

Albumin-bilirubin score: an additional tool for the management of patients with hepatocellular carcinoma

Matteo Serenari, Giorgio Ercolani

Department of Medical and Surgical Sciences, University of Bologna, Bologna, Italy

Correspondence to: Giorgio Ercolani. Department of Medical and Surgical Science, University of Bologna, Bologna, Italy; Director of Department of Surgery, City Hospital in Forlì, Forlì, Italy. Email: giorgio.ercolani2@unibo.it.

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We read with great interest the article written by Feng and colleagues published in the issue of April in *Annals of Translational Medicine* (1). The authors reviewed all the studies that examined the prognostic value of albumin-bilirubin (ALBI) score in patients mostly affected by hepatocellular carcinoma (HCC).

ALBI grade was first proposed by Johnson et al. in 2014 (2) to offer a more objective method to assess liver function in HCC patients, thus eliminating subjective variables such as ascites and encephalopathy which are required in traditional Child-Pugh (CP) classification. Both European Association for the Study of the Liver (EASL) and American Association for the Study of Liver Diseases (AASLD) guidelines for the treatment of HCC endorse the Barcelona Clinic Liver Cancer (BCLC) classification. BCLC staging system consists of five stages and allocates patients according to pre-established prognostic variables. Among them, a preserved liver function defined as CP class A without any ascites is a prerequisite which applies to all treatment options except for liver transplantation. However, ALBI grade seemed to improve the accuracy of prognosis, and ALBI grade 1 patients, as showed in Table 1 of the manuscript, had a significantly better survival compared to those classified as ALBI grade 2/3, even within the same CP class (3). This would mean that such a variation in the prognosis, related to a more precise assessment of the underlying liver function, may potentially change the type of intervention to which a patient could be candidate.

Therefore, ALBI grade represents an additional tool to help clinicians in determining the best treatment option for HCC patients (4). First, among optimal surgical candidates (BCLC 0-A) ALBI score has been demonstrated to better predict the risk of decompensation after hepatectomy (ALBI grade 2/3), including also minimally-invasive liver resections (5), compared to traditional indices (6). Hence, a high-risk subgroup of patients may be identified and benefit more from less invasive treatments such as radiofrequency ablation (RFA) or even to liver transplantation, when survival advantage obtained with liver resection is limited (7). Despite clear advantages and better prognostic value of ALBI grade compared to CP score, some authors considered such a difference not clinically relevant. On the other hand, it should be emphasized that other tests and scores have been validated in the evaluation of patients with HCC waiting for liver resection. MELD score better stratifies cirrhotic patients with HCC for partial hepatectomy than CP classification (8,9). Many Eastern and Western surgeons commonly apply indocyanine green test to select patients for liver resection (10).

Besides BCLC 0-A, patients classified as BCLC B represent an even more heterogeneous population. Although transarterial chemoembolization (TACE) is recommended for these patients, other treatments such as radioembolization, RFA or even surgical resection, have been explored in the past and might be suited to BCLC B patients based on ALBI classification (11,12). On the other side, ALBI-based BCLC may upstage a significant number of patients from CP class B to ALBI grade 3 (or ALBI-based BCLC stage D) (12) (i.e. with an observed survival <3 months), thus optimizing also administration of

experimental therapies in advanced HCC (13).

Though Feng and co-workers report the usefulness and superiority of ALBI grade even in other liver disease, we should remember that at the moment incorporation of hyponatremia into MELD score (MELD-NA) showed the best prognostic factor in patients waiting for liver transplantation (14). Similarly incorporating ICG into MELD (MELD-ICG) score seems to better discriminate intermediate to advanced cirrhotic patients (15).

Anyway, we would like also to highlight the potential role of ALBI score in predicting HCC recurrence. Although the Authors in their review article did not cover this topic, ALBI grade together with other risk factors, has been demonstrated to predict early and *de novo* recurrence after hepatectomy, thus contributing in guiding surveillance follow-up and in the design of clinical trials (16,17).

In conclusion, ALBI grade seems to be a reliable marker of hepatic function, both in HCC and other liver disease. It is much more effective than CP in stratifying patients waiting for liver resection or other treatment and it seems to work as a prognostic factor for recurrence. However, prospective studies comparing ALBI grade to MELD-NA or MELD-ICG in HCC and cirrhotic patients are required to define which might perform better.

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