cambridge.org/cty

## **Original Article**

**Cite this article:** Kara KA, Arslanoğlu E, Yigit F, Arkan C, Yılmaz AA, Akardere ÖF, Cine N, Tuncer E, Çetiner N, and Ceyran H (2024) The effect of albumın and total protein values as nutritional markers after Fontan surgery on prognosis. *Cardiology in the Young* **34**: 734–739. doi: 10.1017/S1047951123003335

Received: 27 July 2023 Revised: 15 August 2023 Accepted: 16 August 2023 First published online: 4 October 2023

**Keywords:** 

Fontan; albumin; nutrition; total protein; protein-losing enteropathy

**Corresponding author:** Kenan Abdurrahman Kara; Email: kenankaradoc@gmail.com

© The Author(s), 2023. Published by Cambridge University Press.



# The effect of albumin and total protein values as nutritional markers after Fontan surgery on prognosis

Kenan Abdurrahman Kara<sup>1</sup>, Ergin Arslanoğlu<sup>1</sup>, Fatih Yigit<sup>1</sup>, Cuneyt Arkan<sup>1</sup>, Abdullah Arif Yılmaz<sup>1</sup>, Ömer Faruk Akardere<sup>1</sup>, Nihat Cine<sup>1</sup>, Eylem Tuncer<sup>1</sup>, Nilüfer Çetiner<sup>2</sup> and Hakan Ceyran<sup>1</sup>

<sup>1</sup>Department of Pediatric Cardiac Surgery, Kartal Kosuyolu Training And Research Hospital, İstanbul, Türkiye and <sup>2</sup>Department of Pediatric Cardiology, Kartal Kosuyolu Training And Research Hospital, İstanbul, Türkiye

## Abstract

Introduction: Children with CHD develop heart failure due to increased pulmonary blood flow, cyanosis, and pulmonary hypertension. The metabolic needs of these children differ from those of healthy children, and malnutrition is common. Protein-losing enteropathy has been reported in 5 to 13% of patients after the Fontan procedure. Serum albumin and total protein levels, which are indicators of the quality of post-operative care, can be useful tools for monitoring and examining the intensive care treatment strategies of these patients. In our retrospective study, the effects of albumin and total protein values, which are two of the markers that give us an idea about diet, nutritional status, and inflammation, on the prognosis of children who underwent the Fontan procedure were investigated. Method: In our study, 127 patients who underwent Fontan procedure in our clinic between 2012 and 2021 were analysed retrospectively. Of the patients, 52.7% (n = 67) were male and 47.3% (n = 60) were female. The mean age is  $5.83 \pm 4.63$ years. Patients who underwent albumin replacement were not included in the study. Results: Although the relationship between pre-operative albumin and total protein values and postoperative mortality was not statistically significant, the inverse correlation of post-operative albumin 1st, 2nd, and 3rd-day values and post-operative total protein 1st, 2nd, and 3rd-day values with mortality was found to be statistically significant. In addition, we found that mortality was statistically high in patients whose total protein amount was below 6.65 mg/dl in the early post-operative period. Conclusion: Albumin and total protein, whose blood levels can vary with diet, can be used as predictors in the early post-operative prognosis of Fontan patients. In addition, when we examined the exitus patients, it was observed that the total protein amount was below 6.65 mg/dl on the post-operative 1st day. Based on this, we think that a diet with high protein content before surgery will help reduce post-operative early mortality.

## Introduction

Children with CHD develop heart failure due to increased pulmonary blood flow, cyanosis, and pulmonary hypertension. As a result, the metabolic needs of these patients differ from those of healthy children. Gastrointestinal malabsorption disorders due to increased oxygen demand and low cardiac output, low-calorie intake, and increased calorie requirement cause malnutrition. The degree of CHD and the degree of malnutrition are directly proportional.<sup>1,2</sup>

Inadequate gastrointestinal absorption causes insufficient serum visceral protein synthesis and inflammation. Thus, low serum albumin, pre-albumin level, and total protein levels occur. Hypoalbuminemia causes liver dysfunction, end-stage liver disease, increased catabolism, nephrotic syndrome, and protein-losing enteropathy in both acute and chronic stages.<sup>3</sup>

Protein-losing enteropathy has been reported in 4–13% of patients undergoing the Fontan procedure. Protein-losing enteropathy is thought to occur secondary to increased systemic venous pressure. Decreased cardiac output, venous congestion, increased mesenteric vascular resistance, and intestinal inflammation may adversely affect the clinical course.<sup>4</sup> In addition, persistent peripheral oedema due to hypoalbuminemia and ascites develops in the abdomen and pleural effusions may occur in the post-operative period. In addition, recurrent infections are seen due to the decrease in immunoglobulin levels and lymphocytes. Dietary regulation, albumin replacement, and steroid therapy in the post-operative period constitute the basic elements of medical treatment.<sup>5</sup> Serum albumin level, which is an indicator of the quality of post-operative care, can be a useful tool for the follow up and examination of intensive care treatment strategies of these patients.<sup>6</sup>

The aim of this study is to investigate the effects of albumin and total protein values, which give us an idea about nutrition and inflammation, on the prognosis of the Fontan procedure.



## **Material and method**

In our study, 127 patients who underwent Fontan procedure in our clinic between 2012 and 2021 were retrospectively analysed and included in the study. We routinely use fresh frozen plasma as a cardiopulmonary bypass prime solution, and we use erythrocyte suspension in patients with a haematocrit below 27. We do not routinely use albumin in cardiopulmonary bypass. We add albumin to the prime solution in patients with albumin levels below 2.5 mg/dl. Patients who underwent albumin replacement were not included in the study. Catheter angiography is routinely performed on patients in the pre-operative period. Pre-operative pulmonary pressures and pulmonary circulation values were evaluated and noted. Alanine amino transferase aspartate amino transferase, creatinine, urea, albumin, and total protein values of the patients included in the study were recorded in the preoperative and post-operative first three days, and their effect on prognosis was evaluated.

Median sternotomy incision was used in all patients. "Fontan" circulation is completed by anastomosis between the inferior vena cava and the right pulmonary artery using a polytetrafluoroethylene graft suitable for the weight, height, and age of the patients. Cardiopulmonary bypass was entered using the superior vena cava, inferior vena cava, and aortic cannula. We do not put a cross-clamp during these anastomoses. When intracardiac procedures such as atrioventricular valve repair were required, the heart was stopped with blood, custadiol, or del nido cardioplegia solutions and the procedure was performed under cross-clamping. At the end of the procedure, patients were appropriately weaned from cardiopulmonary bypass. We routinely cut off the pulmonary antegrade flow of patients during the operation. We monitor the pulmonary pressures in the perioperative period and evaluate the appropriateness of the Fontan circulation intra-operatively.

Intracardiac Fontan procedure was performed in 11 patients and extracardiac Fontan procedure was performed in 116 patients. In the follow up of the patients, a balance between adequate diuresis and <3 mmol/L lactate level was tried to be achieved. Especially colloids have been used to provide cardiac output. Extracorporeal membrane oxygenator support was applied if the low cardiac output syndrome persisted despite volume replacement and inotropic support. Extracorporeal membrane oxygenator need, pre-operative echocardiographic data, systemic ventricular evaluation, post-operative intensive care follow up, and blood gas analysis were noted. Vasoactive-inotrope score score (Vasoactive-inotrope score) was calculated as follows; dopamine dose  $(mg/kg/min) + dobutamine dose (mg/kg/min) + 100 \times dose$ of epinephrine  $(mg/kg/min) + 100 \times dose$  of norepinephrine  $(mg/kg/min) + 10,000 \times dose of vasopressin (U/kg/min) + 10 \times$ milrinone dose (mg/kg/min).

Fenestration was opened in patients who were expected to have high pulmonary pressure and who were found to have high pulmonary pressure in intra-operative measurements. Fenestration was created between the right atrium and the polytetrafluoroethylene graft. A four-millimetre punch was used to open a fenestration between the graft and the right atrium.

Number Cruncher Statistical System 2007 (Kaysville, Utah, USA) programme was used for statistical analysis. While evaluating the study data, descriptive statistical methods (mean, standard deviation, median, frequency, ratio, minimum, maximum) as well as the distribution of the data were evaluated with the Shapiro–Wilk Test. The Mann–Whitney U Test was used to

compare the quantitative data between two groups that did not show normal distribution. Friedman test was used for comparisons of three or more periods, and Wilcoxon test was used for comparisons of two periods. Spearman's correlation analysis was used to determine the relationship between quantitative data. Roc analysis was performed to determine the predictive value of the quantitative data. Logistic regression analysis was used to determine the independent variables affecting the dependent variable. Significance was evaluated at p < 0.01 and p < 0.05 levels.

The study was carried out retrospectively, in accordance with the Declaration of Helsinki, with the permission of the hospital management, taking into account the ethical rules.

## Results

In total, 52.7% (n = 67) of the patients included in the study were male. In terms of systemic ventricle, 74% (n = 94) of the patients were left-sided and 26% (n = 33) were right-sided. Extracorporeal membrane oxygenator was used in 14.1% (n = 18) of the patients, while extracorporeal membrane oxygenator was not required in 85.9% (n = 109) of the patients. While 34.7% (n = 26) of the patients died, 65.3% (n = 49) of the patients survived. While fenestration was performed in 29.1% (n = 37) of the patients, fenestration was not performed in 70.9% (n = 90) of the patients.

Patients' age (years), operative time (min), weight (kilograms), pre-operative  $SpO_2$  (%), post-operative  $SpO_2$ , vasoactive-inotrope score Score, po 24 hours highest BP (mmHG), po 24 hours max lactate level (mmol/dL), po 24 Hours highest central venous pressure (mmHG), ICU stay (days), service hospitalisation time (days), total hospital stay (days), ventilation time (hours), pre-op pulmonary pressure (mmHg), total bypass time (minutes) ), and aortic cross-clamp time (minutes) values are given in the table (Table 1).

Table 2 compares the exitus and living Fontan patients; 24-hours max lactate, ICU hospitalisation time, service hospitalisation time, ventilation time, pre-op pulmonary pressure, total bypass time, aortic cross-clamp time time, Albumin Postop-1. day, Albumin Postop-2. day, and Albumin Postop-3. day values were found to be statistically significant. Pre-op albumin values were statistically insignificant (Table 2).

There are statistically significant changes in albumin, total protein, and creatinine levels on post-operative days (p = 0.001; p < 0.01) (Table 3).

There is a statistically significant difference in alanine amino transferase and aspartate amino transferase levels on postoperative days. (p = 0.001; p < 0.01). It was found statistically significant that the pre-operative alanine amino transferase and aspartate amino transferase values were lower than the other periods (p = 0.001; p < 0.01). It was found statistically significant that the alanine amino transferase and aspartate amino transferase values on the post-operative 1st day were lower than the post-operative 2nd day and post-operative 3rd day (p = 0.001; p < 0.01).

When the lower limit of total protein was taken as 6.65, the sensitivity was determined as 92.3% and the specificity as 81.6% as the reliable cut-off point (Fig. 1).

In the correlation analysis, there is a positive correlation between length of stay in ICU and ventilation time, pre-operative pulmonary artery pressure values, cardiopulmonary bypass time, and aortic cross-clamp time. Correlation coefficients were 0.887– 0.540– 0.554– 0.443, respectively (p < 0.01). There is a positive correlation between ventilation time and pre-operative pulmonary

Та	ble	1.	Averages	of	demograp	hic d	lata
----	-----	----	----------	----	----------	-------	------

	Mean ± Sd	Min–Max (Median)
Age (years)	5,83 ± 4,63	2–23 (4)
Operation time (min)	312,56 ± 97,47	100–580 (310)
Weight (kilogram)	20,17 ± 13,51	8,4–70 (15)
Pre-operative SpO <sub>2</sub> (%)	81,6 ± 4,83	70–91 (81)
Post-operative SpO <sub>2</sub> (%)	89,69 ± 3,67	83–97 (89)
VIS score	13,33 ± 4,1	0-22 (14)
Po 24 hour peak systolic bp (mmHG)	140,71 ± 27,06	90–200 (140)
Po 24 hours max lactate (mmol/dL)	11,6 ± 3,82	4–21 (11)
Po 24 hours highest CVP (mmHG)	18,17 ± 3,62	7–28 (18)
ICU duration (days)	15,73 ± 13,53	3-64 (10)
Service time (days)	10,13 ± 9,69	0-49 (10)
Total length of stay in hospital (days)	25,87 ± 14,37	4–89 (22)
Ventilation time (hours)	10,55 ± 13,91	1-64 (3)
Pre-op pulmonary pressure (mmHg)	12,04 ± 3,7	4-20 (11)
Total bypass (minutes)	118,97 ± 41,27	72–280 (104)
ACC (minutes)	19,24 ± 34,59	0-140 (0)

Po=post-operative; Bp=Blood pressure; ACC=Aortic cross-clamp time; CVP=central venous pressure; VIS=vasoactive-inotrope score.

artery pressure values, and cardiopulmonary bypass time and aortic cross-clamp time. Correlation coefficients were 0.624-0.494-0.393, respectively (p < 0.01).

There was an inverse and weakly significant correlation between the length of stay in the ICU and albumin on the 2nd post-operative day (r = -0.351, p < 0.01). There was an inverse and weakly significant correlation between the maximum lactate level in the first 24 hours post-operatively and the albumin level on the post-operative 1st day (r = -0.324, p < 0.01). There is an inverse and very weak significant correlation between total length of stay in hospital and post-operative 2nd day albumin (r = -0.248, p < 0.05). There was no statistically significant difference between albumin values and the amount of drainage in the first three post-operative days (p > 0.05).

#### Discussion

After Fontan surgery, patients are most at risk of ventricular dysfunction, bronchitis, protein-losing enteropathy, and chronic Fontan failure. When we look at protein-losing enteropathy, we see that the levels of albumin and total protein will have an effect on the prognosis. Pre-operative risk stratification is important and guiding in the follow up of these patients.

Hypoalbuminemia is common in children with paediatric heart disease. Surgical stress is manifested by a decrease in serum albumin concentrations. It has been stated that serum albumin concentration may be associated with the risk of post-operative infection. Hypoalbuminemia is an independent risk factor for patients undergoing paediatric heart surgery.<sup>7,8</sup> In our study, a weak inverse correlation was found between the albumin values on

the post-operative 1st day and the post-operative 24 hour maximum lactate values. This situation was interpreted in favour of the positive effect of albumin on peripheral tissue perfusion.

The effect of nutritional parameters on prognosis in patients undergoing paediatric heart surgery was investigated, but no significant relationship was found. When evaluated together with anthropometric parameters, a link between malnutrition and post-operative morbidity has not been established yet. However, albumin values used to determine the degree of malnutrition have been found to affect post-operative prognosis and mortality.<sup>9,10</sup> In our study, it was found statistically significant that post-operative low albumin values were associated with mortality. The post-operative 1st, 2nd, and 3rd-day values of albumin were found to be higher in the living group compared to the exitus group (p(po1) = 0.01; p(po2) = 0.005 p(po3) = 0.015 p < 0.05). We also believe that post-operative albumin values also have an effect on mortality.

Berbel-Franko et al. stated that the degree of hypoalbuminemia after open heart surgery has negative effects on the development of post-operative complications (especially septic or bleeding complications) and long-term mortality, and they stated that the development of hypoalbuminemia may be due to nutritional and inflammatory factors.<sup>6</sup> In our study, an inverse and weakly significant correlation was found between the length of hospital stay and the albumin value on the second post-operative day. It was concluded that low albumin values affect the length of hospital stay and hospital costs.

Geukers et al. investigated the consequences of normal and excess protein intake after cardiac surgery. They argued that protein intake over 5 g/kg/day in the post-operative period may have side effects such as metabolic acidosis, but although they could not obtain statistically significant results, they argued that normoprotein 2 g/kg/day nutrition in the early period is more appropriate.<sup>11</sup> When the albumin values on the pre-operative and post-operative 1st, 2nd, and 3rd days were compared on a daily basis, it was determined that the albumin values showed a significant decrease in the post-operative period. Considering the effect of albumin values within normal limits on post-operative prognosis, it has been interpreted that the post-operative prognosis of patients whose pre-operative albumin levels are not lower than normal will be better than patients with low pre-operative albumin levels. We did not perform pre-op albumin replacement on the patients in our study group. In addition, we excluded four neonatal hypoplastic patients whose pre-op albümin level was below 2.5 mg/ dl and who had pre-operative albumin replacement. Thus, we think that we cannot directly affect the course of post-operative albumin. The prognostic nutritional index is used as an indicator of the nutritional and immune status of cancer patients calculated according to serum albumin concentration and peripheral blood lymphocyte count. Measurements made with this index have yielded significant results in many patient populations, including heart failure patients undergoing left ventricular assist device implantation, as well as those undergoing gastrointestinal cancer surgery.<sup>12,13</sup> When we investigated the effect of albumin and total protein on the prognosis in our own patient group, we could not obtain a fixed cut-off value for albumin, but when the total protein value was above 6.65 (sensitivity 92.3%; specificity 81.6%), there was a significant decrease in mortality.

It should be recognised that in children requiring surgery for CHD, clinicians and surgeons should carefully consider the child's nutritional status when planning non-emergency surgery for CHD and counselling parents about the potential risks of surgery. Longer

## Table 2. Comparison of parameters by living status

		Ν	Mean ± Sd	Min–Max (Median)	р
Age (Month)	Exitus	19	7,27 ± 6,27	2–23 (4,5)	0,424
	Alive	108	5,06 ± 3,29	2–17 (4)	
Operation time (min)	Exitus	19	353,54 ± 97,05	142–580 (345)	0,014*
	Alive	108	290,82 ± 91,38	0–480 (300)	
Weight (kg)	Exitus	19	24,52 ± 17,67	8,5–70 (16)	0,114
	Alive	108	17,87 ± 10,15	8,4–65 (14)	
Pre-operative SpO <sub>2</sub> (%)	Exitus	19	83,04 ± 4,49	74–91 (84)	0,086
	Alive	108	80,84 ± 4,88	70–91 (81)	
Post-operative SpO <sub>2</sub> (%)	Exitus	19	89,85 ± 3,84	83–97 (89)	0,840
	Alive	108	89,61 ± 3,6	84–97 (90)	
Vis score	Exitus	19	14,77 ± 3,8	8–22 (14,5)	0,058
	Alive	108	12,57 ± 4,09	0-22 (13)	
24 hour peak BP (mmHG)	Exitus	19	141,62 ± 33,53	90–200 (135)	0,880
	Alive	108	140,22 ± 23,3	98–190 (140)	
24 hours max lactate	Exitus	19	13,65 ± 3,66	6-21 (14)	0,001**
	Alive	108	10,51 ± 3,47	4–19 (9)	
24 hours highest CVP (mmHG)	Exitus	19	18,73 ± 2,86	13–24 (18,5)	0,308
	Alive	108	17,88 ± 3,96	7–28 (18)	
ICU length of stay (days)	Exitus	19	26,73 ± 15,33	4–64 (23,5)	0,001**
	Alive	108	9,9±7,71	3–40 (7)	
Post-operative service time (days)	Exitus	19	0 ± 0	0–0 (0)	0,001**
	Alive	108	15,51 ± 7,72	6–49 (15)	
Total hospital stay (days)	Exitus	19	26,73 ± 15,33	4–64 (23,5)	0,738
	Alive	108	25,41 ± 13,98	10-89 (22)	
Ventilation time (days)	Exitus	19	25,19 ± 14,91	4–64 (23)	0,001**
	Alive	108	2,78 ± 2,14	1–10 (2)	
Pre-op pulmonary pressure (mmHg)	Exitus	19	14,31 ± 3,76	5–20 (14,5)	0,001**
	Alive	108	10,84 ± 3,08	4–18 (11)	
Total bypass (minutes)	Exitus	19	139,65 ± 43,33	84–240 (125)	0,001**
	Alive	108	108 ± 35,96	72–280 (95)	
ACC (minutes)	Exitus	19	35,38 ± 44,37	0–140 (7,5)	0,001**
	Alive	108	10,67 ± 24,56	0–105 (0)	
Albumin Pre-op (mg/dL)	Exitus	19	4,46 ± 0,52	3,5–5,63 (4,4)	0,514
	Alive	108	4,49 ± 0,36	3,7–5,2 (4,5)	
Albumin Postop-1 (mg/dL)	Exitus	19	3,3 ± 0,54	2,5–4,4 (3,27)	0,01*
	Alive	108	3,72 ± 0,67	2,4–5,2 (3,8)	
Albumin Postop-2 (mg/dL)	Exitus	19	3,39 ± 0,56	2,1–4,5 (3,37)	0,005**
	Alive	108	3,79 ± 0,57	2,5–4,8 (3,9)	
Albumin Postop-3 (mg/dL)	Exitus	19	3,39 ± 0,58	2,1–4,5 (3,35)	0,015*
	Alive	108	3,75 ± 0,57	2,6-5 (3,88)	

Mann-Whitney U Test \*p < 0,05; \*\*p < 0,01.

		Preop	Postop-1. day	Postop-2. day	Postop-3.day	р
Albumin (mg/dL)	Mean ± Sd	4,48 ± 0,42	3,58 ± 0,66	3,65 ± 0,59	3,63 ± 0,59	0,001**
	Min–Max (Median)	3,5–5,63 (4,5)	2,4–5,2 (3,5)	2,1–4,8 (3,7)	2,1–5 (3,7)	
Total protein (mg/dL)	Mean ± Sd	7,15 ± 0,71	5,62 ± 0,95	5,71 ± 0,81	5,72 ± 0,77	0,001**
	Min–Max (Median)	4,8-8,6 (7,1)	3,3–7,9 (5,5)	3,7–7,5 (5,7)	3,7–7,8 (5,8)	
Creatinin (mg/dL)	Mean ± Sd	0,38 ± 0,11	0,7 ± 0,4	0,85 ± 0,76	0,98 ± 1,02	0,001**
	Min–Max (Median)	0,19–0,82 (0,36)	0,18–2,08 (0,58)	0,22–3,8 (0,5)	0,11-4,2 (0,44)	
ALT (u/L)	Mean ± Sd	16,89 ± 7,64	108,56 ± 387,43	378,2 ± 781,71	465,21 ± 767,41 5	0,001**
	Min–Max (Median)	6-60 (16)	5–3159 (22)	7–4586 (31)	3342 (105)	
AST (u/L)	Mean ± Sd	35,45 ± 9,99	262,79 ± 905,14	889,35 ± 1739,81	1113,4 ± 1668,68	0,001**
	Min–Max (Median)	10-69 (34)	15-7454 (79)	16–7056 (113)	11-7254 (203)	
Drainage (cc)	Mean ± Sd	-	405,47 ± 164,2	290,13 ± 109,17	206,4 ± 127,54	0,001**
	Min-Max (Median)		130-900 (380)	100-700 (270)	0-600 (200)	

Table 3. Comparison of biochemistry measurements by periods for all patients

Friedman test: \*\*p < 0,01, AST=aspartate amino transferase; ALT=alanine amino transferase.



Figure 1. Total protein Roc analysis graph.

hospital stays and higher readmission rates were observed among malnourished children who underwent surgery for CHD. It is argued that nutrition should be improved by making necessary interventions in the pre-operative period in order to improve nutrition and patient outcomes.<sup>14</sup> We evaluate all our patients together with our paediatrician in the pre-operative period, make pre-operative preparations in optimal conditions according to the patient and cardiac pathology, and inform the family.

Feeding difficulties are common in infants with CHD, often presenting with recurrent lung infection, tachypnoea, and sweating. Malnutrition causes low immunity in children with CHD and predisposes them to infections, leading to heart failure, higher rates of complications, and more frequent hospital admissions. Malnutrition and the vicious cycle of infections pose a significant problem in these children. Nutritional assessment and appropriate counselling help improve the situation.<sup>15,16</sup> We follow up with our patients in the post-operative period by calling them for routine controls.

Growth retardation is common in children with single ventricular physiology. In a study by Luo et al. on the nutritional status of children with single ventricle physiology who had undergone Glenn operation, they stated that nutritional status affected clinical results and increased the length of hospitalisation.<sup>17</sup> In our study, there was an inverse and weakly significant relationship between the length of stay in the ICU and the albumin level on the 2nd post-operative day, and we believe that nutrition is an important factor in terms of post-operative prognosis, in line with the literature.

Albumin contributes to Starling forces in capillaries. It also helps to prevent hemodilution associated with cardiopulmonary bypass and preserve erythrocyte morphology. Both of these factors affect microcirculation, and hypoalbuminemia can exacerbate tissue damage, resulting in prolonged hospital stay. However, a clear limit value has not been determined for the values at which albumin transfusion should be performed.<sup>18</sup> Although there was no clear lower limit value for albumin in our study, mortality decreased significantly above 6.65 mg/dl for total protein values in the blood.

Zaccagni et al found that albumin values lower than 5% in the early post-operative period were associated with prolonged chest tube output and prolonged hospital length of stay.<sup>19,20</sup> We attribute our finding of a statistically significant difference between albumin values and the amount of early post-operative drainage to the fact that the main cause of early drainage is bleeding and coagulation factors. We think that the relationship between albumin values and prolonged chest tube output in the late post-operative period may be significant.

Endothelial dysfunction triggered by extracorporeal circulation makes it difficult to maintain fluid homeostasis during and after cardiac surgery. The albumin solution affects the plasma oncotic pressure, which causes an increase in intravascular volume. The increased plasma volume interacts positively with tissue perfusion and the endothelial glycocalyx layer, reducing the leakage from the capillaries to the extravascular space.<sup>21,22</sup> Similarly, in our study, statistically significant results were obtained between the highest lactate value in 24 hours and the albumin value on the post-operative 1st day. Here, the positive effect of albumin on microcirculation is shown.

Rauf et al., in their study in which they compared the patients who received albumin in the cardiopulmonary bypass prime solution and those who did not, found that adding albumin to the prime solution had an effect on hemodynamic stability and preserved platelet count in the post-operative period.<sup>23</sup> Except for special patient groups with severe hypoalbuminemia, we do not prefer to routinely add albumin to the prime solution in our protocol due to high costs.

## Conclusion

Fontan is a palliative, not a corrective surgery with high morbidity and mortality, but we think that it is the best solution that can be offered to patients with functional single ventricles who do not have a known risk factor for today. It can be said that with the contribution of the developing technique, pre-operative evaluation methods and preparatory operations, late complications will be less and long-term results will be better in these patients. Albumin and total protein, the levels of which can vary with the amount and quantity of nutrition, can be used as predictors in the prognosis of these patients. Although the total protein values of 6.65 mg/dl and above in our study group decreased the post-operative mortality statistically significantly, we think that it is only one of the predictors that can affect the post-operative mortality.

## Limitations

The study is a retrospective single-centre study. Albumin and total protein levels are affected by many factors, and blood levels vary, and studies with larger populations are needed to determine more precise limits.

#### Acknowledgements. None.

**Financial support.** This research received no specific grant from any funding agency, commercial, or not-for-profit sectors.

#### Competing interests. None.

#### References

- Mehta NM, Skillman HE, Irving SY, et al. Guidelines for the provision and assessment of nutrition support therapy in the pediatric critically ill patient: society of critical care medicine and American Society for Parenteral and Enteral Nutrition. J Parenter Enteral Nutr 2017; 41: 706–742. DOI: 10.1177/ 0148607117711387.
- Varan B, Tokel K, Yilmaz G. Malnutrition and growth failure in cyanotic and acyanotic congenital heart disease with and without pulmonary hypertension. Arch Dis Child 1999; 81: 49–52. DOI: 10.1136/adc.81.1.49.
- Loftus TJ, Brown MP, Slish JH, Rosenthal MD. Serum levels of prealbumin and albumin for preoperative risk stratification. Nutr Clin Pract 2019; 34: 340–348. DOI: 10.1002/ncp.10271.
- Schleiger A, Ovroutski S, Peters B, et al. Treatment strategies for proteinlosing enteropathy in Fontan-palliated patients. Cardiol Young 2020; 30: 698–709. DOI: 10.1017/S1047951120000864.
- 5. Brancaccio G, Carotti A, D'Argenio P, Michielon G, Parisi F. Protein-losing enteropathy after Fontan surgery: resolution after cardiac transplantation.

J Heart Lung Transplant 2003; 22: 484–486. DOI: 10.1016/s1053-2498(02) 01231-7.

- Berbel-Franco D, Lopez-Delgado JC, Putzu A, et al. The influence of postoperative albumin levels on the outcome of cardiac surgery. J Cardiothorac Surg 2020; 15: 78. DOI: 10.1186/s13019-020-01133-y.
- Ohuchi H, Yasuda K, Miyazaki A, et al. Comparison of prognostic variables in children and adults with Fontan circulation. Int J Cardiol 2014; 173: 277–283. DOI: 10.1016/j.ijcard.2014.03.001.
- Leite HP, Fisberg M, de Carvalho WB, de Camargo Carvalho AC. Serum albumin and clinical outcome in pediatric cardiac surgery. Nutrition 2005; 21: 553–558. DOI: 10.1016/j.nut.2004.08.026.
- Abel RM, Fisch D, Horowitz J, van Gelder HM, Grossman ML. Should nutritional status be assessed routinely prior to cardiac operation? J Thorac Cardiovasc Surg 1983; 85: 752–757.
- Leite HP, Fisberg M, Vieira JG, Carvalho WB, Chwals WJ. The role of insulin-like growth factor I, growth hormone and plasma proteins in surgical outcome of children with congenital heart disease. Pediatr Crit Care Med 2001; 2: 29–35.
- Geukers VG, Dijsselhof ME, Jansen NJ, et al. The effect of short-term high versus normal protein intake on whole-body protein synthesis and balance in children following cardiac surgery: a randomized double-blind controlled clinical trial. Nutr J 2015; 14: 72. DOI: 10.1186/s12937-015-0061-9.
- Yost G, Tatooles A, Bhat G. Preoperative nutritional assessment with the prognostic nutrition index in patients undergoing left ventricular assist device implantation. ASAIO J 2018; 64: 52–55.
- Onodera T, Goseki N, Kosaki G. Prognostic nutritional index in gastrointestinal surgery of malnourished cancer patients. Nihon Geka Gakkai Zasshi 1984; 85: 1001–1005.
- Radman M, Mack R, Barnoya J, et al. The effect of preoperative nutritional status on postoperative outcomes in children undergoing surgery for congenital heart defects in San Francisco (UCSF) and Guatemala City (UNICAR). J Thorac Cardiovasc Surg 2014; 147: 442–450. DOI: 10.1016/j. jtcvs.2013.03.023.
- El-Koofy N, Mahmoud AM, Fattouh AM. Nutritional rehabilitation for children with congenital heart disease with left to right shunt. Turk J Pediatr 2017; 59: 442–451. DOI: 10.24953/turkjped.2017.04.011.
- Schleiger A, Ovroutski S, Peters B, et al. Treatment strategies for proteinlosing enteropathy in Fontan-palliated patients. Cardiol Young 2020; 30: 698–709. DOI: 10.1017/S1047951120000864.
- Luo WY, Xu ZM, Zhang YY, Hong L, Zhang MJ, Zhang YQ. The nutritional status of pediatric patients with single ventricle undergoing a bidirectional glenn procedure. Pediatr Cardiol 2020; 41: 1594–1600. DOI: 10.1007/ s00246-020-02416-w.
- Koertzen M, Punjabi P, Lockwood G. Pre-operative serum albumin concentration as a predictor of mortality and morbidity following cardiac surgery. Perfusion 2013; 28: 390–394. DOI: 10.1177/0267659113488990.
- Zaccagni HJ, Alten JA, Cleveland DC, et al. Early postoperative albumin administration contributes to morbidity after the Fontan operation. Pediatr Cardiol 2016; 37: 1278–1283. DOI: 10.1007/s00246-016-1429-0.
- Schiller O, Goldshmid O, Mowassi S, et al. The utility of albumin level as a marker of postoperative course in infants undergoing repair of congenital heart disease. Pediatr Cardiol 2020; 41: 939–946. DOI: 10.1007/s00246-020-02339-6.
- Mwaura L, Vuylsteke A. Fueling the debate on albumin after cardiac surgery. J Cardiothorac Vasc Anesth 2019; 33: 2928–2929. DOI: 10.1053/j. jvca.2019.04.030.
- Karas PL, Goh SL, Dhital K. Is low serum albumin associated with postoperative complications in patients undergoing cardiac surgery? Interact Cardiovasc Thorac Surg 2015; 21: 777–786. DOI: 10.1093/icvts/ ivv247.
- Rauf A, Joshi RK, Aggarwal N, et al. Effect of albumin addition to cardiopulmonary bypass prime on outcomes in children undergoing openheart surgery (EACPO study)-a randomized controlled trial. World J Pediatr Congenit Heart Surg 2021; 12: 61–69. DOI: 10.1177/21501351 20959088.