

## Understanding Surgical Decision Making in Early Hepatocellular Carcinoma

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### A B S T R A C T

#### Purpose

The choice between liver transplantation (LT), liver resection (LR), and radiofrequency ablation (RFA) as initial therapy for early hepatocellular carcinoma (HCC) is controversial, yet little is known about how surgeons choose therapy for individual patients. We sought to quantify the impact of both clinical factors and surgeon specialty on surgical decision making in early HCC by using conjoint analysis.

#### Methods

Surgeons with an interest in liver surgery were invited to complete a Web-based survey including 10 case scenarios. Choice of therapy was then analyzed by using regression models that included both clinical factors and surgeon specialty (non-LT v LT).

#### Results

When assessing early HCC occurrences, non-LT surgeons (50% LR; 41% LT; 9% RFA) made significantly different recommendations compared with LT surgeons (63% LT; 31% LR; 6% RFA;  $P < .001$ ). Clinical factors, including tumor number and size, type of resection required, and platelet count, had significant effects on the choice between LR, LT, and RFA. After adjusting for clinical factors, non-LT surgeons remained more likely than LT surgeons to choose LR compared with LT (relative risk ratio [RRR], 2.67). When the weight of each clinical factor was allowed to vary by surgeon specialty, the residual independent effect of surgeon specialty on the decision between LR and LT was negligible (RRR, 0.93).

#### Conclusion

The impact of surgeon specialty on choice of therapy for early HCC is stronger than that of some clinical factors. However, the influence of surgeon specialty does not merely reflect an across-the-board preference for one therapy over another. Rather, certain clinical factors are weighed differently by surgeons in different specialties.

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### INTRODUCTION

Hepatocellular carcinoma (HCC) is the sixth most common cancer and, because of its poor prognosis, the third leading cause of cancer death worldwide.<sup>1</sup> Surgical therapy offers the only means of cure, but the choice of appropriate surgical therapy for HCC is controversial, especially for early HCC in the setting of well-compensated cirrhosis.<sup>2-5</sup> Multiple studies have demonstrated that acceptable long-term survival outcomes can be achieved in appropriately selected patients by using liver transplantation (LT),<sup>6-10</sup> liver resection (LR),<sup>11-13</sup> or radiofrequency ablation (RFA),<sup>14-16</sup> and all of these options are used in clinical practice. Furthermore, surgeons who treat HCC come from a variety of training backgrounds and may perform only a subset of these procedures. As such, both clinical factors

and surgeon specialty likely impact the choice of surgical therapy in early HCC. Although previous series have reported outcomes for treatment of early HCC with LT, LR, and RFA, no study has addressed how surgeons use clinical data and their own experience to choose the most appropriate surgical therapy for individual patients.

Conjoint analysis, a technique developed in mathematical psychology<sup>17</sup> and then used widely in marketing research,<sup>18,19</sup> allows healthcare decision making to be systematically studied.<sup>20,21</sup> This technique consists of a choice experiment: key attributes of sample cases are varied, and the variation in respondents' choices in relation to these attributes can be quantified. Conjoint analysis has been used previously to study surgical decision making.<sup>22-30</sup> We sought to use conjoint analysis to elucidate the factors that drive surgical decision making in early

HCC. Our objectives were to characterize the therapeutic preferences of surgeons who treat HCC, to quantify the impact of clinical factors on the choice of surgical therapy, and to understand the interplay between surgeon specialty and clinical factors in determining choice of therapy for early HCC. We hypothesized that both clinical factors and surgeon factors would independently predict choice of therapy.

## METHODS

### Survey Instrument Design

For purposes of this study, early HCC was defined as HCC within the Milan criteria for LT (ie, single tumor < 5 cm or 2 to 3 tumors all < 3 cm, with no evidence of extrahepatic tumor)<sup>6</sup> in the setting of well-compensated cirrhosis (ie, Child-Pugh class A, no varices, no ascites, and no encephalopathy). Standard conjoint analysis methodology was used to design, implement, and analyze the survey in five steps: defining attributes, assigning attribute levels, creating scenarios, obtaining preference data, and estimating model parameters.<sup>19</sup> First, structured interviews were conducted with five academic and community surgeons who treat HCC to identify key clinical factors influencing choice of therapy as well as to determine a clinically meaningful range of values for each factor. Seven key clinical factors were thus identified and were then incorporated into 36 case scenarios by using a fractional factorial design that is based on an orthogonal array.<sup>31</sup> A random subset of 10 case scenarios was chosen for each respondent. In each case, the respondent was asked which procedure he would recommend as initial curative-intent therapy: LR, RFA, LT without bridging therapy (LT – B), or LT with bridging therapy (LT + B). Bridging therapy is the use of an adjunct modality, such as chemoembolization, to reduce the risk of tumor progression while waiting for a transplantation. In addition to the case scenarios, the survey instrument included questions on practice characteristics and on attitudes toward surgical therapies for early HCC.

The resulting survey instrument was then pilot tested and iteratively refined. Survey invitations were e-mailed to surgeons who had an interest in liver surgery (including hepatobiliary surgery and liver transplantation); e-mail addresses were obtained from publicly available sources, such as Web sites and publications.<sup>32</sup> Prespecified eligibility criteria specified that respondents must be practicing surgeons who had completed surgical training and who evaluated at least five patients with HCC per year.

### Statistical Analysis

Descriptive statistics were compared by using Fisher's exact test or the rank-sum test, as appropriate. Choice data were analyzed by using multinomial logistic regression models with robust variance estimates, yielding relative risk ratios (RRRs) that reflected the change in the probability of choosing a particular therapy over an alternative.<sup>33</sup> It was specified a priori that the LT – B and LT + B options would be analyzed both as a combined choice and as separate choices. Interaction terms were included in an all-or-none fashion to assess differences in decision making between groups of respondents. All tests of statistical significance were two sided, and statistical significance was established at  $P < .05$ . Statistical analyses were performed with Stata/MP 10.1 for Windows (StataCorp LP, College Station, TX). The study protocol was deemed exempt from review by the Johns Hopkins Bloomberg School of Public Health institutional review board.

## RESULTS

A total of 1,032 e-mail invitations were sent; 336 eligible and complete responses were received (response rate, 33%). Practice characteristics of the respondents are given in Table 1. Approximately half (54%) of respondents indicated that they currently performed LT for HCC, and these data were used to designate surgeon specialty (non-LT or LT). Of the 155 non-LT surgeons, 149 performed LR, and 130 performed LR

**Table 1.** Practice Demographics and Clinical Characteristics of Survey Respondents

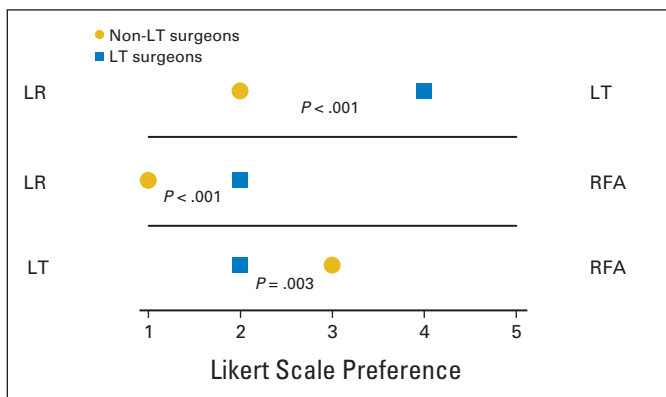
Characteristic	No. of Patients (N = 336)	%
<b>Practice type</b>		
Academic practice or university hospital staff	284	85
Private practice or community hospital staff	52	15
<b>Years in practice</b>		
Median	10	
IQR	4-17	
<b>Fellowship training</b>		
HPB with LT	112	33
HPB without LT	35	10
Surgical oncology	86	26
Transplantation	142	42
None of the above	22	7
<b>No. of patients annually evaluated for HCC</b>		
Median	30	
IQR	20-60	
<b>Perform liver resection for HCC</b>		
Yes	315	94
No	21	6
<b>Perform radiofrequency ablation for HCC</b>		
Yes	267	79
No	69	21
<b>Perform liver transplantation for HCC</b>		
Yes	181	54
No	155	46
<b>Annual liver procedure volumes for HCC*</b>		
<b>Liver resection</b>		
Median	5	
IQR	4-15	
<b>Radiofrequency ablation</b>		
Median	5	
IQR	3-15	
<b>Liver transplantation</b>		
Median	10	
IQR	8-20	
<b>HCC cases discussed at multidisciplinary tumor board</b>		
Yes	311	93
No	25	7

Abbreviations: IQR, interquartile range; HPB, hepato-pancreato-biliary; LT, liver transplantation; HCC, hepatocellular carcinoma.

\*For non-zero volumes only.

as well as RFA for HCC. Of the 181 LT surgeons, 166 also performed LR, and 130 performed all three therapies. Respondents annually evaluated a median of 30 patients with HCC and annually performed a median of 20 surgical procedures for HCC.

When asked to indicate their general preferences for initial surgical therapy for early HCC on a Likert scale, there were significant differences by surgical specialty (Fig 1). Non-LT surgeons preferred LR rather than both LT and RFA, whereas non-LT surgeons were in aggregate neutral in their preference for LT versus RFA. In contrast, LT surgeons preferred LT rather than LR and RFA; however, LT surgeons were strongly in favor of LR compared with RFA. These differences were also manifested in the choice of therapy in case scenarios. In aggregate, without adjustment for surgical specialty or clinical factors, LT was chosen in 53% of cases (41% LT + B; 12% LT – B); LR, in 40%; and RFA, in 7%. However, there was a significant



**Fig 1.** Preferences for initial therapy for early hepatocellular carcinoma (HCC), stratified by surgeon specialty. (“In general, do you prefer A or B as initial surgical therapy for patients with well-compensated cirrhosis and HCC within the Milan criteria?”) Median response on Likert scale indicated for each group. LR, liver resection; LT, liver transplantation; RFA, radiofrequency ablation.

difference in the choices made by non-LT surgeons (50% LR; 41% LT; 9% RFA) and LT surgeons (63% LT; 31% LR; 6% RFA;  $P < .001$ ). That RFA was uncommonly chosen was consistent with the fact that only 43% of respondents considered RFA to be a potentially curative treatment modality for HCC.

The clinical factors used to generate case scenarios are listed in Table 2. Respondents were asked to assume that all patients had HCC

Table 2. Factors Used in Case Scenarios	
Factor by levels	
Age, years	50 65
Tumor number and size	3 tumors; largest, 2.5 cm 1 tumor; 4.5 cm 1 tumor; 2.5 cm
Type of resection required	Left hemi-hepatectomy (segments 2, 3, and 4); FLR, 60% Right posterior sectionectomy (segments 6 and 7); FLR, 70% Left lateral sectionectomy (segments 2 and 3); FLR, 85%
Etiology of cirrhosis*	Chronic hepatitis C Chronic hepatitis B Past alcohol abuse (abstinent > 1 year)
Biologic MELD score	10 (INR, 1.3; TB, 1.3 mg/dL; SCr, 1.0 mg/dL) 8 (INR, 1.1; TB, 1.3 mg/dL; SCr, 1.0 mg/dL) 6 (INR, 1.0; TB, 1.0 mg/dL; SCr, 1.0 mg/dL)
Platelet count, per $\mu\text{L}$	90,000 150,000
Anticipated waiting time for liver transplantation, months	2 5 8

Abbreviations: FLR, future liver remnant; MELD, model for end-stage liver disease; INR, international normalized ratio; TB, total serum bilirubin; SCr, serum creatinine.  
\*All patients were described as having Child’s A cirrhosis.

An otherwise healthy patient presents with newly diagnosed HCC with the following characteristics:

- age 50 years
- one 4.5-cm tumor
- resection would require left lateral sectionectomy (segments 2 and 3), future liver remnant 85%
- Child’s A cirrhosis resulting from previous past alcohol abuse (abstinent > 1 year)
- biologic MELD score 10 (INR 1.3, serum bilirubin 1.3 mg/dL, serum creatinine 1.0 mg/dL)
- platelet count of 150,000 per microliter
- anticipated wait time for a liver transplant is 2 months

Which procedure would you recommend as initial curative-intent therapy?

- Liver resection
- Radiofrequency ablation
- Liver transplantation (without bridging therapy)
- Liver transplantation (with bridging therapy)

**Fig 2.** Sample case scenario. HCC, hepatocellular carcinoma; MELD, model for end-stage liver disease; INR, international normalized ratio.

within the Milan criteria in the setting of well-compensated cirrhosis. Figure 2 depicts a sample case scenario. In regression analyses, all clinical factors demonstrated statistically significant effects on the choice among LR, LT, and RFA (Table 3). The type of resection required, tumor number and size, and platelet count had the largest effects on choice of therapy. LT was more likely to be recommended for patients who would require a major hepatic resection, for those who had multifocal disease, and for those who had low platelet count. Additional analyses were performed to identify factors important in choice of bridging therapy when LT was chosen. The presence of a solitary 4.5-cm tumor versus a solitary 2.5-cm tumor (RRR, 1.59; 95% CI, 1.13 to 2.23) or versus three tumors  $\leq$  2.5 cm (RRR, 1.85; 95% CI, 1.24 to 2.77) increased the choice of LT + B versus LT – B ( $P < .001$ ). Similarly, longer LT waiting times increased the choice of LT + B v LT – B (5 v 2 months: RRR, 3.70; 95% CI, 2.66 to 5.14; 8 v 2 months: RRR, 11.9; 95% CI, 7.41 to 19.1;  $P < .001$ ). No other clinical factor significantly affected the choice of LT + B versus LT – B.

Additional regression analyses were then performed to assess the impact of surgeon specialty, relative to clinical factors, on choice of therapy. First, a variable denoting surgeon specialty was added to the previously described regression model, allowing for an overall difference in the propensity to choose each therapy (Appendix Table A1, online only). Although the effects of clinical factors did not change in this model compared with the previous model, there was a separate, independent effect of surgeon specialty on choice of therapy ( $P < .001$ ). Non-LT surgeons were significantly more likely than LT surgeons to choose LR rather than LT (RRR, 2.67; 95% CI, 2.00 to 3.55) and RFA rather than LT (RRR, 2.64; 95% CI, 1.62 to 4.30). Surgeon specialty did not impact the choice of LR versus RFA (RRR, 1.01; 95% CI, 0.63 to 1.62). With respect to choice of bridging therapy, non-LT surgeons were significantly less likely than LT surgeons to choose LT + B rather than LT – B (RRR, 0.38; 95% CI, 0.25 to 0.58).

To additionally explore the differences in choice of therapy related to surgeon specialty, we examined a third model that allowed the impact of each clinical factor to vary by surgeon specialty (Figure 3). By doing so, the surgeon specialty effect could be split into two components: one part related to how different specialists weigh clinical

**Table 3.** Determinants of Choice of Therapy: Clinical Factors

Factor	LR v LT		LR v RFA		RFA v LT		P
	RRR	95% CI	RRR	95% CI	RRR	95% CI	
Age, years							
50	Ref		Ref		Ref		.046
65	1.11	0.96 to 1.29	0.78	0.57 to 1.05	1.44	1.06 to 1.95	
Tumor number and size							
3 tumors; largest, 2.5 cm	Ref		Ref		Ref		< .001
1 tumor; 4.5 cm	1.72	1.35 to 2.19	3.23	1.57 to 6.64	0.53	0.26 to 1.07	
1 tumor; 2.5 cm	2.27	1.76 to 2.93	0.40	0.24 to 0.68	5.65	3.48 to 9.16	
Type of resection required							
Left hemi-hepatectomy, FLR 60%	Ref		Ref		Ref		< .001
Right posterior section, FLR 70%	1.54	1.23 to 1.94	1.34	0.89 to 2.02	1.15	0.80 to 1.65	
Left lateral section, FLR 85%	4.19	3.31 to 5.31	5.79	3.61 to 9.30	0.72	0.47 to 1.12	
Etiology of cirrhosis*							
Chronic hepatitis C	Ref		Ref		Ref		< .001
Chronic hepatitis B	1.18	0.95 to 1.47	0.75	0.50 to 1.12	1.58	1.08 to 2.32	
Past alcohol abuse	1.47	1.21 to 1.80	1.08	0.74 to 1.59	1.36	0.94 to 1.97	
Biologic MELD score							
10	Ref		Ref		Ref		< .001
8	1.44	1.18 to 1.77	1.46	0.94 to 2.28	0.99	0.65 to 1.49	
6	1.66	1.37 to 2.02	1.56	1.03 to 2.37	1.06	0.71 to 1.59	
Platelet count, per $\mu\text{L}$							
90,000	Ref		Ref		Ref		< .001
150,000	2.32	1.93 to 2.79	1.97	1.43 to 2.73	1.18	0.87 to 1.59	
Liver transplantation waiting time, months							
2	Ref		Ref		Ref		< .001
5	1.30	1.08 to 1.57	0.98	0.60 to 1.60	1.33	0.83 to 2.11	
8	1.70	1.39 to 2.09	1.20	0.77 to 1.88	1.41	0.93 to 2.15	

Abbreviations: LR, liver resection; LT, liver transplantation; RFA, radiofrequency ablation; RRR, relative risk ratio; Ref, referent; FLR, future liver remnant; MELD, model for end-stage liver disease.

\*All patients were described as having Child's A cirrhosis.

data, and another part that was unrelated to the clinical data presented in our scenarios. Again, the overall impact of surgeon specialty (including both components of the effect) was significant ( $P < .001$ ). This analysis revealed that several clinical factors were weighed differently by non-LT and LT surgeons, most notably type of resection required, etiology of cirrhosis, and LT waiting time (Figure 3, with differences by specialty visualized as the horizontal spread of points). When the weight of each clinical factor was allowed to vary by surgeon specialty in the model, the residual effect of surgeon specialty, independent of clinical factors, on the decision between LR and LT was negligible (RRR, 0.93; 95% CI, 0.40 to 2.16). Similarly, there was no significant residual effect on the decision between LT and RFA (RRR, 3.91; 95% CI, 0.82 to 18.7). These results indicated that the effect of surgeon specialty was completely explained by differences in how the two specialties weigh clinical data.

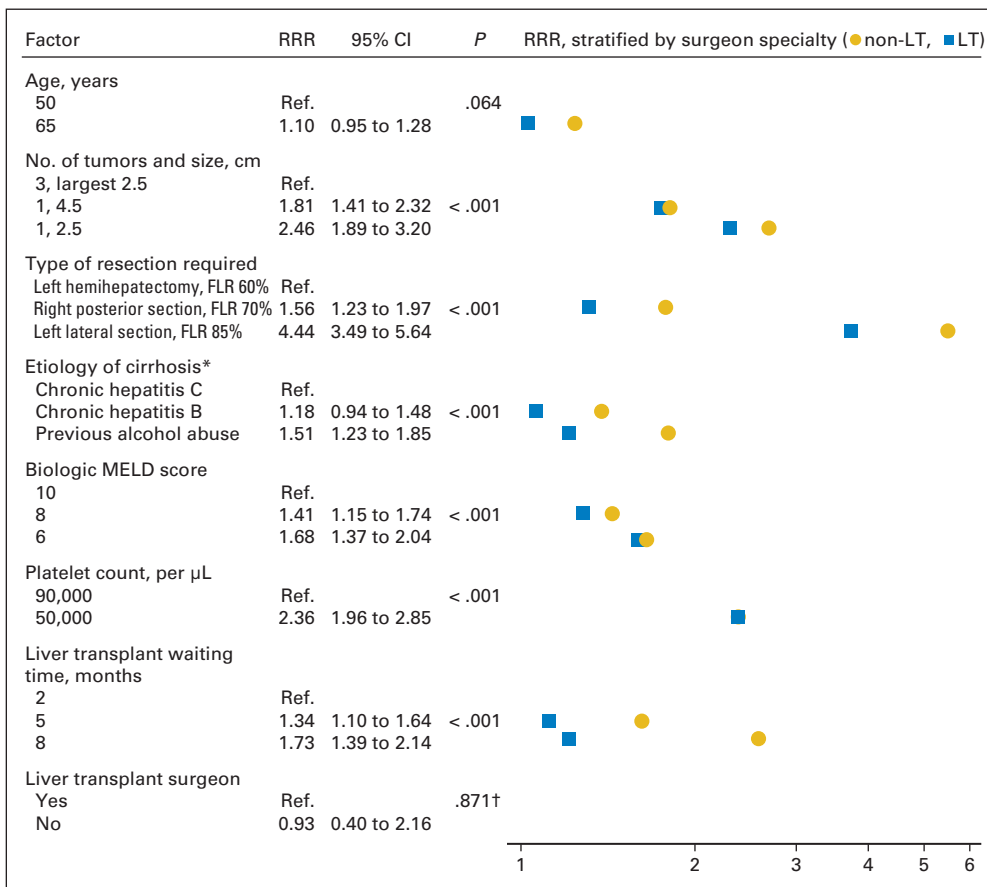
When the choice of LT + B versus LT - B was examined in this fashion, the overall impact of surgeon specialty was again significant ( $P < .001$ ). The impact of LT waiting time varied by surgeon specialty. Non-LT surgeons were sensitive to longer LT waiting times in choosing LT + B v LT - B (5 v 2 months: RRR, 6.77; 95% CI, 4.05 to 11.3; 8 v 2 months: RRR, 20.7; 95% CI, 10.1 to 42.4). In comparison, LT surgeons were relatively less influenced by LT waiting time (LT + B: 5 v 2 months; RRR, 2.32; 95% CI, 1.49 to 3.60; LT - B: 8 v 2 months; RRR, 7.70; 95% CI, 4.01 to 14.8). Notably, even when the weight of each clinical factor was allowed to vary by surgeon specialty, there was still a

residual effect of surgeon specialty on the decision between LT + B and LT - B, with non-LT surgeons favoring bridging therapy one third as often as LT surgeons (RRR, 0.31; 95% CI, 0.13 to 0.75).

Finally, respondents were asked to self-characterize their decision-making strategy for early HCC. Among LT surgeons, 60% reported that they typically decide for or against LT first and then consider other treatment options. Fewer (30%) reported that they consider LR first; 3%, RFA; and 6%, some other strategy. Among non-LT surgeons, 42% reported that they consider LT first; 47%, LR; 3%, RFA; and 7%, some other strategy. Again, there was a significant difference by specialty ( $P = .009$ ).

## DISCUSSION

Variation in choice of therapy in HCC is well documented<sup>34</sup> and likely reflects a collective sense of equipoise regarding the optimal therapy of patients with early HCC.<sup>2-5</sup> Choice of therapy may depend on clinical factors as well as surgeon characteristics. The factors influencing surgeons' choices of therapy for early HCC have not, however, been previously studied. This study is, to our knowledge, the first to analyze surgical decision making in HCC. By using conjoint analysis, we quantified the relative impact of clinical factors on choice of therapy. Interestingly, we also noted a significant impact of surgeon specialty



**Fig 3.** Determinants of choice of liver resection (LR) versus liver transplantation (LT), stratified by and including surgeon specialty. (\*) All patients were described as having Child's A cirrhosis. (†) *P* value for joint significance of all surgeon specialty-related variables < .001. RRR, relative risk ratio; Ref, referent; FLR, future liver remnant; MELD, model of end-stage liver disease.

on decision making, and we were able to quantify the relative contribution of surgeon specialty versus clinical factors to early HCC decision making. In fact, we noted that the impact of surgeon specialty was larger in magnitude than the impact of several clinical factors.

In this study, the choice of surgical therapy for early HCC varied widely, with roughly equal numbers of respondents choosing LR or LT as initial therapy. Unlike in Asia, where RFA is viewed more favorably as a treatment modality for early HCC,<sup>15,16</sup> only 43% of respondents from our American cohort of surgeons considered RFA to be potentially curative for HCC, choosing it in only 7% of cases. Our study identified the need for major hepatic resection, the presence of multifocal disease, and thrombocytopenia as the clinical factors most influencing the choice of LT over LR. The influence of these factors on surgical decision making is consistent with published data that implicate them as important predictors of surgical outcome.<sup>5</sup> For example, more extensive liver resection, especially in the setting of thrombocytopenia and portal hypertension,<sup>35</sup> is associated with increased post-operative morbidity and mortality.<sup>36</sup> Furthermore, LR for multifocal early HCC has been associated with inferior oncologic results.<sup>13</sup> As such, LT may be a better option for patients with these clinical factors.

Of note, surgeon specialty played a role that was at least as important as clinical factors in determining preference for initial therapy. When asked for their general attitudes towards surgery for early HCC, non-LT surgeons indicated a preference for LR, and LT surgeons, for LT. Similarly, analysis of choice data showed that non-LT surgeons were much more likely than LT surgeons to choose LR compared with LT. Such findings may seem self evident and lead one to assume that

the impact of surgeon specialty is merely an example of Maslow's hammer<sup>37</sup> (ie, that surgical specialists use the techniques available to them). Our analyses revealed, however, that the effect of specialty was entirely explained by the fact that certain clinical variables were weighed differently by non-LT versus LT surgeons. For example, in choosing between LR and LT, non-LT surgeons were more likely than LT surgeons to be influenced by LT waiting time and extent of resection required. As such, variations in choice of therapy for early HCC do not appear to reflect dogmatic, specialty-specific preferences for one therapy versus another. Rather, the effect of surgeon specialty on decision making is mediated by the differential impact of certain clinical factors. A notable exception was found, however, in the choice of bridging therapy, for which specialty-specific differences persisted despite accounting for differential weighting of clinical factors.

These data are important for several reasons. First, they begin to shed light on the underlying root causes for variation in choice of therapy for early HCC. They also highlight a potential mechanism for variation in choice of therapy that may be relevant in other cancers. Second, identification of factors that are weighed differently by non-LT and LT surgeons can focus debate and consensus building regarding the appropriate roles of these factors in driving choice of therapy. Our data should not serve as a guide to choosing surgical therapy for early HCC, but they do highlight areas of agreement and disagreement among experts that should stimulate future work. Finally, because these differences in decision making are likely the result of both surgical training and subsequent surgical experience, these data highlight the potential for surgical specialization to result in the



formation of intellectual silos, even in closely related disciplines, and even in an era when presentation of cases at multidisciplinary tumor boards is routine (93% in this study). Reconciling differences in surgical decision making will require high-quality data, not just multidisciplinary discussion, to demonstrate the optimal treatment strategies for specific subgroups of patients.

A particular strength of our study was the use of conjoint analysis to assess provider decision making. Although it has been used extensively to elicit the preferences of patients and communities for health care,<sup>20,21</sup> conjoint analysis has also been used to study decision making by physicians,<sup>38-41</sup> including surgeons.<sup>22-30</sup> Conjoint analysis is a robust technique with the benefits of a strong theoretical underpinning and rigorous, quantitative methodology.<sup>18,19</sup> Conjoint analysis allows clinical factors of interest to be explored in a clean context that is free from constraints, such as institutional policies or insurance regulations, that would influence interpretation of practice data. The Delphi method, which relies on experts' own introspections regarding their decision-making strategies, has been advocated as a means of forging consensus in areas of surgical oncology with substantial practice variation.<sup>42</sup> However, clinical judgment analysis, such as conjoint analysis, generates assessments of decision-making strategies that are more accurate than decision makers' own perceptions of how they use information,<sup>22,38,40</sup> providing a useful reality check of expert opinions.

Several limitations of our study should also be considered. First, this study assessed stated preferences for surgical therapy as opposed to surgeons' actual practice patterns. This approach has the advantage of standardizing case scenarios and reducing confounding, but it has the disadvantage of assessing decision making in a relatively idealized context. For example, referring physicians, such as hepatologists, who were not included in this study, may also play an important role in choice of surgical therapy. Nevertheless, previous work has demonstrated that this approach results in valid assessments of clinical decision making that are predictive of future decisions.<sup>26,38,43</sup> Second, because we do not know the characteristics of survey nonrespondents, we cannot verify that our respondents are representative of all surgeons who treat HCC. Finally, our survey focused on a subgroup of patients with HCC—those with Milan-criteria tumors and well-compensated cirrhosis—who are the focus of greatest controversy in choice of therapy. Although many patients with HCC will not fall into this group, decision making is often simpler in those cases, because

fewer surgical options are generally available to them. Also, because this study focused on initial surgical therapy, other therapeutic strategies, such as LR followed by salvage LT, were not considered.

In conclusion, choice of surgical therapy for early HCC varies widely as the result of both clinical factors and surgeon specialty. The impact of surgeon specialty on choice of therapy is stronger than that of some clinical factors. However, the influence of surgeon specialty results, for the most part, from the differential impact of certain clinical factors rather than an across-the-board preference for one therapy over another. These data should inform future initiatives to understand choice of therapy for patients with HCC as well as other malignancies. High-quality clinical data must be complemented by a deeper understanding of clinical decision making if we are to ensure delivery of consistent, high-quality care for all patients with cancer.

#### AUTHORS' DISCLOSURES OF POTENTIAL CONFLICTS OF INTEREST

The author(s) indicated no potential conflicts of interest.

#### AUTHOR CONTRIBUTIONS

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