A 60-year-old man presented with worsening right upper quadrant pain, diarrhea and weight loss. Computerized tomography revealed multiple lesions in the liver that were suspicious for metastatic disease (Figure 1). In search of a primary lesion, colonoscopy was performed. Two rectal polyps were found and removed with snare cautery. One of the polyps (0.9 cm) showed a poorly differentiated neuroendocrine carcinoma (0.3 cm), arising in a tubulovillous adenoma, invading the muscularis mucosae but not the submucosa (Figure 2). Multiple step sections, exhausting almost all the tissue, were examined, looking for deeper invasion, but none was identified. Although cautery artifact was present at the deep edge of the specimen, it was not severe enough to warrant concern about any destruction of evidence of submucosal tumor. Immunohistochemically, the tumor was positive for neuron-specific enolase and synaptophysin but negative for chromogranin. Positivity for two out of three neuroendocrine markers supported the probability of the neuroendocrine nature of the tumor. Fine-needle aspiration of the liver lesions showed cells similar to the rectal primary. The liver tumor showed an immunohistochemical staining pattern identical to that of the rectal tumor, being positive for neuron-specific enolase and synaptophysin but negative for chromogranin. An extensive radiological work-up, including positron emission tomography, was negative for any other alternative primary lesions. The patient declined therapy and was discharged to hospice care.

Tumor was found at only two sites in this patient (rectum and liver). Since these two lesions were morphologically and immunohistochemically identical, we concluded that the liver lesions represented metastases from the rectum. The absence of other lesions radiologically and endoscopically makes it unlikely that both the rectal and hepatic lesions represent metastasis from a third, occult site.

Although the patient’s presenting complaints were most likely related to metastatic tumor, the cause for the mild diarrhea was not readily apparent. In accordance with the patient’s wishes, further work-up was not carried out, and serum gastrin or vasoactive intestinal peptide levels were not determined. In the absence of this information, the possibility that the diarrhea may have been related to hormonal products of the neuroendocrine tumor must remain speculative.

Poorly differentiated neuroendocrine carcinomas of the gastrointestinal tract carry a worse prognosis than adenocarcinomas, typically metastasizing despite only superficial invasion [2]. They often arise in the setting of an overlying adenoma, as seen in the present case [2, 3]. However, our case is unique because it is an exception to the rule that colorectal tumors that do not infiltrate the submucosa are without metastatic potential [3]. Hepatic metastases in the absence of submucosal invasion can be explained by invasion of capillaries rather than lymphatics [3]. Small advanced colorectal carcinomas are rare. The smallest tumor in a series published in an earlier issue of Endoscopy was 0.8 cm in size [4]. A 0.3-cm nonmetastatic neuroendocrine carcinoma arising in a tu-
bulovillous adenoma has also been described [2]. Minamoto et al. described a 0.5-cm nonpolypoid adenocarcinoma metastatic to lymph node [5]. To the best of our knowledge, the present case represents the smallest and most superficial metastatic colorectal carcinoma described in the literature to date. Histological examination of endoscopically “benign” polyps may reveal an unsuspected primary source of a metastatic carcinoma of unknown origin.

S. Mukhopadhyay¹, A. Marhaba², J. S. Sidhu¹

¹ Department of Pathology, State University of New York Upstate Medical University, Syracuse, New York, USA
² Department of Gastroenterology, United Health Services Hospitals, Johnson City, New York, USA
³ Department of Pathology and Laboratory Medicine, United Health Services Hospitals, Johnson City, New York, USA

References


Corresponding Author

S. Mukhopadhyay, M.D.
Department of Pathology
State University of New York Upstate Medical University
750 East Adams Street
Syracuse, NY-13210
Fax: +1-315-464-7130
E-mail: mukhopas@upstate.edu