



## Evaluation of hepatic perfusion in the liver graft using fluorescence imaging with indocyanine green

Yoshikuni Kawaguchi, Nobuhisa Akamatsu\*, Takeaki Ishizawa, Junichi Kaneko, Junichi Arita, Yoshihiro Sakamoto, Kiyoshi Hasegawa, Norihiro Kokudo\*

*Artificial Organ and Transplantation Surgery Division, Department of Surgery, Graduate School of Medicine, University of Tokyo, Tokyo, Japan*



### ARTICLE INFO

#### Article history:

Received 1 June 2015  
Received in revised form 18 July 2015  
Accepted 28 July 2015  
Available online 31 July 2015

#### Keywords:

Portal vein thrombosis  
Living-donor liver transplantation  
Indocyanine green  
Fluorescence imaging technique

### ABSTRACT

**INTRODUCTION:** Portal vein thrombosis (PVT) after liver transplantation (LT) is one of serious complications and reportedly ranges from 2% to 13%. PVT impairs the blood perfusion to the grafts and causes the graft dysfunction.

**PRESENTATION OF CASE:** A 60-year-old female underwent living-donor LT with the left liver graft for end-stage liver disease related to chronic hepatitis C. After reperfusion, Indocyanine green (ICG)-fluorescence imaging was performed to confirm the graft perfusion, which pointed out an insufficient perfusion on the surface of segment 4. Following intraoperative ultrasonography revealed thrombus in the portal vein of segment 4, which was successfully removed by heparinized saline flush.

**DISCUSSION:** The most of patients with PVT developed graft failure and resulted in retransplantation. This enhances the importance of the surveillance for PVT in the postoperative period as well as the intraoperative period. However, the modality to identify PVT during surgery is limited mainly to intraoperative ultrasound. ICG-fluorescence imaging can visualize regions with impaired hepatic perfusion due to PVT in real time during LT in addition to visualization of hepatic flows of reconstructed vessels and evaluation of regions with venous occlusion.

**CONCLUSION:** ICG-fluorescence imaging can be simply performed with single ICG injection and is expected to have potential roles to enhance the safety of LT.

© 2015 The Authors. Published by Elsevier Ltd. on behalf of Surgical Associates Ltd. This is an open access article under the CC BY-NC-ND license (<http://creativecommons.org/licenses/by-nc-nd/4.0/>).

Portal vein thrombosis (PVT) after liver transplantation (LT) is one of serious complications and reportedly ranges from 2% to 13% [1,2]. PVT impairs the blood perfusion to the grafts and causes the graft dysfunction. The most of patients with PVT developed graft failure and resulted in retransplantation [1,2]. This enhances the importance of the surveillance for PVT in the postoperative period as well as the intraoperative period. However, the modality to identify PVT during surgery is limited mainly to intraoperative ultrasound. Herein, we describe an alternative method to identify PVT during liver transplantation using indocyanine green (ICG)-fluorescence imaging.

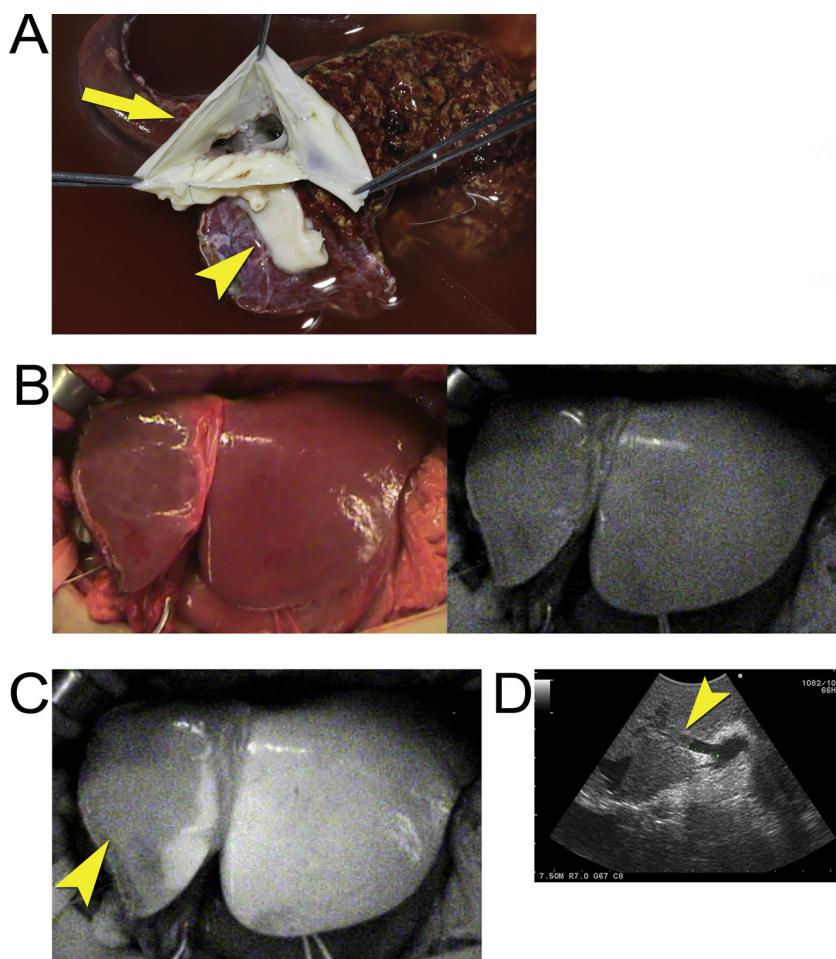
A 60-year-old female underwent living-donor LT for end-stage liver disease related to chronic hepatitis C. The left liver graft with

caudate lobe (372 mL, 36.7% of the recipient standard liver volume) was procured, and venoplasty with a circular cuff vein patch at the stump of the middle and left hepatic veins using cryopreserved veins was performed to avoid outflow obstruction [3] (Fig. 1A). After reconstruction of all hepatic vessels, ICG (0.93 mg; 2.5 µg per 1 mL of graft LV) was administered intravenously to evaluate anastomotic sites of reconstructed hepatic artery and portal vein and venous congestion in the graft related to venous reconstruction [4]. ICG-fluorescence imaging visualized continuous arterial and portal flow passing anastomotic sites. Fluorescence intensity (FI) on the left liver graft surface increased gradually while FI on the surface of segment 4 was lower than that on left lateral sector (Fig. 1B and C, and Supplementary video 1). Doppler intraoperative ultrasonography (D-Ious) showed no evidence of impaired venous flow in the tributary of the segment 4, but revealed thrombus in the portal vein flowing into the segment 4 (Fig. 1D). The heparinized saline was injected and flushed from the portal venous anastomosis, which successfully melted away the thrombus. The postoperative course was uneventful except she suffered from herpes zoster, and the patient was discharged on postoperative days 48 without the development of PVT any more.

**Abbreviations:** D-Ious, Doppler intraoperative ultrasonography; FI, fluorescence intensity; ICG, indocyanine green; LT, liver transplantation; PVT, portal vein thrombosis.

\* Corresponding authors at: Artificial Organ and Transplantation Surgery Division, Department of Surgery, Graduate School of Medicine, University of Tokyo, 7-3-1 Hongo, Bunkyo-ku, Tokyo 113-8655, Japan. Fax: +81 3 5800 8843.

E-mail addresses: [nakamats-tky@umin.ac.jp](mailto:nakamats-tky@umin.ac.jp) (N. Akamatsu), [KOKUDO-2SU@h.u-tokyo.ac.jp](mailto:KOKUDO-2SU@h.u-tokyo.ac.jp) (N. Kokudo).



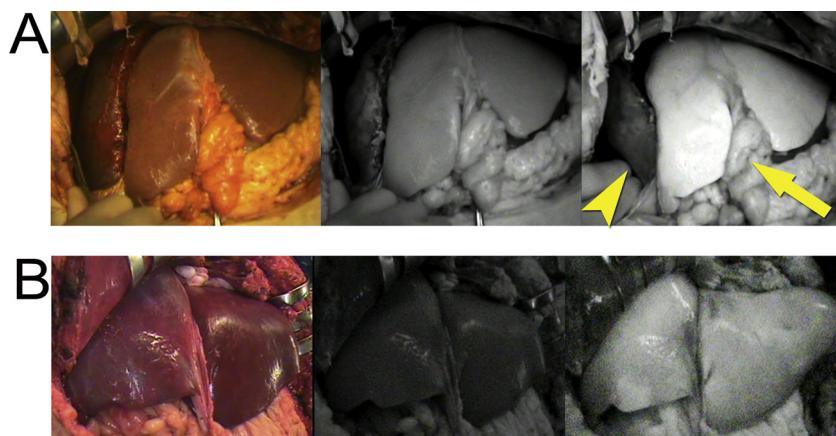
**Fig. 1.** Identification of portal vein thrombus using ICG-fluorescence imaging.

(A) Venoplasty with a circular cuff vein patch (arrow) at the stump of the middle and left hepatic veins which was combined with the stump of the short hepatic vein (arrowhead).

(B) Intraoperative gross appearance of the left liver graft (left) and corresponding fluorescence images prior to intravenous injection of ICG (right).

(C) Fluorescence imaging following intravenous injection of ICG provided demarcation of FI between the segment 4 (arrowhead) and left lateral sector.

(D) IOUS reveals a thrombus in the portal vein flowing into the segment 4 (arrow head).



**Fig. 2.** Application of ICG-fluorescence imaging for auxiliary partial orthotopic and deceased-donor LT.

(A) Fluorescence imaging visualized that the hepatic perfusion in the native liver (arrowhead) was impaired compared to the left liver graft (arrow) in auxiliary partial orthotopic LT (left; gross appearance, middle; fluorescence images before ICG injection, right; fluorescence images 240 s after the injection).

(B) Fluorescence images visualized the homogenous perfusion of the whole liver graft in the deceased-donor LT (left; gross appearance, middle; fluorescence images before ICG injection, right; fluorescence images 240 s after the injection).

Supplementary material related to this article found, in the online version, at doi:[10.1016/j.ijscr.2015.07.031](https://doi.org/10.1016/j.ijscr.2015.07.031)

Fluorescence imaging visualized regions in which hepatic perfusion was impaired due to PVT, and FI was lower in the regions with PVT than those without PVT. Hepatic perfusion levels in liver grafts can be evaluated in real-time by comparing FI values on the liver surface. The liver graft may develop hypoperfusion due to thrombosis, stricture, or occlusion in the portal vein when ICG-fluorescence imaging reveals FI heterogeneity on the liver surface. Previously, our groups demonstrated that FI was lower in veno-occlusive regions than in non-veno-occlusive regions [4,5]. These results imply that ICG-fluorescence imaging, during LT, can be utilized to visualize flows of reconstructed hepatic vessels, regions with venous occlusion, and regions with impaired hepatic perfusion caused by PVT as in this case. ICG-fluorescence imaging was applied to visualize anastomotic sites of arteries and lymphatic vessels in plastic surgery as well. In addition to this direct visualization of the reconstructed vessels, the present case might indicate the potential role of ICG-fluorescence imaging in visualizing the hypoperfusion area of the liver graft intraoperatively, which may help the real-time surveillance of hepatic circulation during LT. This technique was also applied for other type of LT; first, for the auxiliary partial orthotopic LT, in which perfusion of native liver found to be impaired while the satisfactory perfusion of the graft was confirmed (Fig. 2A), and second, for the whole liver graft from deceased-donor, in which the homogenous perfusion of the graft was confirmed (Fig. 2B). We believe that ICG-fluorescence imaging complement D-IOUS and contribute to demonstrate the problems of the graft circulation in real time during LT. These examinations are simply performed with the placement of fluorescence camera above the hepatic hilum or the graft surface after single ICG injection. Hepatic arterial and portal flows were visualized approximately 5–10 s after ICG injection and FI on the liver surface gradually increases spending 300 s and reaches a plateau. One of the drawbacks of this technique is that it cannot be applied repeatedly to evaluate hepatic perfusion of the graft in principle because the fluorescence on the liver surface lasts more than hours. By contrast, hepatic flows of the reconstructed vessels can be repeatedly evaluated using this technique. D-IOUS is the simplest and

most reliable modality for the intraoperative evaluation of hepatic circulation. ICG-fluorescence imaging may have additive roles in demonstrating not only the vessels themselves but also hypoperfusion or venous congestion in liver parenchyma, which may further improve the safety and certainty of LT procedures.

In conclusion, ICG-fluorescence imaging visualized regions with impaired hepatic perfusion due to PVT in real time during LT in addition to visualization of hepatic flows of reconstructed vessels and evaluation of regions with venous occlusion. These evaluations are simply performed with single ICG injection and are expected to be potential roles of ICG-fluorescence imaging to enhance the safety of LT.

### Conflict of interest

The authors disclose no conflicts.

### Financial support

This work was supported by a grant-in-aid for Scientific Research from the Ministry of Education, Culture, Sports, Science, and Technology of Japan and Grants-in-aid for Research on HIV/AIDS, and Research on Measures for Intractable Diseases from the Ministry of Health, Labor, and Welfare of Japan.

### References

- [1] L. Llado, J. Fabregat, J. Castellote, E. Ramos, J. Torras, R. Jordà, et al., Management of portal vein thrombosis in liver transplantation: influence on morbidity and mortality, *Clin. Transplant.* 21 (2007) 716–721.
- [2] J.P. Duffy, J.C. Hong, D.G. Farmer, R.M. Ghobrial, H. Yersiz, J.R. Hiatt, et al., Vascular complications of orthotopic liver transplantation: experience in more than 4200 patients, *J. Am. Coll. Surg.* 208 (2009) 896–903, discussion 03–5.
- [3] T. Hashimoto, Y. Sugawara, S. Tamura, J. Kaneko, N. Motomura, S. Takamoto, et al., One orifice vein reconstruction in left liver plus caudate lobe grafts, *Transplantation* 83 (2007) 225–227.
- [4] Y. Kawaguchi, Y. Sugawara, T. Ishizawa, S. Satou, J. Kaneko, S. Tamura, et al., Identification of veno-occlusive regions in a right liver graft after reconstruction of vein segments 5 and 8: application of indocyanine green fluorescence imaging, *Liver Transplant.* 19 (2013) 778–779.
- [5] Y. Kawaguchi, T. Ishizawa, Y. Miyata, S. Yamashita, K. Masuda, S. Satou, et al., Portal uptake function in veno-occlusive regions evaluated by real-time fluorescent imaging using indocyanine green, *J. Hepatol.* 58 (2013) 247–253.

### Open Access

This article is published Open Access at [sciencedirect.com](https://www.sciencedirect.com). It is distributed under the [IJSCR Supplemental terms and conditions](#), which permits unrestricted non commercial use, distribution, and reproduction in any medium, provided the original authors and source are credited.