LETTER TO THE EDITORS

Delayed kidney transplantation in combined liver– kidney transplantation for polycystic liver and kidney disease

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Dear Editors,

Patient survival after combined liver–kidney transplant (CLKT) is lower compared with liver transplantation (LT) alone. As well as the more complex surgical procedure, coagulopathy, significant peri-operative hypotension requiring high vasopressor doses, and hepatic ischemia–reperfusion injury, all compromise kidney allograft function.

In addition, delayed graft function (DGF) of the kidney graft has been reported to be a negative predictor of patient survival [1].

We report our preliminary experience in delayed kidney transplantation (KT) after a very long cold ischemia time in CLKT first performed in Italy, and to our knowledge reported in Europe on patients with polycystic liver and kidney disease (PLKD). This indication was not clearly discussed in the previous works by Ekser *et al.* [2,3].

Patients with polycystic liver disease show increased risk of peri-operative mortality as a consequence of the much more complicated hepatectomy procedure in the case of a huge liver [4,5].

To minimize the recipient's risk of bleeding and the inferior vena cava (IVC) torsion during liver mobilization, we routinely use the standard technique with IVC replacement and venovenous bypass. However, we usually perform primary abdominal closure, given the low risk of abdominal compartment syndrome in these recipients.

Among the over 2000 liver transplants performed in our high-volume center, the two last consecutive PLKD patients underwent a CLKT with a delayed KT from brain-dead donors in March 2019. The kidney grafts were maintained on continuous hypothermic pulsatile machine perfusion (MP) at 4 °C for 45 and 46 h, respectively, until KT. Continuous venovenous hemodialysis was maintained throughout this period, and recipients were completely weaned from vasopressors before kidney implantation. The delayed KT was performed extraperitoneally in the right iliac fossa through a separate incision. Native nephrectomy was not required in these two recipients.

Donor and recipient characteristics and peri- and postoperative outcomes are reported in Table 1. Patients were discharged with normal liver and kidney function, and neither suffered kidney DGF, defined as the need for dialysis within the first week after KT. No surgical complications related to the dual organ transplantation occurred, and patients are still alive and well 5 months after CLKT. The delayed KT approach was feasible in both two cases and the results of our first experience with the procedure, after an admittedly short follow-up, seem encouraging.

We would like to point out that all strategies aimed at avoiding kidney allograft futility in CLKT should be evaluated and, if feasible, adopted, especially in the uncommon setting of PLKD.

Prompt reporting of all applications of this novel approach would accelerate its widespread use and allows better understanding of whether synergy between the use of hypothermic MP and delayed KT optimizes kidney allograft function in CLKT, and might also allow safe expansion of the kidney donor pool on the basis of extended donor criteria, including kidney donation after circulatory death.

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| | Donor characteristics | Recipient characteristics | Peri-operative outcome | Postoperative outcome |
|--|--|---|--|---|
| | Age, year; sex, male/female Cause of death BMI, kg/m ² KDPI KDPI | Age, year; sex, male/female BMI, kg/m ² MELD score Total native liver weight, g s-Cre, mg/dl Dialysis before transplant Duration of dialysis, months eGFR before transplant Duration of eGFR <30, days | SCS liver, min SCS kidney, min HMP kidney, h Total CIT kidney (SCS + HMP) Peri-transplant CVVH, yes/no Venovenous bypass, yes/no Transplant technique Operative time (LT), min Operative time (KT), min Blood loss (LT), ml | Peak AST/ALT Peak bilirubin, mg/dl DGF kidney, yes/no UOP <40 ml (within 24 h after KT) ICU stay, days Hospital stay, days s-creatinine, mg/dl 1 week 1 month 3 months |
| Case no. 1 | 44; male 28 kg/m ² Head trauma 0.96 46% | 54; female 23 kg/m ² 21 7840 g NA Yes NA NA NA NA | Iransrusion requirements, units 480 min 510 min 53 h, 30 min Yes Yes Starzl technique 560 min 165 min 2200 ml 4 RBC 5 FFP | 2830/3108 4.2 mg/dl No 9 days 2.12 mg/dl 1.78 mg/dl 1.2 mg/dl |
| Case no. 2 | 54 years; female Stroke 27 kg/m ² 1.21 69% | 40 years; female 20 kg/m ² 12 9860 g 2.64 mg/dl No NA 180 days | 1 platelets 520 min 380 min 46 h 52 h, 20 min Yes Yes Starzl technique 600 min 180 min 5400 ml 14 RBC 11 FFP 1 platelets | 535/680 5.8 mg/dl No 5 days 32 days 1.87 mg/dl 1.25 mg/dl 1.25 mg/dl |
| BMI, body mass mated glomerular ney transplantatio | index; CIT, cold ischemia time; CV filtration rate; HMP, hypothermic m. n; LT, liver transplantation; MELD, mo | VH, continuous venovenous hemoc achine perfusion; ICU, intensive care odel for end-stage liver disease; NA, r | Jialysis; DGF, delayed graft function; unit; KDPI, kidney donor profile index; not available; RBC, red blood cell; SCS, | FFP, fresh frozen plasma; eGFR, esti- KDRI, kidney donor risk index; KT, kid- static cold storage; UOP, urine output. |

Letter to the Editors

Conflicts of interest

The authors have declared no conflicts of interest.

Informed consent

Informed written consent to publish these clinical cases were obtained from the patients.

REFERENCES

- Hibi T, Sageshima J, Molina E, et al. Predisposing factors of diminished survival in simultaneous liver/kidney transplantation. Am J Transplant 2012; 12: 2966.
- Ekser B, Mangus RS, Kubal CA, et al. Excellent outcomes in combined liverkidney transplantation: impact of kidney donor profile index and delayed

kidney transplantation. *Liver Transpl* 2018; **24**: 222.

- 3. Ekser B, Chen AM, Kubal CA, *et al.* Delayed kidney transplantation after 83 hours of cold ischemia time in combined liver-kidney transplant. *Transplantation* 2019. [Epub ahead of print]
- 4. Gedaly R, Guidry P, Davenport D, *et al.* Peri-operative challenges and long-term

outcomes in liver transplantation for polycystic liver disease. *HPB (Oxford)* 2013; **15**: 302.

 van Keimpema L, Nevens F, Adam R, et al. Excellent survival after liver transplantation for isolated polycystic liver disease: an European Liver Transplant Registry study. *Transpl Int* 2011; 24: 1239.