Introduction:

Portal hypertension (PH) and its complications are the main cause of morbidity and mortality in pediatric patients with cirrhosis or end-stage liver disease. Those patients should require a threshold value of hepatic venous pressure gradient of 10 to 12 mmHg for the development of varices. The aim of this work was to evaluate the influence of splenectomy with disconnection of short veins of stomach by using Child-Turcotte-Pugh score in a small case series of four pediatric patients with liver disease or cirrhosis.

Patients:

From July 2016 to August 2017, 4 patients with liver disease were diagnosed with hypersplenism or splenomegaly and portal hypertensive gastropathy. Underwent laparoscopic or open procedure, we performed splenectomy with disconnection of azygos or stomach short veins (vascular disconnection).

Outcomes:

The patients 1, 2 and 4 had a good outcome, the platelet account was recovered and no ascites and encephalopathy. Prothrombin time was perfectly and they not required variceal endoscopic treatment at month after the procedure. Endoscopic valuation showed variceal gastropathy hypertensive mild, that not required endoscopic therapy and the medical therapies with non-selective Beta-blocker were decreased; the Child-Pugh class B was changed to class A. Patient 3: presented the same outcome, even though the Child-Pugh class A was not altered.

Conclusion:

The splenectomized pediatric patient requires careful monitoring in the early postoperative period to identify some complications. Fortunately, in this study we observed a good outcome, amelioration of the portal hypertension gastropathy and a better function of liver.

Key Words: Portal hypertension- Liver function- Splenectomy- Vascular disconnection- Variceal bleeding

INTRODUCTION

Portal hypertension (PH) and its complications are the main cause of morbidity and mortality in pediatric patients with cirrhosis or end-stage liver disease. In fact, hypersplenism or splenomegaly like PH complication is considered a risk of leukopenia, bleeding, variceal formation, variceal bleeding and capsular disruption. However, the triad of splenomegaly, leukopenia and liver cirrhosis, is known as Banti’s disease. The initial event that leads to the development of portal hypertension is the increased intrahepatic resistance that is due to architectural disturbances and sinusoidal contractile elements. Patients with cirrhosis and sinusoidal portal hypertension should require a threshold value of hepatic venous pressure gradient of 10 to 12 mmHg for the development of varices. The mortality rate from variceal bleeding in cirrhotic patients with portal vein system thrombosis is 30%-70%. Finally, Portal hypertension in patients with liver cirrhosis is characterized primarily by increased intrahepatic vascular resistance and increased splanchnic blood flow.
The Role of Splenectomy with Vascular Disconnection in Pediatric Patients with Hypersplenism, Portal Hypertension and Liver Disease

Cirrhotic patients with portal hypertension have different levels of hypersplenism. The decrease of hepatic blood flow further damaged to the liver function, even lead to liver failure, hepatic encephalopathy and even death. Under an adequate conditions, near of 30% of the whole body blood platelets stored in the spleen, but it can increase to 80% in view of the hypersplenism, splenomegaly. Therefore, splenectomy has been an option indicated to patients who suffer from hypersplenism with chronic liver diseases. Furthermore, several reports described a decrease in the portal inflow and pressure after splenectomy. The aim of this work was to evaluate the influence of splenectomy with disconnection of azygos or short veins of stomach (vascular disconnection) by using Child-Turcotte-Pugh score and classification in a small series of four Pediatric Patients with liver disease or cirrhosis, portal hypertension and hypersplenism in our Medical Center (Pediatric Hospital “Dr. Silvestre Frenk Freund”, National Medical Center Siglo XXI/ México).

Patients

From July 2016 to August 2017, 4 patients with liver disease were diagnosed with hypersplenism or splenomegaly and portal hypertensive gastropathy due to portal hypertension in our gastro-transplantation department required treatments. Underwent laparoscopic or open procedure, we performed splenectomy with disconnection of azygos or stomach short veins (vascular disconnection). The inclusion criteria were as follows: clinical bleeding history, splenomegaly secondary to hypersplenism and thrombocytopenia (platelets account (PLT) < 40X10^4/L), Child-Pugh liver function. Retrospectively collected preoperative data of the patients, age, gender, and weight, etiology of cirrhosis, ascites, encephalopathy, platelet count (PLT), prothrombin time (PT), international normalized ratio (INR), encephalopathy, albumin, serum bilirubin levels (TBIL), alanine transaminase (ALT), glutamic oxaloacetic transaminase (AST). The four patients were received Pneumovax Vaccine from 15 to 20 days before the procedure with intravenous antibiotics as a preoperative prophylaxis. Endoscopy procedure, computed abdominal tomography and Doppler ultrasound were evaluated.

Case 1

A 3-year-old male of 15 kg of weight and 1.2 cm of size, was referred to our center with 2 years’ history of hypersplenism secondary to portal hypertension and portal cavernous transformation (PCT). During this time, he had prior history of esophageal variceal and portal hypertensive gastropathy. Physical examination showed evidence of hepatosplenomegaly and slight ascites. The patient had a good answer with the treatments for PH but he has continued with thrombocytopenia. He had TBIL 1.8 mg/dl; INR 1.7, PT of 17 seconds, albumin serum of 3.3 g/dl, ALT 200 UI/L, AST 416 UI/L, PLT 38 X 10^4 and she had Child-Pugh class B. Abdominal ultrasound and computed tomography angiography revealed portal cavernous, permeability of the spleen vasculature and the larger of the spleen (figures: 1, 2). The patient was evaluated and we performed a laparoscopic splenectomy with ligation of short stomach veins.

Fig1. Computed abdominal tomography angiography.  
Fig2. spleen size
Case 2

A 9-year-old male of 25.6 kg of weight and 1.32 cm of size, with multiples surgical procedures secondary gastroesophageal reflux, hemangioma and portal hypertension, he was referred to our department with 5 years' history of hypersplenism, portal hypertensive gastropathy (PHG), PCT and cirrhosis. The child had history of variceal bleeding due to severely gastroesophageal varices requiring ligation variceal endoscopic and medical therapy with non-selective beta-blockers. Physical examination showed evidence of hepatosplenomegaly, moderate ascites and abdominal distension. Preoperative laboratory tests showed serum bilirubin 1.26 mg/dl and INR 1.34; TP of 13.4 seconds, albumin serum of 4.5 g/dl and grade 2 of encephalopathy. However, with the medical therapy, the patient continued with thrombocytopenia with platelet count of 33 X10⁴ and recurrent variceal bleeding. He had Child-Pugh class B and severe varices in the last endoscopic before the procedure. Abdominal ultrasound and computed tomography angiography (CTA) revealed cirrhosis, PCT, permeability of the spleen vasculature and the larger of the spleen (figures: 3, 4). The patient was evaluated and we performed an open splenectomy with ligation of short stomach veins and liver biopsy.

Fig3. Doppler of portal vein  
Fig4. CTA showed PCT and spleen size

Case 3

A 5-month-old female of 3.5 kg of weight and 0.56 cm of size, with progressive familial intrahepatic cholestasis type 2 (PFIC2); she was referred to our department with hypersplenism secondary to portal hypertension and splenomegaly. Her Physical examination showed evidence of hepatosplenomegaly and slight ascites. She had TBIL of 18 mg/dl and INR 1.7; PT of 17 seconds, albumin serum of 3.3 g/dl, ALT of 200 UI/L, AST of 416 UI/L, platelets account (PLT) 6 X 10⁴ and she had Child-Pugh class A.

Fig5. Portal Doppler ultrasound  
Fig6. CTA showed spleen size.
The Role of Splenectomy with Vascular Disconnection in Pediatric Patients with Hypersplenism, Portal Hypertension and Liver Disease

Underwent the medical treatments for the PH, she continued with thrombocytopenia with platelet count below 40 X 10^4. Abdominal ultrasound and computed tomography revealed liver disease; permeability of the spleen vasculature and the larger of the spleen (figures: 5, 6). She was evaluated and we performed a laparoscopic splenectomy (LS) with ligation of short stomach veins (LSSV) and liver biopsy.

**Case 4**

A 9.6-year-old female of 30 kg of weight and 1.3 cm of size, with cholestasis and cirrhosis secondary to biliary atresia was referred to our department. She had history of Kasai procedure and multiples endoscopic therapy for portal hypertensive variceal bleeding. Her Physical examination showed evidence of splenomegaly and hypersplenism secondary to portal hypertension and Slight ascites. She had serum bilirubin total of 0.49 mg/dl, INR 1.2, TP of 11 seconds, albumin serum of 3.9 g/dl, grade 2 of encephalopathy and she had Child-Pugh class B. Platelets account 10 x 10^3, AST 69 and ALT 74 UI/L. Furthermore, the medical treatments for the PH she continued with thrombocytopenia with platelet count below 40 X10^4. Abdominal ultrasound and computed tomography revealed cirrhosis, permeability of the spleen vasculature and the larger of the spleen (figures: 7, 8). The patient was evaluated and we performed an open splenectomy with ligation of short stomach and azygos veins and liver biopsy.

**Postoperative Outcomes**

The patient 1 had a good outcome and not required transfusion after the procedure, the platelets account were recovered, no ascites and encephalopathy, prothrombin time was normalized, international normalized ratio (INR) was 1.4 and serum bilirubin levels were < 1 and he not required variceal endoscopic treatment at month after the procedure. The Child –Pugh class passed to A. Six months after the surgical procedure open splenectomy with disconnection of short gastric and azygoportal veins; patient 2 presented laboratory tests ammonium serum 48 ( range 11 – 48) with previous of 78, platelet account was 441 x 10^4, without ascites and encephalopathy; prothrombin time was 9.8 ( range 9.3 – 12.3). INR 0.98, total bilirubin serum 0.54 mg/dl. Endoscopic valuation showed variceal gastropathy hypertensive mild, that not required endoscopic therapy and the medical therapy with non–selective Beta-blocker was decreased.

<table>
<thead>
<tr>
<th>Table1. Characteristics of Patients Pre and Postsplenectomy</th>
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<td><strong>Clinical Characteristics</strong></td>
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<tr>
<td>Etiology</td>
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<td>-PCT</td>
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<td>-HS</td>
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<td>-Cirrhosis</td>
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## The Role of Splenectomy with Vascular Disconnection in Pediatric Patients with Hypersplenism, Portal Hypertension and Liver Disease

<table>
<thead>
<tr>
<th>Child-Pugh Class</th>
<th>Platelets:</th>
<th>AST(TGO): UI/L</th>
<th>ALT(T GO): UI/L</th>
<th>INR</th>
<th>PHG</th>
<th>Beta-blockers doses</th>
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<tr>
<td>B</td>
<td>38 X 10⁴</td>
<td>416</td>
<td>200</td>
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<td>Moderate</td>
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</tr>
<tr>
<td>A</td>
<td>33 X 10⁴</td>
<td>37</td>
<td>25</td>
<td>1.34</td>
<td>Severe</td>
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<tr>
<td>B</td>
<td>6 X 10⁴</td>
<td>417</td>
<td>200</td>
<td>1.7</td>
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<tr>
<td>B</td>
<td>10 x 10⁴</td>
<td>69</td>
<td>74</td>
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<td>Severe</td>
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<th>Surgical Procedure</th>
<th>LS/LSSV</th>
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<th>OS/LSSV/LAV</th>
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<td>Outcomes</td>
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<tr>
<td>Child-Pugh class</td>
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<td>A</td>
<td>A</td>
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<tr>
<td>Platelets:</td>
<td>195X10⁴</td>
<td>441 X 10⁴</td>
<td>492 X10⁴</td>
<td>270 X10⁴</td>
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<td>AST(TGO): UI/L</td>
<td>71</td>
<td>45</td>
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<td>390</td>
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<tr>
<td>ALT(T GO): UI/L</td>
<td>38</td>
<td>39</td>
<td>0.98</td>
<td>209</td>
</tr>
<tr>
<td>INR</td>
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<td>0.98</td>
<td>1.2</td>
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<td>PHG</td>
<td>Mild</td>
<td>Mild</td>
<td>Mild</td>
<td>Decreased</td>
</tr>
<tr>
<td>Beta-blockers doses</td>
<td>Decreased</td>
<td>Decreased</td>
<td>Decreased</td>
<td>Decreased</td>
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The Child-Pugh class B was passed to class A. Patient 3: presented a good outcome. Platelet account was 250 x 10⁴, prothrombin time was 11 (range 9.3-12.3), INR 1.2, total bilirubin serum 10 mg/dl. The Child-Pugh class A was the same. AST was 390 and ALT was 209 UI/L. She had a mild portal hypertension without variceal bleeding during the time of follow-up. Patient number 4, six months after the procedure had serum ammonium 73 (range 11-48), platelet account 270 x 10⁴. Prothrombin time was 11 (range 9.3-12.3), INR 0.94, total serum bilirubin 0.79 mg/dl. The B Child-Pugh classification passed to class A. AST-ALT were 59/56 UI/L respectively, and without encephalopathy. The level of portal hypertensive gastropathy observed in the endoscopic control was decreased to moderate.

### Discussion

The spleen is an important lymphoid organ, which has the function of storing blood, hematopoiesis, and immunizing power; however removal the spleen by laparoscopic or open procedures is an effective invasive treatment for spleen rupture, portal hypertension, hypersplenism, splenomegaly and idiopathic thrombocytopenic purpura. Even though, after splenectomy it can bring high risk of infection, immune function decline, and thrombocytosis.4,7,8 in this study they were not observed those complications. Generally, vaccines (pneumococcal vaccine, haemophilus influenza type b conjugate vaccine, and meningococcal conjugate vaccine and N. meningitidis) were administered 2 weeks before splenectomy to all patients.4,6-10 However all patients in our study had pneumococcal vaccine from 2 to 3 weeks before the splenectomy. The aim of present study was to evaluate relationships between degree of portal hypertension and follow-up, during six months after laparoscopic or open splenectomy with disconnection of short stomach or azygos veins.

The histological characteristics findings of PHG are dilated capillaries and venules in the mucosa and submucosa without erosion, inflammation, or fibrinous thrombi.11-13 Therefore, our patients were classified under the classification of portal hypertensive gastropathy according to new Italian Endoscopic Club and the commonly endoscopy classification: that’s mean Mosaic like pattern: mild-diffusely pink areola, moderate-flat red spot in centre of pink areola, severe-diffusely red areola, red lesions of variable diameter, flat or slightly protruding, discrete o confluent.12 respectively, in the commonly endoscopy classification; F1, small straight varices; F2,
enlarged tortuous varices that occupy less than one third of the lumen; and F3, large coil-shaped varices that occupy more than one-third of the lumen.3 Our patients had moderate to severe PHG that’s mean F2 to F3 respectively.

Under physiologic condition, after splenectomy; there are proliferations of platelet megakaryocyte in bone marrow, induced platelets rapidly increased 2-6 times in the blood circulation. This effect was observed in our patient after the splenectomy.10,14 On the other hand, the risk of sepsis is greatest during the first 2 years after splenectomy. Children under 5 years old at the time of splenectomy have a greater risk for sepsis than older children and adult; therefore, in our study they had 2 patients < 5 years without infections in the first 6 months after splenectomy.4,11,15 Pancreatic fistula is another rare complication in patient after laparoscopic splenectomy with prevalence range from 4.5% to 16% in patient with hematological diseases; Tsutsumi report that the incidence of pancreatic fistula after laparoscopic splenectomy in patient with hypersplenism due to liver cirrhosis is not low. However, we had not this complication in our patients.14

Hypertension gastropathy commonly encountered in patients with cirrhosis, with prevalence between 20% and 80% and observed more disease. In fact, hepatic venous pressure gradient (HVPG) is higher and systemic vascular resistance lower in patients with PHG. Prostaglandins, Nitric oxide synthase and local ischaemia also appear to play a role.3,12,16 Portal hypertension gastropathy is the most common gastric mucosal injury in patients with liver cirrhosis. Furthermore, there is an increase in total gastric blood flow but with a changed distribution: a relative decrease in blood flow to the mucosa and increased blood flow to the submucosa, muscle, and serosal layers.16 On the other hand, idiopathic non-cirrhotic intrahepatic portal hypertension is characterized by occlusion of the small intrahepatic portal vein radicles.17 Other disease that playing an important role in the portal hypertension is the portal cavernoma (PC) caused gastrointestinal bleeding among children, some studies mentioned at least one episode of upper gastrointestinal bleeding was present in approximately 80% of patients with PC or better known like cavernous transformation of the portal vein (CTP).7 In our review, fifty percent of patients had CTP with hypertensive gastropathy and gastrointestinal bleeding that is not different in literature report.

There are various proposal of treatment for PH; Kalambokis George N et al, in its brief report suggested that terlipressin should be considered the vasoconstrictor treatment of choice when cirrhotic patients with PH present with variceal bleeding or hepatorenal syndrome (HRS) in our series of four patients we used beta blockers and our patients had not HRS.9 In our study, Patients 1, 2 and 4 had more than two episodes of bleeding. The indications for splenectomy in cirrhotic patients are somewhat controversial.15 However, the most common indications of splenectomy are for hereditary spherocytosis, idiopathic thrombocytopenic purpura, sickle cell anemia and hematologic diseases; all of our patients had hypertensive gastropathy due to portal hypertension and other disease of the liver; we had one patient with sickle cell anemia that presented a good outcome after a laparoscopic splenectomy.11

Portal hypertension might eventually develop with esophagogastric varices, splenomegaly, and hypersplenism.7 Like it was in our study, Guo H. reported presented nine cases with open retroperitoneal communicating branches (venous plexus of retzius), to prevent variceal bleeding in its study. However, in this same study they performed disconnection in the gastric fundus and lower esophagus in combination with splenectomy was performed in 36 patients with CTP and they observed thrombus formation in eight patients among them; surgical thrombus removal was performed in those patients. In fact, in our small series of cases we performed the disconnection in the mayor curvature (Coronary Venous and in lower esophagus) of the stomach with splenectomy and we had not thrombi in six months of following. For patients with cirrhosis, portal hypertensive bleeding and hypersplenism, open splenectomy with azygoportal disconnection is generally accepted as the most effective approach. However, one of the most serious perioperative complications during laparoscopic splenectomy disconnection (LSD) is rapid loss of large volumes of blood. In our series of cases, patients one and two had LSD without loss of large volumes of blood.8
The normal diameter of the portal vein ranges from 6 to 10 mm; Li et al. demonstrated that portal vein diameter > 13 mm was an independent risk factor of portal thrombosis after open splenectomy in cirrhotic patients with hypersplenism. However, some authors reported systemic inflammatory response syndrome experienced in 54% and the portal vein thrombosis experienced in 17% of the patients. In this study one patient had systemic inflammatory with good answer to the treatment. Portal vein thrombosis is a potentially lethal complication after splenectomy with reported incidence of 6% in children and more than 55% in adults. The mechanism and risk factors of this complication are poorly understood, especially in cirrhotic patients with portal hypertension. Fortunately, in our four pediatric patients, this complication was not observed.

Severe complications such as upper gastrointestinal bleeding, refractory ascites, jaundice, hepatic encephalopathy, bowel edema acute ischemic necrosis, ileus and other serious complications were not observed also in this study. Some adult studies have demonstrated with splenectomy, portal vein pressure is reduced, which alleviates esophagogastric varices and prevents esophagogastric variceal bleeding. Furthermore, several important aspects of chronic liver diseases still remain to be fully studied. Splenectomy improves also liver function. Similarly situations, were observed in our four pediatric patients. Bai et al. reported that, azygoportal disconnection can effectively reduce the recurrence rate of esophagogastric variceal bleeding, which is a major cause of death in patients with portal hypertension due to cirrhosis in adult population similarly effects were observed in our series of pediatric patients. Finally, Yamamoto et al. observed that the liver function had changed to Child-Pugh class C after progression of liver disease with portal vein invasion at 1 year after splenectomy. In our four patients with portal hypertension we observed diminution of portal hypertension and Child-Pugh class at 6 months of the procedure was maintained or change to a better score. The mechanism underlying this functional amelioration of the liver after splenectomy has yet to be fully elucidated. Furthermore, platelet-derived serotonin has recently been shown to be important in liver regeneration.

CONCLUSION

The splenectomized pediatric patient requires careful monitoring in the early postoperative period to identify some complications like formation of portal thrombi and infections; in this study we observed a good outcome on the platelets account without bleeding, and amelioration of the portal hypertension gastropathy similarly a better function of liver or prolongation of the time of liver deterioration. Through this time of follow-up any patient required variceal treatment by endoscopic method and the risk of variceal bleeding was decreased. The weakness of this study was the number of patients and carefully patient selection.

REFERENCES


The Role of Splenectomy with Vascular Disconnection in Pediatric Patients with Hypersplenism, Portal Hypertension and Liver Disease


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