## Meeting abstract

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# Clinical and pathologic factors associated with development of hepatocellular carcinoma in patients with hepatitis virus-related cirrhosis: a long-term follow-up study

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#### Background

Hepatocellular carcinoma (HCC) represents >90% of primary liver neoplasms and develops mainly in patients with liver cirrhosis. Risk factor identification for development of HCC in patients with cirrhosis possesses great clinical relevance due to its high incidence and poor prognosis when detected at advanced stages. The aim of our study was to identify HCC development-associated risk factors in a cohort of patients with hepatitis virus-related chronic liver disease and cirrhosis.

#### Materials and methods

Patients with a diagnosis of hepatitis virus-related cirrhosis from January 1980 to January 2000 were included. Patients were followed with abdominal ultrasound and determination of alpha-fetoprotein levels, physical examination, and routine biochemical tests every 3–6 months. The endpoint in this study was defined as development of HCC. Liver histology was evaluated according to the French METAVIR Cooperative Study Group (METAVIR) score.

#### Results

Two hundred and eighty two patients met the inclusion criteria; the majority of these (86%) had a serologic diagnosis of hepatitis C virus, and only 14% had hepatitis B

virus at the time of diagnosis of cirrhosis, while 56 and 37% were classified as Child A and B, respectively, and only 7% as Child C. Histological activity was mild in 59% of patients, and moderate and severe in 41%. Mean annual incidence was 1.87%, and 22 and 35% of patients developed HCC at 10 and 15 years of follow-up, respectively. Diagnosis of HCC was made by histopathology in 37% and by tumoral lesion-associated alpha-fetoprotein elevation confirmed by imaging studies in 63%. In multivariate analysis, we found three variables associated with HCC: moderate to severe histological activity; platelet count <105 × 10<sup>3</sup>/mm<sup>3</sup>, and alpha-fetoprotein >5 ng/mL (see Table 1). We divided patients into two groups according to regression coefficient: low and high-risk; patients assigned to the low-risk group showed 5-, 10-, and 15-year HCC incidences of 3.4, 6.4, and 6.4%, respectively, in contrast to patients from the high-risk group, who showed incidences of 17.8, 33.5, and 56.8%, respectively.

#### Conclusion

We found three HCC-associated variables: histological activity; platelet count and alpha-fetoprotein levels. Patients with high risk for developing hepatocellular carcinoma must be considered candidates for closer follow-up.

Variables	Hazard ratio (95% confidence interval)	Coefficients	P value (log rank)
Platelet count	3.7 (1.2–11)	1.3	0.025
$\alpha$ -fetoprotein	5.41 (1.86–15.7)	1.23	0.02
Histological activity	7.6 (1.7–33)	2.03	0.007

### Table I: Variables with independent predictive value for HCC in the multivariate analysis