Resected Intrahepatic Cholangiocarcinoma with Anaphylactic Shock from a Preoperative Liver Function Test before Hepatectomy

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Abstract
The indocyanine green test is a reliable liver function examination before major hepatectomy, and anaphylaxis is rarely a concern. A 65-year-old male patient without epigastralgia was diagnosed with a 2.2-cm intrahepatic cholangiocarcinoma. He had no history of allergic reactions. Some liver dysfunction was indicated by the laboratory data; however, there was no marked obstructive jaundice and the liver functional reserve was maintained by technetium-99m galactosyl serum albumin. The indocyanine green test was routinely performed, but the patient immediately demonstrated severe anaphylaxis due to indocyanine green administration. He had cardiorespiratory arrest, but recovered after immediate resuscitation. Although acute renal and respiratory failure was significant, the patient recovered at day 10 after the event, and his liver function and other organ functions were improved. Then, the scheduled left hepatectomy with caudate and extrahepatic duct resection was successfully performed without issues. The patient exhibited no allergic response against the administration of antibiotics or other drugs and the postoperative course was uneventful. The patient was discharged.
on day 17. The tumor was diagnosed as stage III intrahepatic cholangiocarcinoma and R0 re- section was accomplished. Preoperative management, including the liver functional loading test, should be carefully carried out before major hepatectomy.

Introduction

In the field of hepatic surgery, the functional liver reserve is usually precisely examined before hepatectomy to avoid post-hepatectomy liver failure, particularly for major hepatectomy. In Japan, the indocyanine green (ICG) test has been widely applied as the gold standard test even in patients with Child-Pugh class A disease [1]. According to this parameter, the extension of hepatectomy is limited [2]. This preoperative test has recently been applied to detect the existence of liver tumors as the photodynamic eye test [3]. Administration of ICG is supposed to be safe even in patients with a chronically injured liver; however, although anaphylaxis from ICG has rarely been reported, it may lead to severe organ damage [4]. We herein report a successfully resected case of primary intrahepatic cholangiocarcinoma with severe anaphylactic shock leading to organ failure at the time of the preoperative liver test before hepatectomy.

Case Report

A 65-year-old male patient without epigastralgia was diagnosed with intrahepatic cholangiocarcinoma with peripheral biliary obstruction. He had no remarkable coexisting disease and no history of allergic reactions. The physical examination showed no abnormalities and laboratory data demonstrated liver dysfunction. At the time of admission, increased serum levels of aminotransferase enzymes (aspartate aminotransferase 90 IU/L, alanine aminotransferase 157 IU/L; normal range <325 IU/L) were observed, but the bilirubin level was normal (total bilirubin 1.0 mg/dL). The serum level of carcinoembryonic antigen was increased to 21.4 ng/mL (normal range <5 ng/mL). The liver uptake ratio at 15 min by technetium-99m galactosyl serum albumin (99mTc GSA) was within the normal range at 95% [5]. Cardiorespiratory and renal function was almost normal (BUN 10 mg/dL, creatinine 0.7 mg/dL).

Contrast-enhanced computed tomography (CT) revealed a 2.2-cm irregular low-density space-occupying lesion at the left hepatic duct with obstructive dilation of the peripheral biliary trees (Fig. 1). Although endoscopic biliary drainage was performed, no allergic reactions to the contrast medium for imaging analysis were observed. The ICG test was routinely carried out; however, the patient immediately exhibited unconsciousness and cardiorespiratory arrest due to anaphylaxis from ICG administration. Resuscitation was rapidly performed and consciousness was recovered at 34 min after resuscitation. The serum BUN and creatinine levels were markedly increased (BUN 36 mg/dL, creatinine 2.6 mg/dL), the urinary volume was decreased to 60 mL per 24 h, and arrhythmia was observed. By management in the intensive care unit using inotropic agents, an artificial ventilator, and diuretic treatment, the patient recovered at day 10 after the event.

Liver function and other organ functions were rapidly improved (BUN 17 mg/dL, creatinine 0.9 mg/dL, total bilirubin 0.7 mg/dL, and alkaline phosphatase 1,694 IU/L), and the scheduled left hepatectomy with caudate and extrahepatic duct resection was successfully
performed (operating time 9 h 32 min, blood loss 1,190 mL). The patient demonstrated no allergic response to the administration of antibiotics or other drugs and the postoperative course was uneventful. The patient was discharged at day 17. Histologically, the tumor was diagnosed as stage III intrahepatic cholangiocarcinoma and R0 resection was accomplished.

Discussion/Conclusion

The ICG test is supposed to be a safe and reliable liver function test in Japan [1, 6], and this test is easily performed not only as a preoperative test but also for photodynamic diagnosis during operations [3]. On the other hand, anaphylactic reactions have been previously reported [4], but the prevalence is very low. In our experience, only 1 case out of approximately 650 cases exhibited mild hives. Although this patient had no allergic reactions to the contrast medium for imaging analysis, he exhibited anaphylaxis from ICG testing. Olsen et al. [7] reported a case of anaphylactic shock following ICG angiography in a patient without iodine allergy, so we should keep in mind that serious allergic reactions to ICG may occur with or without iodine allergy.

Allergic responses seem to vary, and in the present case, the patient developed renal and respiratory failure once in a critical situation. Fortunately, he recovered within 1 week. Although abandoning the scheduled major hepatectomy was a concern due to multiorgan injuries by cardiac arrest and anaphylactic shock, his systemic condition was recovered. By recalling this event, it is necessary to check for allergic shock after administration in cases with ICG testing.

Preoperative problems or coexisting diseases may affect postoperative complications. $^{99m}$Tc GSA is as reliable a liver function test as the ICG test, and GSA is a safe drug [5]. Allergic reactions have not been reported yet. This test showed the distribution of active and living hepatocytes. Previous reports have demonstrated that the liver uptake ratio from $^{99m}$Tc GSA scintigraphy could be converted into an ICGR15 value even in cases of obstructive jaundice [8]. Therefore, patients with a history of allergy can be evaluated with this reliable test, although the cost is high compared with the ICG test.

In conclusion, preoperative management, including the liver functional loading test, should be carefully carried out before major hepatectomy.

Statement of Ethics

The authors have no ethical conflicts to disclose.

Disclosure Statement

The authors have no conflicts of interest to declare.
Author Contributions

K. Higuchi and A. Nanashima wrote this paper. N. Imamura, K. Yano, and Y. Tsuchimochi performed the surgery and perioperative management. M. Hiyoshi, T. Hamada, T. Wada, K. Tsuchiya, F. Kawano, T. Ikeda, and S. Takeno contributed to the patient’s care.

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Fig. 1. a, b Enhanced CT with contrast medium and endoscopic cholangiography showed a low-density mass-forming lesion in the left liver obstructing the left hepatic duct (arrowhead). Left hepatectomy was scheduled following this imaging finding.