Curable Resection in Gastric and Lymph Node Metastases From Melanoma

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ABSTRACT

We herein report a rare case of gastric and regional lymph node metastasis of cutaneous malignant melanoma that underwent curative resection. The patient, a 68-year-old man, was first diagnosed as having cutaneous malignant melanoma of the right forearm in 2005. He had extensive skin excision and axillary lymph node dissection and had undergone adjuvant chemotherapy. Six years after the primary surgery, gastrointestinal endoscopy revealed gastric metastasis of a malignant melanoma. As there was no other metastasis found, laparoscopic-assisted distal gastrectomy with lymph node dissection was performed. Microscopic findings showed diffuse melanin granule growth invading the muscularis propria of the stomach. Micrometastases of the lymph nodes were observed that were not detected by preoperative examination. Seventeen months have passed without recurrence. We conclude that regional lymph node dissection should be performed with gastrectomy whenever distant metastases are not observed, because there is a possibility of micrometastases, which cannot be detected preoperatively.

Key Words: Gastrectomy, Gastric metastasis, Laparoscopic surgery, Lymph node metastasis, Melanoma.

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INTRODUCTION

Metastatic melanoma is a poor prognosis disease for which multiple metastases are often observed. Metastatic malignant melanoma of the gastrointestinal tract is said to be seen in 4% of patients.¹ Autopsy data indicate that the small intestine was the most commonly involved metastatic site (58%), followed by the stomach (26%).² However, most gastrointestinal metastasis are not discovered due to lack of digestive symptoms and mostly are accompanied with other multiple distant metastases.

We report a case with gastric and regional lymph node metastasis of malignant melanoma, which underwent curative resection by laparoscopic surgery. To our knowledge, this is the first case to report a curative resection of gastric metastasis and lymph node metastases of malignant melanoma.

CASE REPORT

A 68-year-old man was first diagnosed as having cutaneous malignant melanoma of the right forearm in 2005. He had extensive skin excision and axillary lymph node dissection. According to American Joint Committee on Cancer,³ the histology was T4b, N0, M0, Stage IIc, and the patient underwent 4 courses of dacarbazine, nimustine, vincristine, and interferon-beta therapy as adjuvant chemotherapy. He had periodic survey for 5 years without recurrence.

He had gastrointestinal endoscopy for medical checkup in 2011, and an elevated tumor with ulcer and brownish punctate distribution was observed in the posterior wall of the lower gastric body (**Figure 1**). From the upper gastrointestinal examination, a "bull's eye" configuration was observed and gastric metastasis was suspected. The bi-



Figure 1. Gastrointestinal endoscopy findings showed elevated tumor with ulcer and brownish punctate distribution in the posterior wall of the lower gastric body.

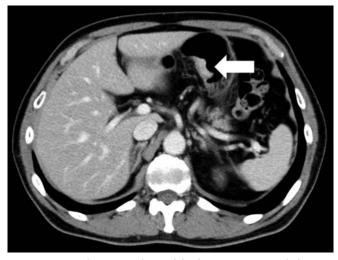


Figure 2. PET/CT scan showed high-intensity spot (white arrow) in the lesser curvature of the stomach. No other high-intensity spot was observed. CT, computed tomography; PET, positron-emission tomography.

opsy confirmed the diagnosis as gastric metastasis of the malignant melanoma.

The laboratory test results were within normal limits except for slight elevation of the serum lactic dehydrogenase level 227 IU/L and the elevation of 5-S-cysteinyldopa 28.5 nmol/L. A contrast computed tomography (CT) scan showed tumor stain in the lesser curvature of

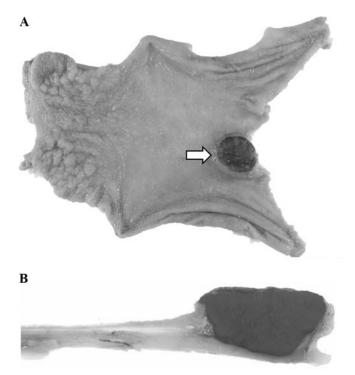


Figure 3. A. The tumor (white arrow) was 2 cm in diameter with ulceration on the surface. **B.** The division surface showed sharp marginated tumor.

the stomach. Lymph nodes' swelling and ascites were clinically negative on imaging. Positron-emission tomography (PET)/CT scans were also performed and a high-intensity spot was only seen in the lesser curvature of the stomach **(Figure 2)**.

As there were no other metastases, laparoscopic-assisted distal gastrectomy with lymph node dissection was performed. The black tumor was 2 cm in diameter and sharply marginated with ulceration on the surface in the resected specimen (**Figure 3**). Pathological findings of the tumor showed abundance of melanin granule with expansive growth and invasion to the muscularis propria of the stomach (**Figures 4 and 5**). The junctional change in the overlying epithelium was not observed. Elastica van Gieson stain and D2–40 stain were performed; however, vascular invasion was beyond determination. Two perigastric lymph nodes metastases, stations no. 3 and no. 7 according to the Japanese Classification of Gastric Carcinoma, 4.5 were also observed in the subcapsular sinus of lymph nodes (**Figure 6**).

Postoperative course was satisfactory, and he started dacarbazine, nimustine, cisplatin, and tamoxifen therapy. Chest, abdominal, and brain contrast CT and gastrointes-

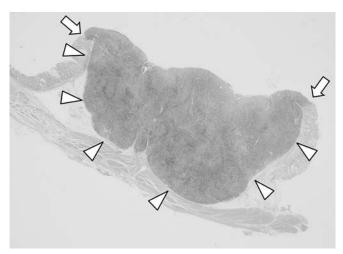


Figure 4. The tumor was microscopically sharply marginated and localized (white arrowheads). No junctional change in the overlying epithelium was observed (white arrow). (Hematoxylin and eosin stain.)

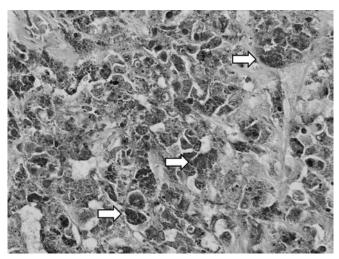


Figure 5. Pathological findings of the tumor showed abundance of melanin granule with expansive growth (white arrows). (Hematoxylin and eosin stain.)

tinal endoscopy were performed every 6 months and PET was performed 1 year after operation. Seventeen months have passed without recurrence.

DISCUSSION

There have been only few reports of patients with metastatic melanoma in the gastrointestinal tract who underwent surgery. **Table 1**^{6–13} summarizes the cases with gastrointestinal tract metastases from cutaneous malignant melanoma reported from 2000 to 2012. As reported, the small intestine was the most commonly involved metastatic site. In most

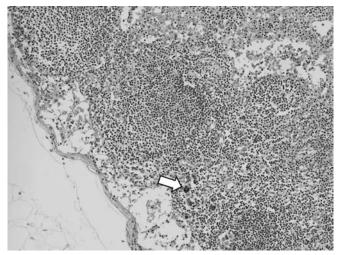


Figure 6. Micrometastases (white arrow) were observed in the subcapsular sinus of lymph nodes. (Hematoxylin and eosin stain.)

cases, detailed examination was performed due to the digestive symptoms, and distant metastases were also detected. Because most cases had multiple distant metastases, reduction therapy were performed to control the digestive symptoms. The cases in which the metastatic site was localized and curative surgical resection was performed had better prognosis than the noncurative surgical resection cases did.

Three types of metastatic malignant melanoma in the gastro-intestinal tract are observed by gastrointestinal endoscopy: (1) nodules varying in size, usually appearing to arise on the crest of normal range, often ulcerated at tip, and invariably melanotic; (2) raised submucosal tumor masses with ulcerated centers; (3) mass lesions with varying degrees of necrosis and melanosis. ¹⁴ Types 1 and 2 were also recognized in our case. The difference between these 3 types of the appearance might be the size of the tumor. Enlargement of the tumor might cause the ulceration and necrosis of the center. Histological changes between primary and metastatic malignant melanoma is the absence of junctional change in the overlying epithelium.¹⁵

The PET scanning for detecting melanoma metastasis had high sensitivity of 84% to 94% and specificity of 83% to 97%, whereas CT showed 55% to 58% sensitivity and 70% to 84% specificity. 16,17 PET is the most sensitive method compared with conventional imaging, particularly in soft tissues, lymph nodes, and liver. It also helps to detect sites that are not routinely evaluated by CT. 17,18 False-negative scans are observed in 4% of all scans, due to the small size, <0.3 to 0.5 cm, or the tumor burden, consisting of only a few cells, that were unable to be resolved by PET scanning. 18,19

			The Cases W	ith Gastrointestinal	Table 1. Tract Metastases	Table 1. The Cases With Gastrointestinal Tract Metastases From Cutaneous Malignant Melanoma	alignant Me	lanoma	
					Metastatic Sites				
Author (yr)	Age, yrs	Sex	Primary Lesion	Digestive Symptom	GI Tract	Others	Stage, AJCC	Treatment	Survival From the Diagnosis of GI Metastasis
Târcoveanu et al ⁶ (2009)	53	M	Chest	Abdominal pain	Small intestine	Brain	NS	Laparotomy	Dead, 4 mo
Târcoveanu et al ⁶ (2009)	63	щ	NS	Abdominal pain	Small intestine	1	NS	Laparotomy	Alive, 5 mo
Tsilimparis et al 7 (2009)	43	M	Upper back	Abdominal pain (due to perforation)	Small intestine	Thalamus, lung, liver, gall bladder	IIIA or more	Laparotomy, DTIC	Alive, 8 mo
Park et al ⁸ (2009)	74	M	Upper back	GI bleeding	Small intestine	1	IIIA or more	Laparotomy	Alive, NS
Albert et al 9 (2007)	69	П	Left lower leg	GI bleeding	Duodenum, small intestine	Skin, LNs	IIB or more	Laparotomy, IFN	Alive, 14 mo
Liang et al ¹⁰ (2006)	72	M	Lower back	Melena	Small intestine	Skin, lung, brain, stomach, abdominal LNs	IA or more	Laparotomy, radiation, vaccine, molecular target therapy	Dead, <12 mo
Belagyi et al ¹¹ (2006)	22	ш	Back	None	Small intestine, stomach	Bilateral ovary, left sternocleidomastoid muscle, pancreas, subcutaneous tissue	NS	Laparotomy, radiation, CCNU, DTIC, VCR, BLM, IFN	Dead, 4 mo
Loualidi et al 12 (2004)	9	M	Back	GI bleeding	Duodenum	Skin, LNs	IA or IB	Palliative care	Dead, 13 mo
Tatlidil and Mandelkem ¹³ (2001)	99	ш	Left shoulder	Abdominal pain, fecal occult blood	Small intestine	Subcutaneous tissue, chest wall, mediastinum, abdominal LNs	IIIA or more	Exploratory laparotomy, DTIC	Dead, 12 mo
Tatlidil and Mandelkern ¹³ (2001)	53	П	Left scalp	Abdominal pain, cramping	Small intestine, colon	Skin, gall bladder liver, femur, LNs	IA or more	Laparotomy, chemotherapy	Dead, 11 mo
Tatlidil and Mandelkern ¹³ (2001)	36	M	Chest	NS	Small intestine	Mesenteric, LNs	IA or more	Laparotomy, DTIC, CDDP, VCR, IL-2, IFN, tamoxifen, vaccine	Alive, 48 mo
Tatlidil and Mandelkern ¹³ (2001)	55	M	Left ear lobe	GI bleeding	Small intestine	Liver, lung, pelvis, iliac crest, LNs	IA or more	Laparotomy, DTIC, CDDP, BCNU, tamoxifen	Dead, <12 mo
Otowa et al (2014)	89	M	Right forearm	None	Stomach	INS	IIC	Laparotomy, DTIC, CDDP, ACNU, tamoxifen	Alive, 17 mo

AJCC, American Joint Committee on Cancer; BCNU, carmustine; BLM, bleomycin; CCNU, lomustine; CDDP, cisplatin; DTIC, dacarbazine; F, female; GI, gastrointestinal bleeding; IFN, interferon; IL-2, interleukin-2; INs, lymph nodes; M, male; mo, months; NS, not specified; VCR, vincristine.

Most patients have distant metastases; however, if the metastasis is localized and resectable, surgical resection should be tried. Some previous studies show longer median survival when curative surgery was performed than for those with nonsurgical treatment.^{20,21}

Though chemotherapy is widely used as an alternative treatment, it has been difficult to improve the prognosis. Recently, immunotherapy with ipilimumab has been shown to improve overall survival²² and molecular target therapy with vemurafenib has been shown to improve progression free and overall survival in patients with advanced malignant melanoma.²³ The combined modality using the new treatments might improve the prognosis.

CONCLUSIONS

In our case, lymph node metastasis was observed in the subcapsular sinus. This might suggest that resection of the metastasis lesion with lymph nodes dissection is recommended for curative surgery even if the PET scan is negative and might lead to a long disease-free interval.

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