



**Stanford**  
MEDICINE

# Surgical Management of Hepatocellular Carcinoma

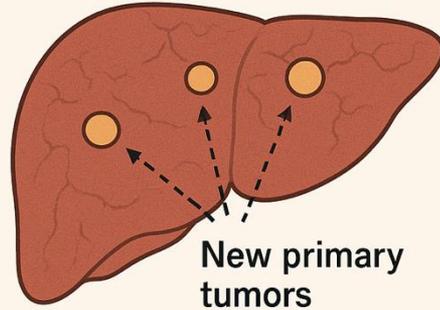
## What is the Role of Liver Transplantation?

**Kazunari Sasaki, MD**

# What makes HCC surgical treatment distinct?

## Patterns of Tumor Recurrence After Liver Resection for Hepatocellular Carcinoma

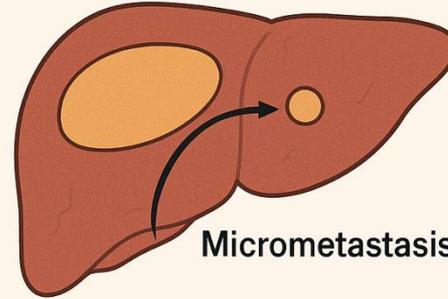
### Multicentric Recurrence



New primary tumors

No direct link to original tumor

### Recurrence from Resected Tumor (Intrahepatic Metastasis)



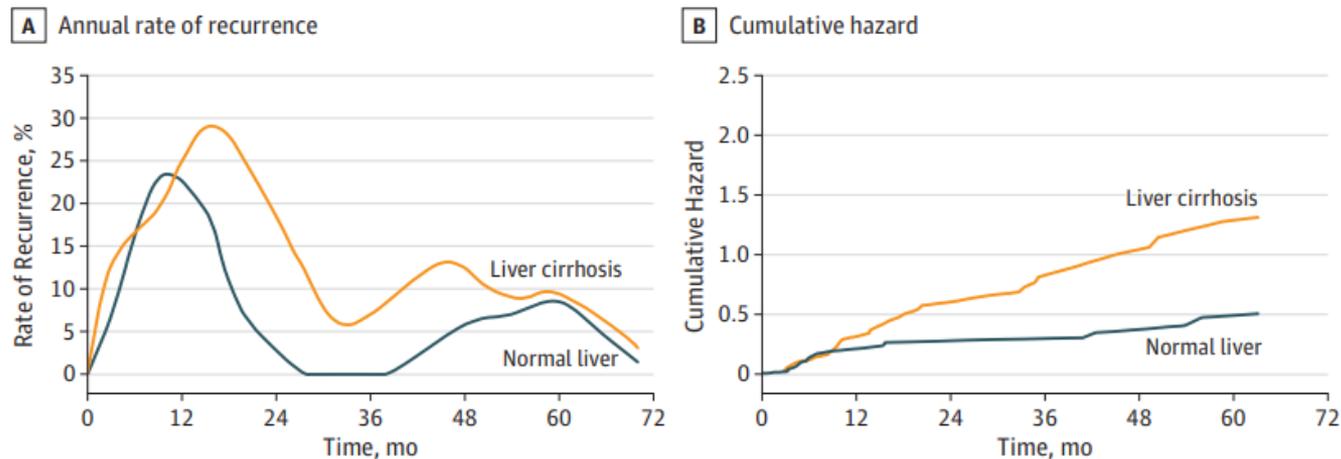
Micrometastasis

Recurring tumor  
Same clonal origin

# Effect of Background Liver Cirrhosis on Outcomes of Hepatectomy for Hepatocellular Carcinoma

Kazunari Sasaki, MD; Junichi Shindoh, MD, PhD; Georgios A. Margonis, MD, PhD; Yujiro Nishioka, MD; Nikolaos Andreatos, MD; Akinari Sekine, MD; Masaji Hashimoto, MD, PhD; Timothy M. Pawlik, MD, MPH, PhD

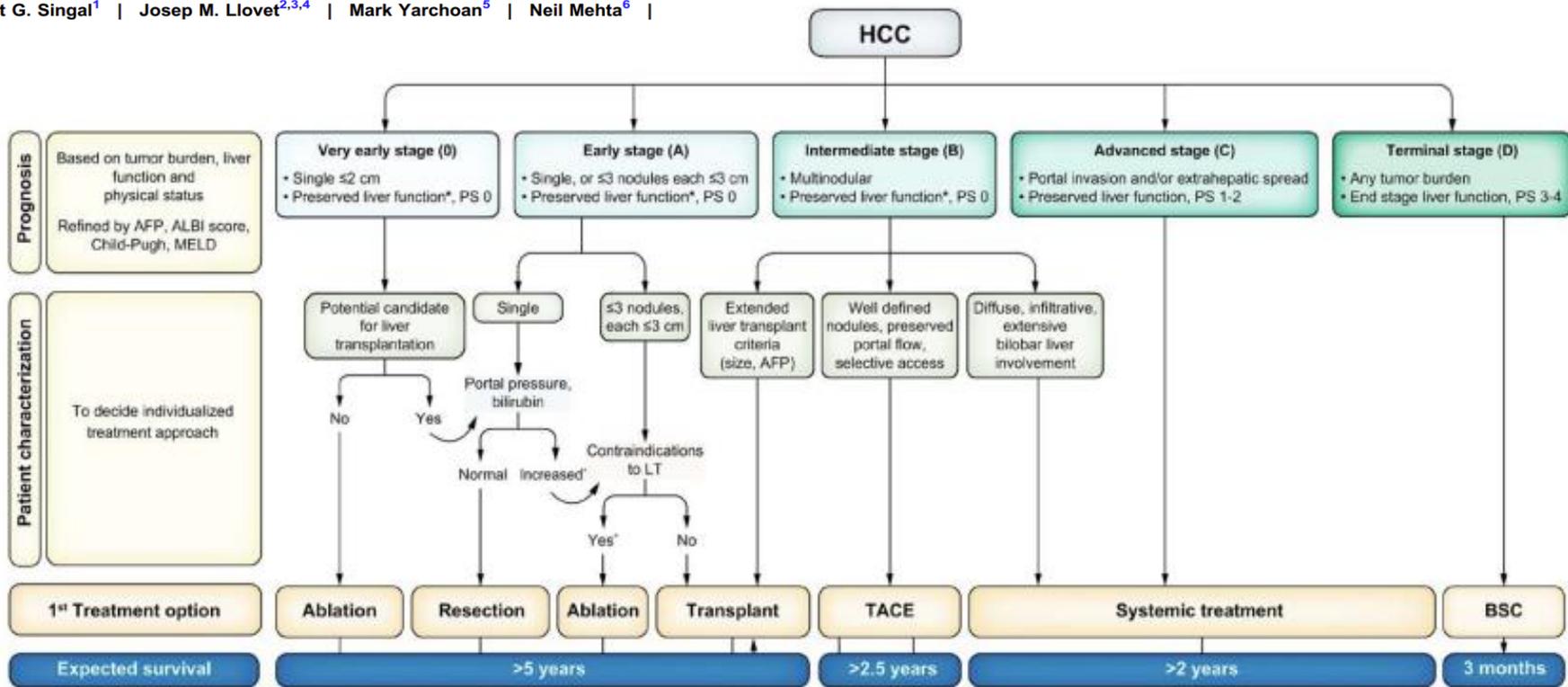
Figure 4. Annual Recurrence Rate and Cumulative Hazard of Recurrence



The recurrence rate among patients in the LC group remained consistently **6% to 15%** higher than that in the NL group

# AASLD Practice Guidance on prevention, diagnosis, and treatment of hepatocellular carcinoma

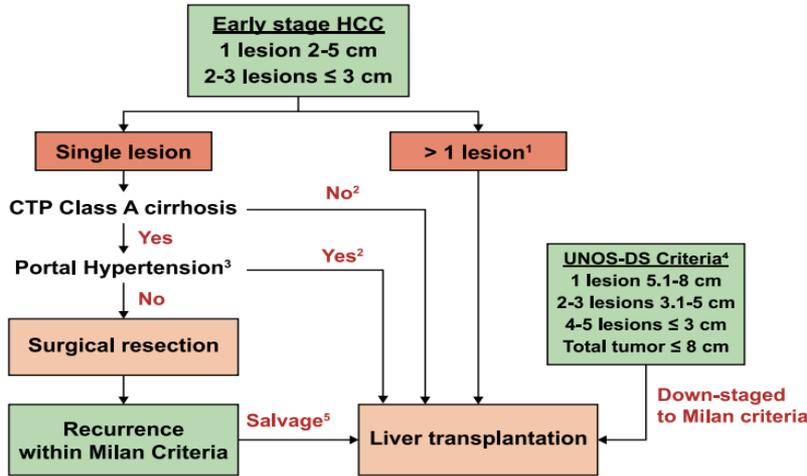
Amit G. Singal<sup>1</sup> | Josep M. Llovet<sup>2,3,4</sup> | Mark Yarrow<sup>5</sup> | Neil Mehta<sup>6</sup> |



# AASLD Practice Guidance on prevention, diagnosis, and treatment of hepatocellular carcinoma

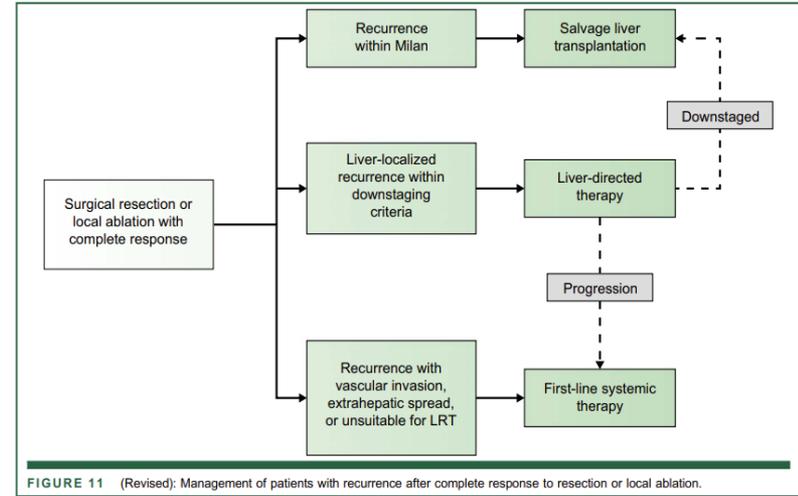
Amit G. Singal<sup>1</sup> | Josep M. Llovet<sup>2,3,4</sup> | Mark Yarchoan<sup>5</sup> | Neil Mehta<sup>6</sup> |

## Algorithm for surgical treatment of early stage HCC



**FIGURE 10** Algorithm for surgical treatment of early-stage hepatocellular carcinoma (HCC). Abbreviations: CTP, Child-Turcotte-Pugh; UNOS-DS, United Network for Organ Sharing Down-Staging.

<sup>1</sup>In non-liver transplant (LT) candidate, can consider surgical resection



**FIGURE 11** (Revised): Management of patients with recurrence after complete response to resection or local ablation.

1 lesion >5 cm and ≤8 cm, OR 2–3 lesions, at least one >3 cm and ≤5 cm, with total tumor diameter ≤8 cm, OR 4–5 lesions, all ≤3 cm, with total tumor diameter ≤8 cm, No macrovascular invasion or extrahepatic spread must be present.

# Allocation policy

Year	Policy Update	Description
2002	MELD Implementation	MELD system replaces CTP for liver allocation. T2 HCC patients assigned <b>MELD 29</b> ; T1 patients assigned <b>MELD 24</b> (exception points).
2003	Point Adjustment	T2 reduced to <b>MELD 24</b> , T1 reduced to <b>MELD 20</b> .
2004	No Exception for T1	T1 HCC lesions no longer eligible for MELD exception.
2005	T2 MELD Exception Reduced	T2 lesion MELD exception lowered to <b>22</b> .
2015	6-Month Delay Introduced	Patients must wait <b>6 months</b> after listing before exception points are granted. Aimed to assess tumor biology.
2019	MMaT-3 Policy	MELD exception based on <b>Median MELD at Transplant - 3 points</b> in transplant center's region, increasing equity.
2020	Standardized Review	Creation of <b>National Liver Review Board (NLRB)</b> for consistent review of HCC exceptions.
2023	LI-RADS Imaging Requirement	Imaging for exception requests must follow <b>LI-RADS</b> criteria. Enhances standardization and diagnostic accuracy.



# Major transformations in the US LTx environment

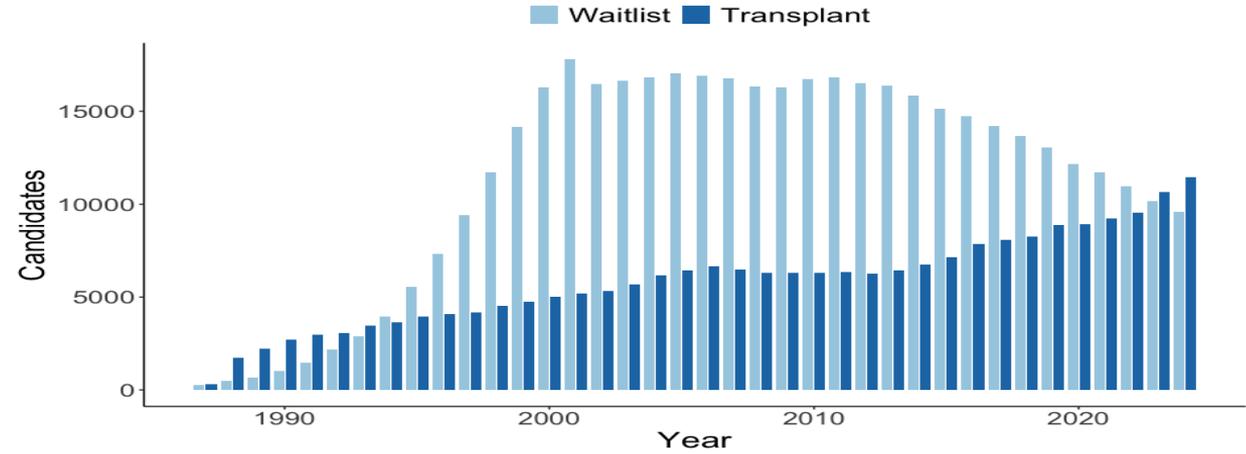
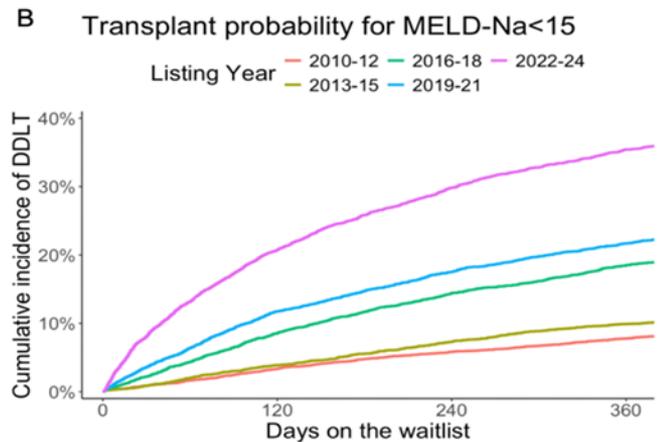
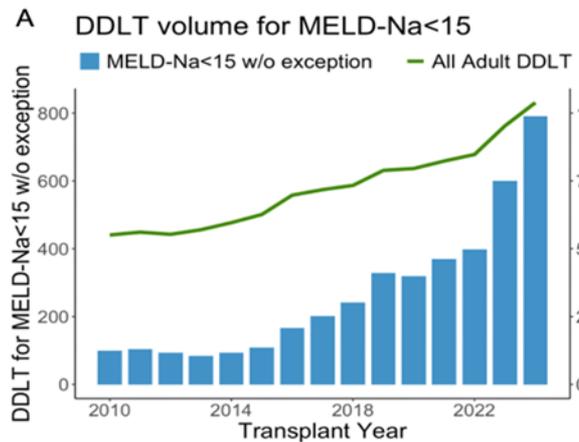
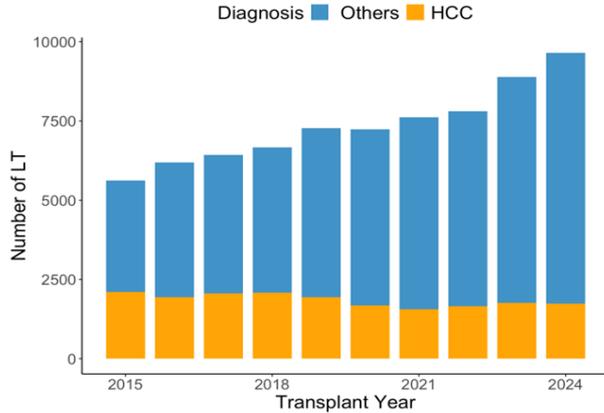


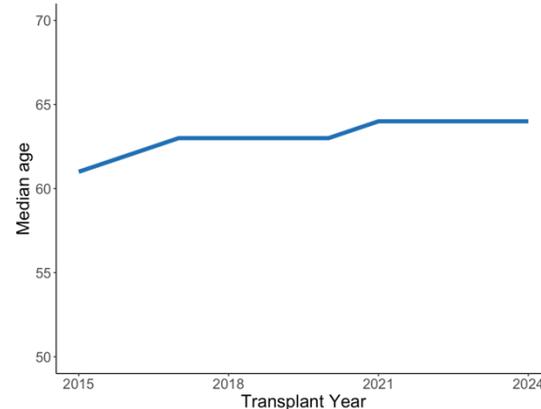
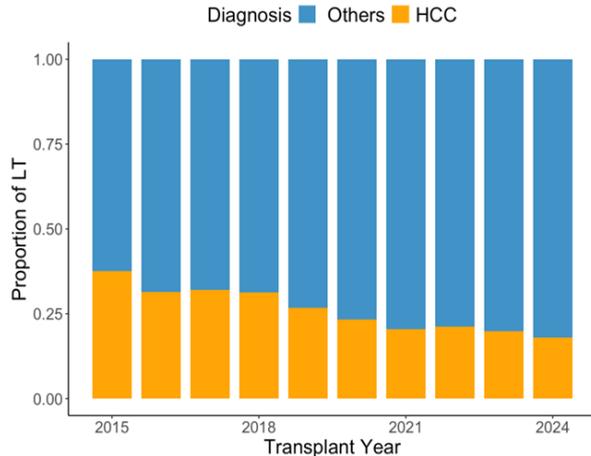
Figure 1



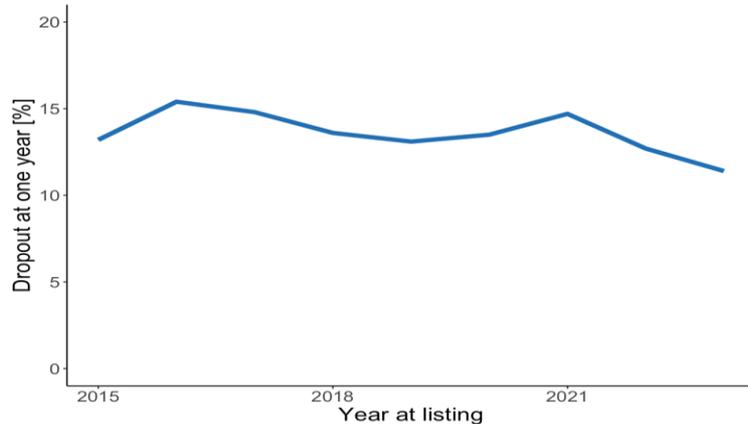
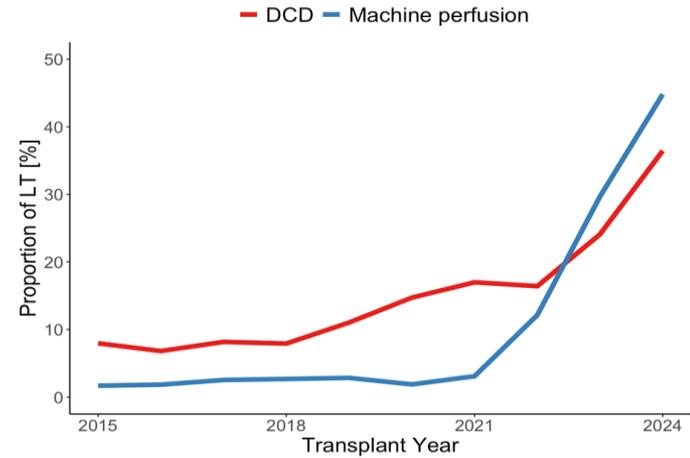
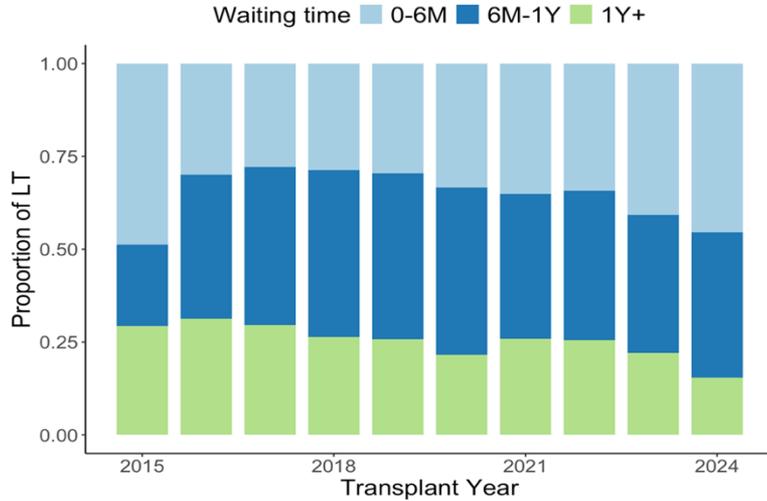
# Is there an increase in transplants for HCC?



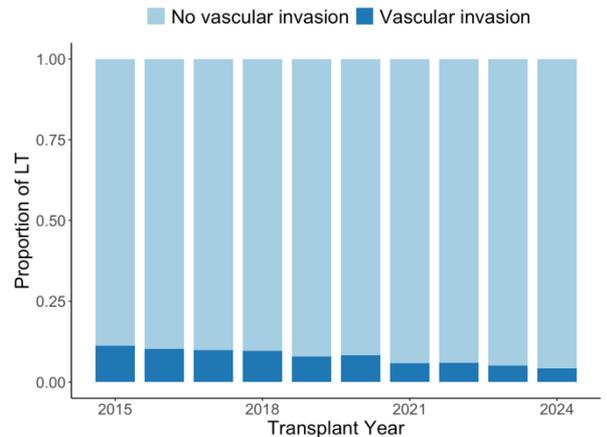
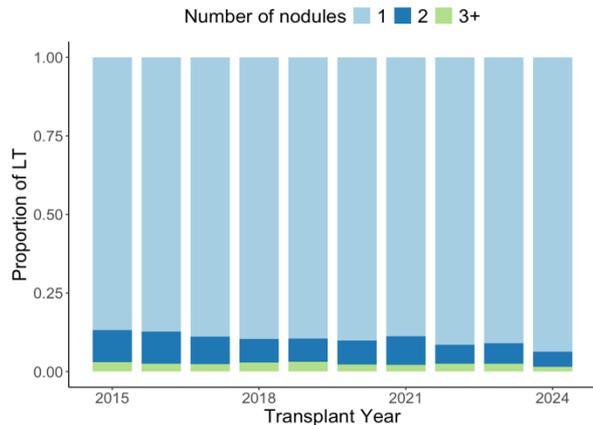
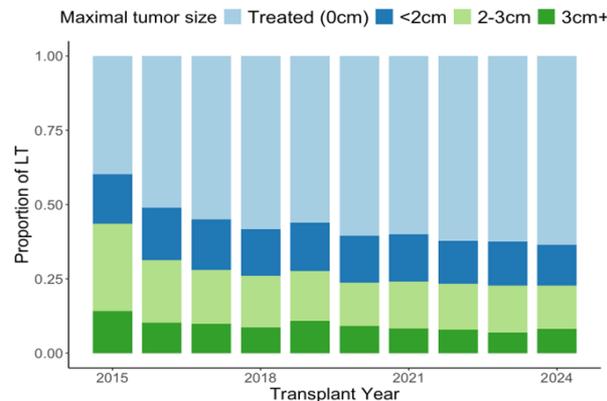
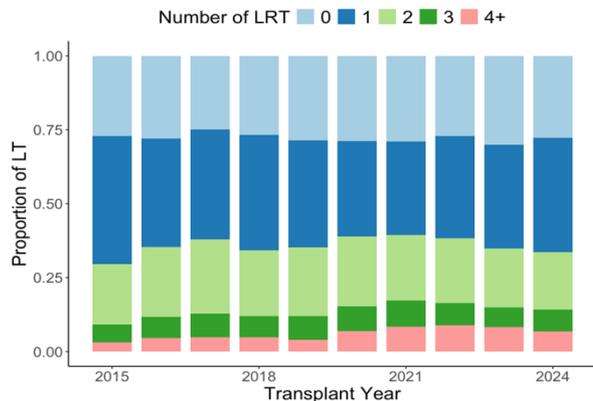
**Absolute/percentage decreased**  
**Median Pt age increased significantly**



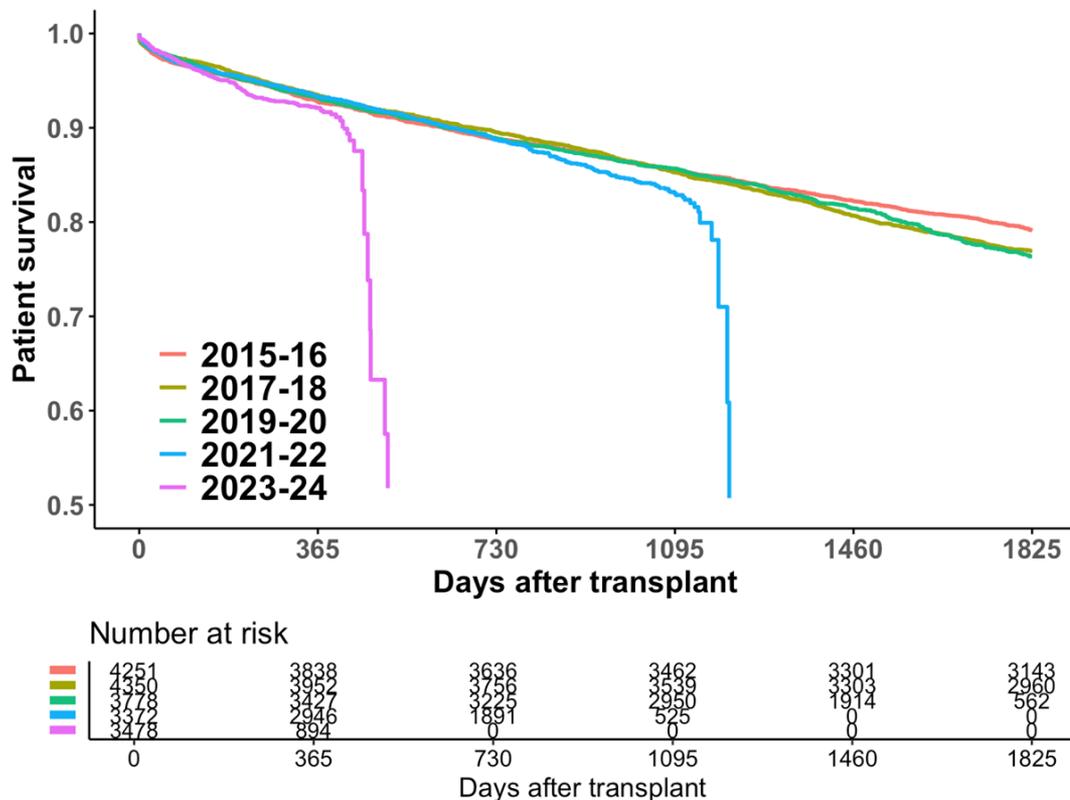
# Does Transplant for HCC increase ?



# Transplant for HCC tumor characteristics



# Post Transplant Survival



**1-year: 93-94%**  
**3-year: 86%**  
**5-year: 76-79%**

OS	1yr	3yr	5yr
2015-16	93.7	86.7	79.2
2017-18	94.6	86.1	76.9
2019-20	93.5	86.2	76.4
2021-22	94.3	83.9	
2023-24	92.2		



# Time to Expand Selection Criteria for MELD Exception Points in Liver Transplantation for Hepatocellular Carcinoma

Chase J Wehrle, MD,<sup>1</sup> Jiro Kusakabe, MD, PhD, MPH,<sup>2</sup> Toshihiro Nakayama, MD,<sup>3</sup> Charles Miller, MD,<sup>1</sup> Koji Hashimoto, MD, PhD,<sup>1</sup> Timothy M. Pawlik, MD, PhD, MPH,<sup>4</sup> Kazunari Sasaki, MD, PhD,<sup>3</sup> Vincenzo Mazzaferro, MD, PhD,<sup>5</sup> Andrea Schlegel, MD, MPH,<sup>1</sup> and Federico Aucejo, MD<sup>1</sup>

## Drawbacks of Milan Criteria based system

### Milan Criteria: **Low prediction power**

- Made in 1996 using 1991-1994 patients
- Only assesses tumor morphology
- Originally generated by pathology specimen
- Binary approach (yes/no)
- Increasing penetration of LRT

### Exception point: **Too arbitrary**

- No supportive scientific evidence
- Does not reflect individual risk of drop-out
- Cannot balance between HCC patients and non-HCC patients



# Development and validation of the HALT-HCC score to predict mortality in liver transplant recipients with hepatocellular carcinoma: a retrospective cohort analysis

Kazunari Sasaki\*, Daniel J Firl\*, Koji Hashimoto, Masato Fujiki, Teresa Diago-Usó, Cristiano Quint  
Federico N Aucejo, Charles M Miller

THE LANCET  
Gastroenterology & Hepatology  
Volume 2, Issue 8, August 2017, Pages 595-603

## HEPATOLOGY

Charting the Path Forward for Risk Prediction in Liver Transplant for Hepatocellular Carcinoma: International Validation of HALTHCC Among 4,08 Patients

Clinical Gastroenterology and Hepatology 2024;22:2044–2052

### HEPATOLOGY

Continuous Risk Score Predicts Waitlist and Post-transplant Outcomes in Hepatocellular Carcinoma Despite Exception Changes

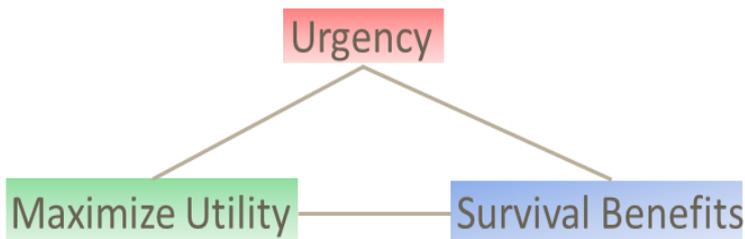


## HEPATOLOGY

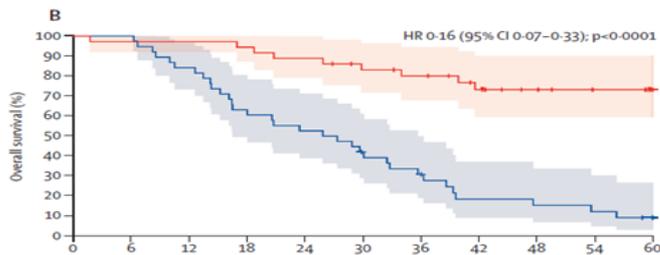
Reframing the Approach to Patients With Hepatocellular Carcinoma: Longitudinal Assessment With Hazard Associated With Liver Transplantation for HCC (HALTHCC) Improves Ablate and Wait Strategy

Continuous score which can predict both wait list mortality and post LT outcomes

# Survival benefit by Liver Transplant



Liver transplantation plus chemotherapy versus chemotherapy alone in patients with permanently unresectable colorectal liver metastases (TransMet): results from a multicentre, open-label, prospective, randomised controlled trial

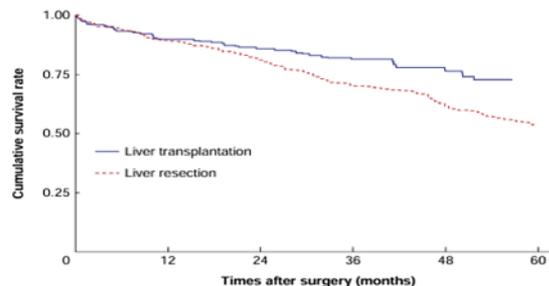


Number at risk (number censored)	0	6	12	18	24	30	36	42	48	54	60
Chemotherapy only	38 (0)	38 (0)	32 (0)	24 (0)	20 (0)	15 (1)	11 (1)	6 (2)	5 (2)	4 (2)	2 (3)
LT plus chemotherapy	36 (0)	35 (0)	35 (0)	34 (0)	32 (0)	28 (2)	26 (3)	21 (6)	17 (10)	14 (13)	10 (17)



## Liver transplantation for elderly patients with early-stage hepatocellular carcinoma

Yutaka Endo<sup>1</sup>, Kazunari Sasaki<sup>2</sup>, Zorays Moazzam<sup>1</sup>, Henrique A. Lima<sup>1</sup>, Laura Alaïmo<sup>1</sup>, Muhammad Musaab Munir<sup>1</sup>, Chanza F. Shaikh<sup>1</sup>, Austin Schenk<sup>1</sup>, Minoru Kitago<sup>3</sup> and Timothy M. Pawlik<sup>1\*</sup>



No. at risk	0	12	24	36	48	60
Transplantation	215	187	159	125	98	78
Resection	1907	1613	1323	967	671	454

Fig. 4 Kaplan-Meier survival curves after propensity score overlap weighting adjustment for elderly patients with stage I-II hepatocellular carcinoma undergoing liver transplantation versus surgical resection