



# Prediction of the Risk of Lymph Node Metastases in Early Gastric Cancer: Contrast-Enhanced Harmonic Endoscopic Ultrasonography May Help

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## To the Editor:

We have read with interest the article by Shin *et al.*<sup>1</sup> reporting their experience in the evaluation of clinicopathologic features associated with lymph node metastases (LNM) in submucosal papillary gastric cancer.

The authors found that the LNM rate tended to be higher in papillary early gastric cancer (P-EGC) than in other differentiated types of EGC and that the rate increased to 25.6% when there was invasion of the submucosal layer.

Lymphovascular invasion was the only factor significantly associated with LNM in submucosal P-EGC. Moreover, location in the lower third of the stomach and elevated gross appearance were independent factors associated with lymphovascular invasion in submucosal P-EGC. Conversely, the depth of submucosal invasion, which is a known predictive factor against LNM in classic EGC, was not significantly associated with LNM.

These findings raise concern about the appropriateness of the current European guidelines that recommend the use of the same criteria for endoscopic submucosal dissection (ESD) as a treatment or both P-EGC and other differentiated EGCs.<sup>2</sup> The feasibility of ESD for P-EGC is still debatable, in particular when it meets the expanded criteria.

Accurate assessment of the depth of invasion of EGC is critical for selecting the appropriate treatment option. Although endoscopic ultrasonography (EUS) has become the preferred tool for the locoregional staging of gastric cancer,<sup>3</sup> there is no consensus on the accuracy of EUS for the evaluation of the invasion depth of EGC.<sup>4-6</sup>

However, several studies have shown the optimal accuracy of contrast-enhanced harmonic EUS (CH-EUS) for the differential diagnosis of benign and malignant lesions.<sup>7-10</sup> In particular, the diagnostic accuracy of CH-EUS in the differential diagnosis of lymph nodes was comparable to those of EUS elastography and EUS-guided fine needle aspiration.<sup>11</sup> Therefore, the characterization of lymph nodes could represent the main target of EUS during EGC staging rather than the T parameter, especially for P-EGC.

Several studies performed on surgical specimens reported a worse prognosis associated with P-EGC than with other differentiated types.<sup>2,12</sup> However, the treatment outcomes of ESD for P-EGC have not been precisely documented.<sup>12</sup>

A recent Korean study<sup>13</sup> evaluated the short- and long-term outcomes after ESD in P-EGC; the curative resection rate of P-EGC was significantly lower than those of well differentiated and moderately differentiated EGC (49.4% vs 72.5% and 93.7%, respectively), although it increased to 72.5% for mucosal (T1a) cancer. Despite the poor short-term outcomes, the long-term outcomes of ESD for P-EGC were favorable once



curative resection was achieved (no LNM, no extragastric recurrences and a low metachronous recurrence).

As discussed by the authors, qualitative endoscopic criteria evaluation and accurate staging should be adopted for the assessment of submucosal invasive P-EGC, although lymphovascular invasion is difficult to predict. Indeed, accurate staging, together with a radical endoscopic or surgical resection of gastric neoplasia can significantly improve patients' clinical outcomes.<sup>14</sup>

In P-EGC accurate N staging is crucial due to the high rate of LNM even in the early stages. We think that EUS evaluation, together with CH-EUS, should therefore be included in the staging process for P-EGC. Further large-scale studies are needed to demonstrate whether the correct staging yield and prediction of lymphovascular invasion could be further improved.

## CONFLICTS OF INTEREST

No potential conflict of interest relevant to this article was reported.

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