# World Journal of Clinical Cases

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CASE REPORT

## Long-term survival of the Sister Mary Joseph nodule originating from breast cancer: A case report

Koji Kanayama, Maki Tanioka, Yoshitsugu Hattori, Takuya Iida, Mutsumi Okazaki

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#### **Abstract**

#### **BACKGROUND**

A Sister Mary Joseph nodule (SMJN) is an uncommon cutaneous metastasis found in the umbilicus, indicating an advanced malignancy. SMJNs typically originate from intra-abdominal sources, rarely from breast cancer. Diagnosis suggests a poor prognosis with a median survival of approximately 8 mo after detection. Managing patients with SMJNs is challenging, as most receive limited palliative care only. The optimal strategy for long-term survival of these patients remains unclear.

#### CASE SUMMARY

A 58-year-old female, previously diagnosed with right breast cancer 17 years ago and underwent breast-conserving surgery, adjuvant radiotherapy, and endocrine therapy, presented with a 2-cm umbilical nodule. Thirteen years previously, metastases were detected in the right supraclavicular, infraclavicular, hilar, and mediastinal lymph nodes. An umbilical nodule emerged four years before the date of presentation, confirmed as a skin metastasis of primary breast cancer upon excisional biopsy. Despite initial removal, the nodule recurred and grew, leading to her referral to our hospital. The patient underwent extensive excision of the umbilical tumor and immediate abdominal wall reconstruction. Endocrine therapy was continued postoperatively. Five years later, no local recurrence was observed, and the patient continued to work full-time, achieving over 9 years of survival following SMJN diagnosis.

#### CONCLUSION

This case study aimed to identify the optimal strategy for achieving extended survival outcomes in patients with SMJN through comprehensive treatment. We presented a case of the longest survival in a patient after undergoing a multidisciplinary treatment regimen. Our findings underscore the significance of adopting

a multimodal treatment approach comprising timely and wide excision along with adjunctive therapy. This approach can control the disease, prolong survival, and improve the quality of life in patients with SMJN.

Key Words: Sister Mary Joseph nodule; Breast cancer; Cutaneous metastasis; Long-term survival; Abdominal wall reconstruction; Case report

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Core Tip: Sister Mary Joseph nodules (SMJN) typically present as firm, irregular periumbilical nodules, which are uncommon yet significant symptoms that indicate an advanced stage of malignancy with a poor prognosis. Although most patients receive limited treatment, the optimal systemic treatment strategy for long-term survival remains controversial. While treatment decisions should be based on the patient's overall health and condition, a multimodal treatment approach comprising timely and wide excision along with adjunctive therapy has the potential to control the disease, prolong survival, and improve the quality of life of patients with SMJNs without peritoneal dissemination.

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#### INTRODUCTION

The Sister Mary Joseph nodule (SMJN) is an unusual cutaneous metastasis in the umbilicus. It originates from intraabdominal or pelvic sources and is rarely associated with breast cancer[1]. On physical examination, SMJN is an uncommon yet significant sign indicating an advanced stage of malignancy and a poor prognosis. The median survival duration after SMJN diagnosis was 7.9 mo, with a 95% confidence interval of 6.7 to 9.1 mo[2]. Because managing patients with SMJN is challenging, they generally receive limited treatments, such as palliative surgery or best supportive care. However, the optimal systemic treatment strategy for long-term survival remains controversial. This report presents a rare case of a patient diagnosed with SMJN originating from breast cancer who survived for over 9 years after receiving treatment with a multidisciplinary approach. The patient remained active in society, working full-time.

#### **CASE PRESENTATION**

#### Chief complaints

A painless 2-cm hard nodule with erosions in the umbilical region, which was first detected four years ago.

#### History of present illness

At age 41 (17 years ago), she underwent a right breast-conserving surgery (invasive ductal carcinoma, ER+, PR+, HER-, T2N2M0). The patient was treated with adjuvant chemotherapy (four cycles of Epirubicin and Cyclophosphamide), followed by adjuvant radiotherapy (50Gy/25Fr) and endocrine therapy (Tamoxifen). At age 45 (13 years ago), metastases were detected in her right supraclavicular to infraclavicular lymph nodes, right hilar lymph nodes, and mediastinal lymph nodes on positron emission tomography-computed tomography (PET-CT). Consequently, she was prescribed an LH-RH agonist (Leuprorelin) with Tamoxifen as a second-line treatment. Over time, the tumor showed a trend of reduction in both lymph node and lung metastases compared to their state before treatment, leading to the discontinuation of Leuprorelin. At age 53 (5 years ago), Tamoxifen was switched to Anastrozole as progressive enlargement of the right supraclavicular lymph node was found. One year later, Leuprorelin was resumed because the CT scan showed an increasing number of nodules in both the upper and lower lobes of the right lung. At age 54 (4 years ago), an umbilical nodule emerged, for which an excisional biopsy was performed, confirming skin metastasis originating from the primary breast cancer (adenocarcinoma, metastatic, ER+, PR+, HER-). Although endocrine therapy with Leuprorelin and Exemestane was initiated, the nodule recurred and gradually increased in size, leading to her referral to our hospital.

#### History of past illness

The patient has no significant medical history.

#### Personal and family history

No notable abnormalities: Never smoked, no alcohol consumption, and no known allergies to drugs or food.



#### Physical examination

A hard, localized, and poorly mobile nodule accompanied by erosion was detected in the umbilical region. The nodule measured approximately 2 cm in diameter, with no redness, heat, pain, or tenderness (Figure 1A and B). A lymph node approximately 2 cm in diameter was palpable subcutaneously above the right clavicle, with no other palpable lymph nodes including the axillary or inguinal regions.

#### Laboratory examinations

Her laboratory results were normal, with the carcinoembryonic antigen level at 3.1 ng/mL (normal reference value: 5 ng/ mL), except for a slight elevation in the carbohydrate antigen (CA) 15-3 Level to 59 U/mL (normal reference value: 21 U/ mL).

#### Imaging examinations

PET scans revealed increased 18F-fluorodeoxyglucose uptake in the umbilicus (SUVmax, 4.0), right supraclavicular fossa (SUVmax, 7.2), and bilateral lung fields (SUVmax, 1.2) (Figure 1C). Additionally, the PET-CT scan indicated no evident nodules suggesting tumor metastasis to the peritoneum in the abdominal cavity, nor was there any evidence of abdominal or pelvic fluid (Figure 1D).

#### FINAL DIAGNOSIS

A puncture biopsy was performed, resulting in a diagnosis of recurrent metastatic adenocarcinoma originating from breast cancer. Histological examination confirmed adenocarcinoma with a foamy tubular arrangement (Figure 2). Immunohistochemistry was positive for ER and GCDFP-15, negative for PR and HER, and showed a low proliferative index of 7% for neoplastic cells on Ki67.

#### TREATMENT

The tumor was excised with a 15 mm horizontal margin, involving combined resection of the peritoneum and falciform ligament (Figure 3A). No tumor was exposed to the abdominal cavity (Figure 3B and C), and no adhesions were observed in the intra-abdominal organs. There was no evidence of peritoneal dissemination. Given the large size of the defect, simple closure was challenging; therefore, we reconstructed the abdominal wall using a component separation technique (Figure 3D-F). Postoperative endocrine therapy with Toremifene citrate was initiated followed by adjuvant radiation therapy (45Gy/15Fr) for the enlarged right subclavian lymph nodes.

#### OUTCOME AND FOLLOW-UP

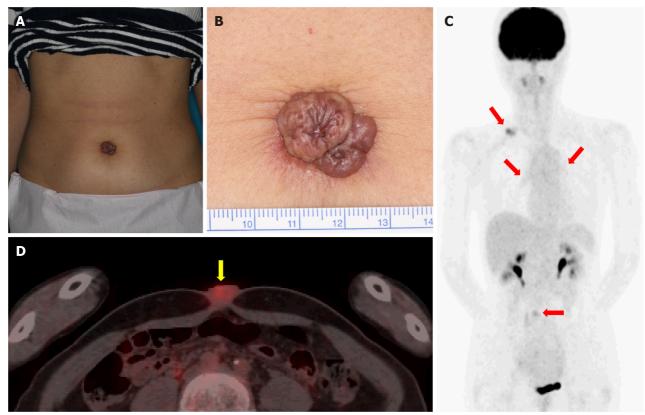
The wound healed completely, enabling the patient to return to work. Five years postoperatively, there was no evident local recurrence in the umbilical region, and the patient continued to work full-time, five days a week (Figure 4).

#### DISCUSSION

This report presents a case of long-term survival in a patient with an SMJN originating from breast cancer. SMJNs typically present as firm, irregular periumbilical nodules, usually measuring 0.5-2 cm. They may be painful, ulcerated, bloody, or suppurative[3]. For patients presenting with an umbilical nodule, the differential diagnosis should include primary umbilical neoplasms, metastatic umbilical nodules, umbilical hernias, umbilical endometriosis, keloids, pyoderma gangrenosum, and foreign bodies. Notably, SMJN may be the sole presenting complaint in some otherwise healthy patients. However, other patients may present with a poor clinical state and additional physical signs, such as ascites and pleural effusion[4]. A focused history and examination of the chest, abdomen, and regional lymph nodes can help identify the source of the primary neoplasm. Patients presenting with SMJN after treatment for a known neoplasm should be evaluated for recurrence. Biopsy of the umbilical nodule offers a convenient method for obtaining tissue samples for histological diagnosis. CT scan and magnetic resonance imaging can determine the extent of the malignancy.

Cutaneous metastases occur in up to 9% of malignancies, with only 10% localized to the umbilicus [5,6]. SMJN is typically associated with primary neoplasms of the gastrointestinal and genitourinary tracts. Other reported primary sites include the lungs, pancreas, liver, gallbladder, breast, kidney, prostate, and testicle [7,8]. However, the source of the primary neoplasm may remain unidentified in up to 30% of patients[5]. Only 1.7% of umbilical metastases originate from breast tissues[9].

The etiology of SMJN remains unclear. Proposed hypotheses include direct extension of the tumor to the umbilicus as well as lymphatic or hematogenous spread[10]. A literature review showed that 21 out of 23 (91%) patients with SMJN at presentation experienced peritoneal dissemination[11]. Although the mechanism of tumor spread from the breast to the umbilicus remains unclear, extensive vascular and lymphatic connections between the umbilicus and axillary and



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Figure 1 A 58-year-old woman presented with Sister Mary Joseph's nodule. A and B: A 2-cm hard, localized, and painless nodule with erosions observed in the umbilical region; C: Positron emission tomography (PET) demonstrates increased 18F-fluorodeoxyglucose uptake in the umbilicus, right supraclavicular fossa, and bilateral lung fields (red arrow); D: PET-computed tomography scan revealed increased 18F-fluorodeoxyglucose uptake in the umbilicus (yellow arrow), but there were no nodules or abdominal/pelvic fluid suggesting tumor metastasis to the peritoneum.

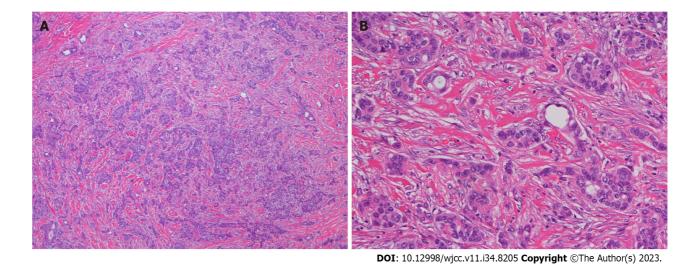
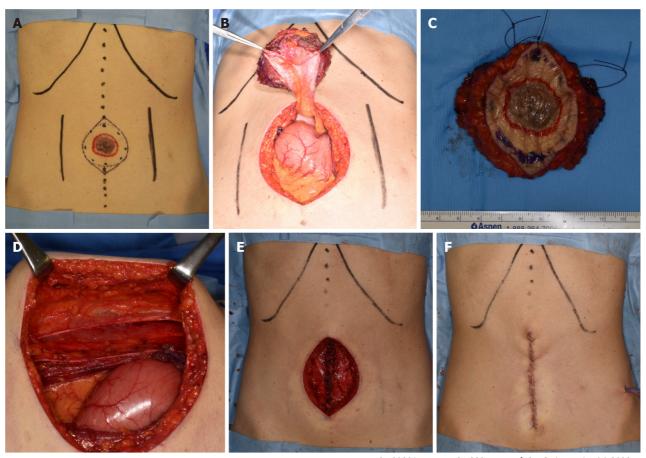


Figure 2 Pathology of the metastatic specimens. A: Morphology (Hematoxylin and eosin, × 40); B: Morphology (Hematoxylin and eosin, × 200) confirming an adenocarcinoma in a foamy, tubular arrangement in metastatic lesions.

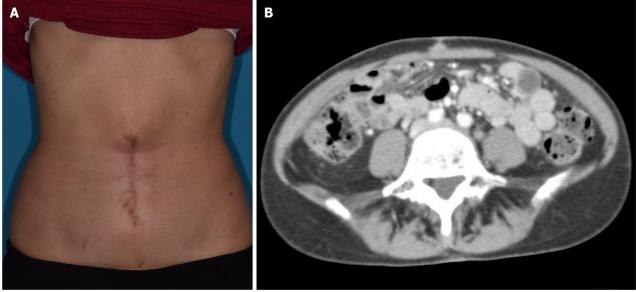
thoracic regions may facilitate the migration of breast tumor cells[5,6,12,13].

Patients diagnosed with SMJN typically have a poor prognosis: fewer than 30% of treated patients survive beyond 1 year, and fewer than 15% survive beyond 2 years[12]. To our knowledge, no report has described a patient with SMJN who survived for over 9 years. The recommended treatment approaches for SMJN remain controversial. Some physicians advocate an aggressive regimen combining surgery with adjunctive therapy [12,14], while others recommend avoiding surgery due to the poor prognosis [15]. Majmudar et al [12] demonstrated that an aggressive multimodality treatment approach, including both surgery and chemotherapy, improved survival (average survival 17.6 mo) compared to those



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Figure 3 Perioperative images of tumor resection and immediate abdominal wall reconstruction. A: Tumor excision with a 15 mm horizontal margin, encompassing a combined resection of the peritoneum and the falciform ligament; B: Image showing no exposure of the tumor to the abdominal cavity; C: Macroscopic image of the resected tissue; D-F: Abdominal wall reconstruction using a component-separation technique.



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Figure 4 Five-year postoperative images. A: No evident local recurrence in the umbilical region; B: Computed tomography scan shows no nodules, suggesting tumor recurrence in the umbilical region or metastasis to the peritoneum in the abdominal cavity.

treated with either surgery alone (average survival 7.4 mo) or adjunctive therapy alone (average survival 10.3 mo); patients who received no therapy had the lowest average survival of 2.3 mo. The prognosis of patients with SMJN is variable and may be most directly affected by the primary tumor type and location. Patients with a primary tumor originating from the pancreas had the poorest prognosis, with a median overall survival rate of approximately 3 mo from the date of diagnosis of umbilical metastasis, while those with primary tumors originating from the ovaries or endometrium had a median survival rate of approximately 10 mo<sup>[2]</sup>.

When determining a management strategy, it is crucial to consider the patient's preferences, clinical condition, and the etiology of the primary malignancy. Although palliative care may be the only viable option for certain patients, others, when carefully selected, may benefit from more aggressive interventions such as surgery, chemotherapy, and radiotherapy. In the present case, the initial approach was local resection followed by endocrine therapy. However, due to disease progression, more extensive surgery was required. Timely and wide excision can prevent peritoneal dissemination and treat patients with SMJN. Peritoneal dissemination often distresses patients, manifesting as abdominal distension, constipation, pain, nausea, and vomiting. As the disease progresses, the patient's quality of life (QOL) can be impaired as a result of poor oral intake due to bowel obstruction, jaundice resulting from bile duct stenosis, and hydronephrosis caused by ureteral constriction. While such treatment decisions must align with the patient's overall health and condition, this strategy has the potential to control the disease, extend survival, and enhance the QOL in patients with SMJN.

The current study has limitations; as this is a report on just one patient's outcome, the results may not be generalizable to the broader population of patients with SMJNs originating from breast cancer. Additional studies with larger sample sizes are needed. Future studies could investigate potential biomarkers or clinical factors that could help identify SMJN patients who are more likely to respond positively to aggressive treatment approaches, similar to the case presented. This could lead to more personalized treatment plans.

#### CONCLUSION

This case study aimed to identify the optimal strategy for achieving extended survival outcomes in patients with SMJN through comprehensive treatment. We presented a case of the longest survival in a patient after undergoing a multidisciplinary treatment regimen. Our findings underscore the significance of adopting a multimodal treatment approach comprising timely and wide excision along with adjunctive therapy. This approach can control the disease, prolong survival, and improve the QOL in patients with SMJN.

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