT: 0-1	0.641(0.219-1.940) 0.420	
AF	No	CONUT: ≥ 5	(Ref.)	0.872
		CONUT: 2-4	0.460(0.248-0.876) 0.016	
		CONUT: 0-1	····■ 0.467(0.242-0.920) 0.025	
	Yes	CONUT: ≥ 5	(Ref.)	
		CONUT: 2-4	← <b>0</b> .322(0.069-1.648) 0.152	
		CONUT: 0-1	← 0.358(0.070-1.984) 0.220	
mrgency PCI	No	CONUT: ≥ 5	(Ref.)	0.100
		CONUT: 2-4	·── <b>─</b> ── 0.453(0.228-0.939) 0.028	
		CONUT: 0-1	<b>└──■</b> 0.611(0.299-1.293) 0.184	
	Yes	CONUT: ≥ 5	(Ref.)	
		CONUT: 2-4	► 0.476(0.152-1.499) 0.198	
		CONUT: 0-1	← ■ 0.182(0.048-0.655) 0.010	
MI	No	CONUT: ≥ 5	(Ref.)	0.075
		CONUT: 2-4	← ■ 0.222(0.077-0.692) 0.007	
		CONUT: 0-1	0.315(0.110-0.980) 0.037	
	Yes	CONUT: ≥ 5	(Ref.)	
		CONUT: 2-4	0.589(0.304-1.172) 0.122	
		CONUT: 0-1	0.490(0.237-1.030) 0.056	

Fig. 2 Subgroup analysis of the effect of CONUT score on CA-AKI incidence in the matched cohort

Fig. 3 Incidence rate of CA-AKI in patients with various combination of CONUT score and anemia



aggravation of renal ischemia and hypoxic injury to the outer medulla in the kidney [30]. Combined with malnutrition, the capacity of scavenging oxygen free radical is reduced, which may aggravate cell toxicity and further impair the kidney function [31].

The mechanism of the connection between the CONUT score and CA-AKI has not been fully investigated. However, each component of CONUT score has been demonstrated the relationship with kidney injury. First, serum albumin levels, which are twice the weight of other indicators in CONUT score, can predict the risk of CA-AKI. A meta-analysis including about 68,000 subjects confirmed that lower serum albumin is an independent predictor both of AKI and death after AKI in patients undergoing cardiac surgery or acute coronary interventions [6]. Furthermore, a retrospective study found that in patients treated with PCI, the serum albumin level was significantly lower in the CA-AKI group (3.52 + 0.40 vs. 3.94 + 0.39 mg/dL, p < 0.001) and it was an independent predictor of CA-AKI[5]. Second, although hypercholesterolemia is a well-established risk factor for cardiovascular disease in the general population [32], the correlation between low serum cholesterol and adverse outcomes has been reported for patients with renal failure. Obialo et al. evaluated a 3-year retrospective study of patients with acute kidney injury and demonstrated that survival was higher among patients with cholesterol > 150 mg/dL than those whose levels were < or = 150 mg/dL (p < 0.001) [10].Another study also showed that lower total cholesterol (TC) levels over time were significantly associated with worse survival (HR 1.66, 95% CI 1.11-2.47) in patients with chronic kidney diseases who underwent peritoneal dialysis [33]. Finally, lymphocyte count may represent a marker of the inflammation response [34], which is a significant and crucial factor in the pathogenesis of CA-AKI [35]. A low relative lymphocyte count was shown to be independently associated with worse prognosis in patients with CAD [36]. Additionally, several studies have shown that higher ratio of neutrophil/lymphocyte is related to a greater risk of CA-AKI in patients who underwent PCI [11, 37], suggesting that a low lymphocyte count may contribute to the development of CA-AKI. Thus, the CONUT score, reflecting not only the status of malnutrition but also the degree of inflammation, may be more appropriate for early detection of elderly patients with high risk of CA-AKI.

# Limitation

First, this study was a single-center cohort with a relatively small sample size. A potential patient selection bias may be existed. Second, the CONUT score was evaluated only at the time of admission, and we did not assess the effect of the change of score during the observation period. Thirdly, elderly patients comprised major proportion of participants in our study. Therefore, our results may not be applied to younger patients. Furthermore, based on this study, it is unclear whether the stratified risk classes need an nutritional intervention.

# Conclusion

We find that moderate-severe malnutrition is strongly associated with high risk of CA-AKI in elderly patients who underwent PCI. Moreover, the risk of CA-AKI related with CONUT score was stronger in patients with anemia. Further studies are required to determine whether nutrition support improves clinical outcomes in this population.

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## Declarations

**Conflict of interest** The authors declare that there are no conflict of interests.

**Ethical approval** All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional or national research committee and with the 1964 Helsinki Declaration and its later amendments or comparable ethical standards. Approval for this study was approved by the ethics committee of the Fujian Provincial Hospital, China (ethics approval number: K2012-01-011).

**Informed consent** Informed written consents were obtained from all patients after providing them a detailed written description of the potential benefits and risks associated with the study.

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