World J Clin Cases 2022 November 26; 10(33): 12066-12461





#### **Contents**

Thrice Monthly Volume 10 Number 33 November 26, 2022

#### **MINIREVIEWS**

12066 Review of risk factors, clinical manifestations, rapid diagnosis, and emergency treatment of neonatal perioperative pneumothorax

Zhang X, Zhang N, Ren YY

#### **ORIGINAL ARTICLE**

#### **Clinical and Translational Research**

12077 Integrative analysis of platelet-related genes for the prognosis of esophageal cancer

Du QC, Wang XY, Hu CK, Zhou L, Fu Z, Liu S, Wang J, Ma YY, Liu MY, Yu H

12089 Comprehensive analysis of the relationship between cuproptosis-related genes and esophageal cancer

Xu H, Du QC, Wang XY, Zhou L, Wang J, Ma YY, Liu MY, Yu H

12104 Molecular mechanisms of Baihedihuang decoction as a treatment for breast cancer related anxiety: A network pharmacology and molecular docking study

Li ZH, Yang GH, Wang F

12116 Single-cell RNA-sequencing combined with bulk RNA-sequencing analysis of peripheral blood reveals the characteristics and key immune cell genes of ulcerative colitis

Dai YC, Qiao D, Fang CY, Chen QQ, Que RY, Xiao TG, Zheng L, Wang LJ, Zhang YL

#### **Retrospective Study**

12136 Diagnosis and treatment of tubal endometriosis in women undergoing laparoscopy: A case series from a single hospital

Jiao HN, Song W, Feng WW, Liu H

12146 Different positive end expiratory pressure and tidal volume controls on lung protection and inflammatory factors during surgical anesthesia

Wang Y, Yang Y, Wang DM, Li J, Bao QT, Wang BB, Zhu SJ, Zou L

12156 Transarterial chemoembolization combined with radiofrequency ablation in the treatment of large hepatocellular carcinoma with stage C

Sun SS, Li WD, Chen JL

12164 Coexistence of anaplastic lymphoma kinase rearrangement in lung adenocarcinoma harbouring epidermal growth factor receptor mutation: A single-center study

Zhong WX, Wei XF

#### Contents

#### Thrice Monthly Volume 10 Number 33 November 26, 2022

#### **Observational Study**

Prognostic values of optic nerve sheath diameter for comatose patients with acute stroke: An observational 12175 study

Zhu S, Cheng C, Wang LL, Zhao DJ, Zhao YL, Liu XZ

12184 Quality of care in patients with inflammatory bowel disease from a public health center in Brazil

Takamune DM, Cury GSA, Ferrás G, Herrerias GSP, Rivera A, Barros JR, Baima JP, Saad-Hossne R, Sassaki LY

12200 Comparison of the prevalence of sarcopenia in geriatric patients in Xining based on three different diagnostic criteria

Pan SQ, Li XF, Luo MQ, Li YM

#### **Prospective Study**

12208 Predictors of bowel damage in the long-term progression of Crohn's disease

Fernández-Clotet A, Panés J, Ricart E, Castro-Poceiro J, Masamunt MC, Rodríguez S, Caballol B, Ordás I, Rimola J

#### **Randomized Controlled Trial**

12221 Protective effect of recombinant human brain natriuretic peptide against contrast-induced nephropathy in elderly acute myocardial infarction patients: A randomized controlled trial

Zhang YJ, Yin L, Li J

#### **META-ANALYSIS**

12230 Prognostic role of pretreatment serum ferritin concentration in lung cancer patients: A meta-analysis

Gao Y, Ge JT

#### **CASE REPORT**

12240 Non-surgical management of dens invaginatus type IIIB in maxillary lateral incisor with three root canals and 6-year follow-up: A case report and review of literature

Arora S, Gill GS, Saquib SA, Saluja P, Baba SM, Khateeb SU, Abdulla AM, Bavabeedu SS, Ali ABM, Elagib MFA

Unusual presentation of Loeys-Dietz syndrome: A case report of clinical findings and treatment challenges 12247

Azrad-Daniel S, Cupa-Galvan C, Farca-Soffer S, Perez-Zincer F, Lopez-Acosta ME

12257 Peroral endoscopic myotomy assisted with an elastic ring for achalasia with obvious submucosal fibrosis: A case report

Wang BH, Li RY

12261 Subclavian brachial plexus metastasis from breast cancer: A case report

Zeng Z, Lin N, Sun LT, Chen CX

12268 Case mistaken for leukemia after mRNA COVID-19 vaccine administration: A case report

Lee SB, Park CY, Park SG, Lee HJ

Orthodontic-surgical treatment of an Angle Class II malocclusion patient with mandibular hypoplasia and 12278 missing maxillary first molars: A case report

Π

Li GF, Zhang CX, Wen J, Huang ZW, Li H

#### Contents

#### Thrice Monthly Volume 10 Number 33 November 26, 2022

12289 Multiple cranial nerve palsies with small angle exotropia following COVID-19 mRNA vaccination in an adolescent: A case report

Lee H, Byun JC, Kim WJ, Chang MC, Kim S

12295 Surgical and nutritional interventions for endometrial receptivity: A case report and review of literature Hernández-Melchor D, Palafox-Gómez C, Madrazo I, Ortiz G, Padilla-Viveros A, López-Bayghen E

12305 Conversion therapy for advanced penile cancer with tislelizumab combined with chemotherapy: A case report and review of literature

Long XY, Zhang S, Tang LS, Li X, Liu JY

Endoscopic magnetic compression stricturoplasty for congenital esophageal stenosis: A case report 12313 Liu SQ, Lv Y, Luo RX

12319 Novel hydroxymethylbilane synthase gene mutation identified and confirmed in a woman with acute intermittent porphyria: A case report

Zhou YQ, Wang XQ, Jiang J, Huang SL, Dai ZJ, Kong QQ

12328 Modified fixation for periprosthetic supracondylar femur fractures: Two case reports and review of the literature

Li QW, Wu B, Chen B

12337 Erbium-doped yttrium aluminum garnet laser and advanced platelet-rich fibrin+ in periodontal diseases: Two case reports and review of the literature

Tan KS

12345 Segmental artery injury during transforaminal percutaneous endoscopic lumbar discectomy: Two case

Cho WJ, Kim KW, Park HY, Kim BH, Lee JS

12352 Pacemaker electrode rupture causes recurrent syncope: A case report

Zhu XY, Tang XH, Huang WY

12358 Hybrid intercalated duct lesion of the parotid: A case report

Stankevicius D, Petroska D, Zaleckas L, Kutanovaite O

12365 Clinical features and prognosis of multiple myeloma and orbital extramedullary disease: Seven cases report and review of literature

Hu WL, Song JY, Li X, Pei XJ, Zhang JJ, Shen M, Tang R, Pan ZY, Huang ZX

12375 Colon mucosal injury caused by water jet malfunction during a screening colonoscopy: A case report

Patel P, Chen CH

12380 Primary malignant pericardial mesothelioma with difficult antemortem diagnosis: A case report

Oka N, Orita Y, Oshita C, Nakayama H, Teragawa H

12388 Typical imaging manifestation of neuronal intranuclear inclusion disease in a man with unsteady gait: A case report

Ш

Gao X, Shao ZD, Zhu L

#### **Contents**

#### Thrice Monthly Volume 10 Number 33 November 26, 2022

12395 Multimodality imaging and treatment of paranasal sinuses nuclear protein in testis carcinoma: A case

Huang WP, Gao G, Qiu YK, Yang Q, Song LL, Chen Z, Gao JB, Kang L

12404 T1 rectal mucinous adenocarcinoma with bilateral enlarged lateral lymph nodes and unilateral metastasis: A case report

Liu XW, Zhou B, Wu XY, Yu WB, Zhu RF

12410 Influence of enhancing dynamic scapular recognition on shoulder disability, and pain in diabetics with frozen shoulder: A case report

Mohamed AA

12416 Acute myocardial necrosis caused by aconitine poisoning: A case report

Liao YP, Shen LH, Cai LH, Chen J, Shao HQ

12422 Danggui Sini decoction treatment of refractory allergic cutaneous vasculitis: A case report

Chen XY, Wu ZM, Wang R, Cao YH, Tao YL

12430 Phlegmonous gastritis after biloma drainage: A case report and review of the literature

Yang KC, Kuo HY, Kang JW

12440 Novel TINF2 gene mutation in dyskeratosis congenita with extremely short telomeres: A case report

Picos-Cárdenas VJ, Beltrán-Ontiveros SA, Cruz-Ramos JA, Contreras-Gutiérrez JA, Arámbula-Meraz E, Angulo-Rojo C, Guadrón-Llanos AM, Leal-León EA, Cedano-Prieto DM, Meza-Espinoza JP

12447 Synchronous early gastric and intestinal mucosa-associated lymphoid tissue lymphoma in a Helicobacter pylori-negative patient: A case report

Lu SN, Huang C, Li LL, Di LJ, Yao J, Tuo BG, Xie R

#### **LETTER TO THE EDITOR**

12455 Diagnostic value of metagenomics next-generation sequencing technology in disseminated strongyloidiasis

ΙX

Song P, Li X

12458 Diagnostic value of imaging examination in autoimmune pancreatitis

Wang F, Peng Y, Xiao B

#### Contents

#### Thrice Monthly Volume 10 Number 33 November 26, 2022

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CASE REPORT

## T1 rectal mucinous adenocarcinoma with bilateral enlarged lateral lymph nodes and unilateral metastasis: A case report

Xian-Wei Liu, Bing Zhou, Xiao-Yu Wu, Wen-Bing Yu, Ren-Fang Zhu

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#### **Abstract**

#### **BACKGROUND**

There are a few cases of lateral lymph node (LLN) metastasis (LLNM) of T1 rectal cancer. Moreover, LLNM is easily missed, especially in patients with early-stage rectal cancer. To our knowledge, the possibility of bilateral LLNM before surgery has not been reported in previous studies.

#### CASE SUMMARY

A 36-year-old woman underwent endoscopic submucosal dissection at a local hospital owing to a clinical diagnosis of a rectal polyp. The pathology report showed a diagnosis of T1 rectal mucinous adenocarcinoma. She was considered to have bilateral LLNM after the examination at our hospital. Laparoscopic total mesorectal excision plus bilateral LLN dissection was performed and the pathological outcomes indicated unilateral LLNM. The patient received longcourse adjuvant chemoradiotherapy with no recurrence or metastasis observed during the 1-year follow-up period.

#### **CONCLUSION**

T1 rectal cancer could lead to LLNM and possibly, bilateral LLNM. Therefore, adequate clinical evaluation is essential for these patients.

Key Words: T1 rectal cancer; Lateral lymph node metastasis; Lateral lymph node dissection; Brief literature review; Endoscopic submucosal dissection; Case report

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Core Tip: T1 rectal cancer is rarely accompanied by lymph node metastases, and even fewer lateral lymph node (LLN) metastases (LLNM). To our knowledge, the published case reports to date have mainly reported cases of heterochronous LLNM, only two cases with simultaneous metastases, and only one case of missed LLNM after endoscopic submucosal dissection (ESD). For patients with no residual tumor after ESD, in whom LLNM is suspected, it is also inconclusive whether only LLN dissection could be performed.

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#### INTRODUCTION

The probability of lateral lymph node (LLN) metastasis (LLNM) in locally advanced rectal cancer is 10%-25%[1-3], whereas the corresponding probability of bilateral LLNM is only 2.8%-3.5%[1]. It has been reported that the rate of pelvic extra-regional LNM of T1 rectal cancer is 5.4%[4]. Previous studies have reported that the incidence of LLNM in patients with T1 rectal cancer ranges from 0.5% to 0.9%[5]. However, there have been limited studies in this field, therefore, and more studies are needed. Among the five case reports published to date [6-10], three had focused on heterochronous metastasis and two had reported on simultaneous metastasis. Moreover, four cases had unilateral solitary LLNM (only one LLN had cancer metastasis), and one case had unilateral multiple LLNM (several LLNs at one side had cancer metastasis). In one patient, LLNM was suspected to have been missed before total mesorectal excision (TME) and was found 6 mo postoperatively. In another patient, LLNM was missed before endoscopic submucosal dissection (ESD). The detailed information of the five cases is presented in Table 1. To our knowledge, we report the second case, in which LLNM was missed before ESD, and the first case, where bilateral LLNM was suspected.

#### CASE PRESENTATION

#### Chief complaints

A 36-year-old woman underwent a routine physical examination at a local hospital.

#### History of present illness

Colonoscopy revealed a polypoid lesion with a diameter of 1.5 cm, and the lower edge was located 3 cm from the anal verge. Because the lesion was more likely to be a polyp, ESD was performed at that time. Pathology revealed a mucinous adenocarcinoma, with moderate differentiation and submucosal invasion. There was no residual tumor at the basal edge or in the resection mucosa. The patient visited our hospital immediately for consultation on whether further treatment was needed.

#### History of past illness

The woman was previously in good health.

#### Personal and family history

The patient's family history was unremarkable.

#### Physical examination

Physical examination revealed no abnormalities except several metal clips that could be palpated on digital rectal examination.

#### Laboratory examinations

We evaluated the carcinoembryonic antigen and carbohydrate antigen-199 levels, which were found to be 2.54 ng/mL and 12.73 ku/L, respectively. Our pathologists also confirmed the pathologic finding of the local hospital.

#### Imaging examinations

We performed abdominal contrast-enhanced computed tomography (CT) and chest CT for this patient, in addition to a pathological consultation. Magnetic resonance imaging of the pelvis was not performed

Table 1 The detailed information of the previous five cases and our case						
	Case 1	Case 2	Case 3	Case 4	Case 5	Case 6
Ref.	Hara et al[6], 2008	Sueda <i>et al</i> [7], 2013	Ogawa et al[8], 2016	Tanishima et al[9], 2017	Zhang et al[10], 2020	Ours, 2022
Sex	Male	Female	Female	Male	Male	Female
Age (yr)	61	41	35	56	45	36
Risk factors of LNM						
Depth of invasion (µm)	-	-	3000	Head invasion	-	-
Histological type	Well	Moderately	Moderately	Moderately	Well	Moderately
Budding	-	-	-	1	-	-
Time after 1 <sup>st</sup> surgery (mo)	22	6	Simultaneous	6	Simultaneous	Simultaneous
Treatment	Extended, LLND	Extended, LLND	TME + LLND	LLND	TME + LLND	TME + LLND
Bilateral or unilateral LLND	Unilateral	Unilateral	Unilateral	Unilateral	Unilateral	Bilateral
Isolated or multiple LLNM	Isolated	Isolated	Isolated	Multiple	Isolated	Multiple
Adjuvant therapy after LLND	None	None	Tegafururacil + leucovorin	FOLFOX	XELOX + radiotherapy	XELOX + radiotherapy
Follow-up period (mo)	44	12	48	30	10	12
Prognosis	Alive	Alive	Alive	Alive	Alive	Alive

LNM: Lymph node metastasis; TME: Total mesorectal excision; LLND: Lateral lymph node dissection; LLNM: Lateral lymph node metastasis; FOLFOX: Fluorouracil, leucovorin, and oxaliplatin; XELOX: Oxaliplatin plus capecitabine.



Figure 1 Computed tomography findings. A: Computed tomography (CT) imaging findings for the two left enlarged lateral lymph nodes; B: CT imaging findings for the right enlarged lateral lymph node; C: CT imaging findings for the largest mesorectal lymph node (arrow).

because there was no residual tumor. The chest CT scan revealed no sign of distant metastasis. However, the abdominal contrast-enhanced CT showed suspected enlarged bilateral LLNs and some enlarged mesorectal LNs (MLN) (Figure 1). Among these, there were two enlarged LLNs in the left lateral area, both located in the distal internal iliac region (263D)[11], with short-axis diameters of 7.1 mm and 6.2 mm, respectively. An enlarged LLN was noted in the right lateral area (distal internal iliac region), with a short-axis diameter of 5.3 mm. These three LLNs had at least two of three malignant features, including a round, irregular border and shape, and heterogeneous density, consistent with a positive LN diagnosis[12]. The maximum short-axis diameter of the enlarged MLN was 4.8 mm, which was insufficient for a diagnosis of LNM.

#### FURTHER DIAGNOSTIC WORK-UP

After discussions with the multidisciplinary team and after obtaining the consent of the patient and her family, laparoscopic TME with bilateral lateral lymph node dissection (LLND) was performed. The

patient recovered uneventfully and was hospitalized for 10 d with no complications. There was no residual tumor in the rectal specimen and the distal and proximal margins were both negative; only partial mucosal hemorrhage with scattered acute and chronic inflammatory cell infiltration was noted. A total of 31 LNs were harvested, including nine MLNs, 13 left LLNs, and nine right LLNs. The MLNs and right LLNs were confirmed to have no metastasis, but the two enlarged left LLNs were confirmed to have metastasis (Figure 2).

#### FINAL DIAGNOSIS

T1 rectal mucinous adenocarcinoma with simultaneous LLNM.

#### **TREATMENT**

The patient received chemotherapy involving six cycles of XELOX (oxaliplatin plus capecitabine) and radiotherapy (50.4 Gy; 28 fractions).

#### OUTCOME AND FOLLOW-UP

There was no sign of recurrence or metastasis, and the patient's urinary and sexual functions were normal during the 1-year follow-up.

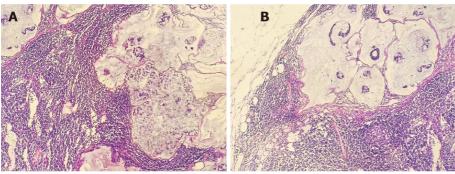
#### DISCUSSION

A meta-analysis suggested that there is only a 1.9% risk of LNM in T1 rectal cancers with low-risk criteria[13], which was defined as submucosal invasion of  $\leq 1000 \, \mu \text{m}$  without poor differentiation, lymphovascular invasion, or budding [14]. Thus, the probability of LLNM is even lower in this type of cancer. Most patients with T1 rectal cancer with a diameter of  $\leq 2$  mm can be cured with ESD[15]. However, the main problem is that it is often difficult to assess the low-risk criteria prior to ESD[16]. Moreover, LLNM is easily missed, especially in patients with early-stage rectal cancer [9].

In our case, the patient achieved the standard clinical cure because she had no risk factors. If the patient did not take the initiative to come to our hospital for a consultation, the LLNM would most likely have been missed. Moreover, she was even suspected to have bilateral LLNM, although it was finally confirmed that she had only unilateral LLNM. Patients with T1 rectal cancer are at the risk of LLNM and may even have bilateral LLNM. ESD may guarantee a clinical cure for the majority of patients with T1 rectal cancer. However, a few patients may need more in-depth clinical examinations to ensure that there is no missed diagnosis, not only for LLNM, but also for liver or lung metastasis [17,18]. Therefore, we suggest that if the relevant examinations cannot be completed before ESD for evaluating the patient's condition, these examinations should be completed after ESD and according to the standard management for advanced rectal cancer, even if the patients are pathologically diagnosed with T1 stage and have no risk factors. Such an approach would avoid the possibility of missing LLNM or distant metastases

At present, for advanced rectal cancer with suspected LLNM, although the National Comprehensive Cancer Network guidelines recommend neoadjuvant chemoradiotherapy (nCRT) combined with TME [19], and Japanese guidelines recommend preventive LLND combined with TME[11], neither method can effectively control the lateral local recurrence [20]. Therefore, more and more scholars recommend nCRT combined with TME and selective LLND with the in-depth clinical research[21]. However, there is no guideline or consensus for the treatment of T1 rectal cancer with suspected LLNM. Moreover, the prognosis of advanced rectal cancer with LLNM is significantly worse than that without LLNM, therefore, LLND is necessary[22]. However, no prognostic data have been reported in T1 rectal cancer with LLNM because not enough cases have been reported. Furthermore, we believe that LLND is important in T1 rectal cancer with suspected LLNM according to the prognostic data of advanced rectal cancer.

In addition, in view of the report of Zhang et al[10] and the present study, although simultaneous LLNM was considered after ESD, no residual tumor or mesorectal LNM was observed after TME and LLND. Therefore, LLND may only be possible in such patients (without considering residual tumor and mesorectal LNM) after adequate assessment of the patient's status. Instead, the "watch and wait" strategy can be implemented for the primary tumor and MLN. However, we have no clinical practice experience regarding this, and our theory needs to be confirmed by more cases or multicenter studies.



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Figure 2 Pathological imaging for the two positive left lateral lymph nodes (hematoxylin and eosin × 100). A and B: Pathological images of two positive lateral lymph nodes, respectively. The lymphatic structure is destroyed, and numerous mucous lakes are formed in which floating adenosine cells are observed.

#### CONCLUSION

T1 rectal cancer could lead to LLNM and possibly, bilateral LLNM. Therefore, adequate clinical evaluation is essential in patients with this type of cancer.

#### **FOOTNOTES**

Author contributions: Liu XW, Zhou B and Wu XY participated in data collection and manuscript writing; Liu XW, Yu WB and Zhu RF completed the surgery; and all authors read and approved the final manuscript.

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