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Risk Factors And Long-term Prognosis Of BCLC Stage 0/A Hepatocellular Carcinoma For Beyond Milan Recurrence After Hepatectomy: A Multicenter Observational Study

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Background: Data on recurrence are important to inform surveillance and improve long—term surgical outcomes for patients with hepatocellular carcinoma (HCC). We sought to identify risk factors and long—term prognosis among patients who experienced beyond—Milan recurrence (BMR) after hepatectomy for early—stage HCC.

Methods: Patients who underwent hepatectomy for BCLC stage 0/A HCC were identified from a multi-institutional database. Predictors of BMR and risk factors associated with post-recurrence survival (PRS) among patients with BMR were assessed using univariate and multivariate Cox-regression analyses.

Results: Among 753 patients, 138 (18.3%) developed BMR at a median follow-up of 51.8 months. Regular surveillance (interval follow-up ≤ 3 months within 1 year and ≤ 6 months in subsequent years after surgery) was not carried out for 53 (38.4%) patients who developed BMR. On multivariate analysis, increased risk of BMR was independently associated with preoperative alpha-fetoprotein level $\rangle 400$ ng/mL, tumor size $\rangle 5.0$ cm, multifocal disease, microvascular invasion, and no/irregular recurrence surveillance. Median PRS among patients with BMR was only 8.4 months (95%CI:7.0–9.8 months). Among patients who developed BMR, Child-Pugh grade B/C, early recurrence within 1 year after surgery, macrovascular invasion/distant metastasis, and non-curative treatment of recurrence were independent risk factors associated with worse PRS.

Conclusions: Nearly 1 in 5 patients developed BMR following hepatectomy for early-stage HCC. Patients with BMR had a median survival of less than 1 year after diagnosis of the recurrence. Regular surveillance is an important and actionable measure to decrease BMR and, in turn, improve long-term survival among patients treated with hepatectomy for HCC.

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