



HAL
open science

Reply to: “Combined heart-liver transplantation for congestive hepatopathy with bridging fibrosis: Is it warranted?”

Manon Allaire, Anna Sessa, Jean François Cadranel, Pascal Lebray

► To cite this version:

Manon Allaire, Anna Sessa, Jean François Cadranel, Pascal Lebray. Reply to: “Combined heart-liver transplantation for congestive hepatopathy with bridging fibrosis: Is it warranted?”: Combined Heart-Liver Transplantation: A need to identify the good candidates beyond liver fibrosis classification?. *JHEP Reports Innovation in Hepatology*, 2021, pp.100327. 10.1016/j.jhepr.2021.100327 . hal-03282352

HAL Id: hal-03282352

<https://hal.sorbonne-universite.fr/hal-03282352>

Submitted on 9 Jul 2021

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L'archive ouverte pluridisciplinaire **HAL**, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d'enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

Journal Pre-proof

Combined Heart-Liver Transplantation: A need to identify the good candidates beyond liver fibrosis classification ?

Manon Allaire, Anna Sessa, Jean François Cadranel, Pascal Lebray



PII: S2589-5559(21)00103-8

DOI: <https://doi.org/10.1016/j.jhepr.2021.100327>

Reference: JHEPR 100327

To appear in: *JHEP Reports*

Received Date: 6 May 2021

Revised Date: 22 June 2021

Accepted Date: 28 June 2021

Please cite this article as: Allaire M, Sessa A, Cadranel JF, Lebray P, Combined Heart-Liver Transplantation: A need to identify the good candidates beyond liver fibrosis classification ?, *JHEP Reports* (2021), doi: <https://doi.org/10.1016/j.jhepr.2021.100327>.

This is a PDF file of an article that has undergone enhancements after acceptance, such as the addition of a cover page and metadata, and formatting for readability, but it is not yet the definitive version of record. This version will undergo additional copyediting, typesetting and review before it is published in its final form, but we are providing this version to give early visibility of the article. Please note that, during the production process, errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

© 2021 Published by Elsevier B.V. on behalf of European Association for the Study of the Liver (EASL).

Combined Heart-Liver Transplantation: A need to identify the good candidates beyond liver fibrosis classification ?

Manon Allaire^{1,2}, Anna Sessa^{1,3}, Jean François Cadranel⁴, Pascal Lebray¹

1. Sorbonne Université, Service d'Hépatologie, Hôpitaux *Universitaires* Pitié Salpêtrière - Charles Foix, AP-HP, Paris, France.
2. Inserm U1149, Centre de Recherche sur l'Inflammation, France Faculté de Médecine Xavier Bichat, Université Paris Diderot, Paris, France.
3. Department of Hepatology and Gastroenterology, Policlinico Federico II, Napoli, Italy
4. Service d'Hépatogastroentérologie de nutrition et d'Alcoologie, Groupe Hospitalier Public du Sud de l'Oise, Creil France

Correspondance and reprints to:

Dr Manon ALLAIRE
Service d'hépatogastroentérologie et oncologie
Groupe Hospitalier Pitié-Salpêtrière
47-83 Boulevard de l'Hôpital
75013 Paris, France
tél: 01.42.16.10.34 - fax: 01.42.16.11.60
mail: manon.allaire@aphp.fr

Acknowledgement : Non

Conflict of interests' statement: nothing to declare regarding this work

Financial support: none

Number of tables: 0

Number of figures: 1

Electronic word count: 626 words excluding references

Author contributions:

Study concept and design: Allaire, Cadranel, Lebray and Sessa

Drafting of the manuscript: Allaire, Cadranel, Lebray and Sessa

Final approval of the version to be submitted : Allaire, Cadranel, Lebray and Sessa

Abbreviations:

CLHT: combined liver heart transplant

HCC: hepatocellular carcinoma

HT: heart transplant

Journal Pre-proof

We thank Izzy et al. (1) for their interest in our article about the management of congestive hepatopathy (2), but also for their participation in improving the knowledge of this disease by identifying diabetes as an independent predictive factor of graft failure and mortality after combined heart-liver transplantation (CHLT) (3). We fully agree that careful selection of the patients with congestive hepatopathy who will benefit from CHLT is mandatory in the era of organ shortage and with regard to the survival observed after such a procedure. However, data are sparse as CHLT represent less than 10% of combined transplantation, 20-30 cases in the USA per year and less than 10 per year in France. Therefore, the identification of good candidates remains a major issue. In most cases, CHLT in the United States has been performed in patients with familial amyloid neuropathy or congenital heart diseases, dealing with a younger population of patients. In these series, indications for CHLT were mainly based on indirect criteria of cirrhosis with explants finding a low proportion of true cirrhosis, which can contribute to the good outcome of these patients (4). Data are missing regarding patients with mixed liver lesions such as acute or chronic congestive lesions in combination with chronic advanced liver disease linked to metabolic syndrome, alcohol, or chronic viral hepatitis, a specific population of patients that constitute the majority of our French cohort (5). They present an increased risk of liver complications following heart transplantation (HT) alone, justifying the consideration of CHLT in case of cirrhosis or even bridging fibrosis (especially for those with liver failure), and hepatocellular carcinoma (HCC).

Itself, the term “bridging cirrhosis” needs further discussion as accurate evaluation for fibrosis stage remains challenging in the setting of end-stage heart failure. The diagnostic performance of non-invasive tests remains low, and individual variability exists when performing a liver biopsy. Moreover, a high risk of complications including bleeding must be considered while indicating biopsy when no specific anatomopathological classification exists with low diagnostic performance. However, specific attention to periportal or perisinusoidal fibrosis appears important to prompt an investigation for the presence of another cause of liver disease which may favor fibrosis progression through cirrhosis and its related complications.

Even when the accurate classification of liver fibrosis in HT candidates is available, at least three questions persist:

- Challenging reports showing regression of fibrosis was noticed after HT alone, one for Fontan associated liver disease and the other for idiopathic dilated cardiomyopathy (6,7),
- To date, no study is available about the outcome of HT for patients with real cirrhosis, while survival results are similar between CHLT and single HT.
- Further analysis of the influence of preoperative acute liver failure (Delta MELD) and postoperative risk factors of fibrosis progression and HCC is needed to identify the patients who will benefit from CHLT or HT alone. Interestingly, MELD score and ascites were independently associated with low postoperative survival in patients who underwent single HT and who did not present cirrhosis. In this series, dilated and ischemic cardiopathies were the main indication for HT (5).

Consequently, decision-making in the context of F3 or even F3-F4 fibrosis, while respecting a definition based on histological criteria, can only be done in the context of a multi-collegial and multidisciplinary decision considering the presence of coexisting acute and chronic liver disease and risk factors of fibrosis liver progression.

References

1. Izzy M, Alexopoulos S, Shinina A. Combined Heart-Liver Transplantation For Congestive Hepatopathy with bridging Fibrosis: Is it warranted. *JHEPReport*. 2021 doi.org/10.1016/j.jhepr.2021.100292
2. Sessa A, Allaire M, Lebray P, Medmoun M, Tiritilli A, Iaria P, et al. From congestive hepatopathy to hepatocellular carcinoma, how can we improve patient management? *JHEPReport*. 2021 DOI:<https://doi.org/10.1016/j.jhepr.2021.100249>
3. Lee S, Matsuoka L, Cao S, Groshen S, Alexopoulos SP. Identifying Predictors of Outcomes in Combined Heart and Liver Transplantation. *Transplant Proc*. 2019;51(6):2002-8.
4. Rizvi SSA, Challapalli J, Maynes EJ, Weber MP, Choi JH, O'Malley TJ, et al. Indications and outcomes of combined heart-liver transplant: A systematic review and met-analysis. *Transplantation Reviews*. 2020;34(2):100517.
5. Lebray P, Varnous S, Pascale A, Leger P, Luyt CE, Ratzu V, et al. Predictive value of liver damage for severe early complications and survival after heart transplantation: A retrospective analysis. *Clin Res Hepatol Gastroenterol*. 2018;42(5):416-26.
6. Bouchardy Judith, Meyer Philippe, Yerly Patrick, Blanche Coralie, Hullin Roger, Giostra Emiliano, et al. Regression of Advanced Liver Fibrosis After Heart Transplantation in a Patient With Prior Fontan Surgery for Complex Congenital Heart Disease. *Circulation: Heart Failure*. 2018;11(11):e003754.
7. Crespo-Leiro MG, Robles O, Paniagua MJ, Marzoa R, Naya C, Flores X, et al. Reversal of Cardiac Cirrhosis Following Orthotopic Heart Transplantation. *American Journal of Transplantation*. 2008;8(6):1336-9.