

## Case report

# Ascending colon cancer associated with deposited ova of *Schistosoma japonicum* in non-endemic area



Kensuke Nakatani, M.D.<sup>a</sup>, Takaharu Kato, M.D., Ph.D.<sup>a,b,\*</sup>, Shinichiro Okada, M.D., Ph.D.<sup>a</sup>, Risa Matsumoto, M.D.<sup>a</sup>, Kazuhiro Nishida, M.D.<sup>a</sup>, Hiroyasu Komuro, M.D.<sup>a</sup>, Maki Iida, M.D., Ph.D.<sup>c</sup>, Shiro Tsujimoto, M.D., Ph.D.<sup>c</sup>, Toshiyuki Suganuma, M.D., Ph.D.<sup>a</sup>

<sup>a</sup> Department of surgery, Yokosuka general hospital Uwamachi, 2-36 Uwamachi Yokosuka City, Kanagawa, 238-8567, Japan

<sup>b</sup> Department of surgery, Saitama Medical Center, Jichi Medical University, 1-847 Amanuma-cho, Omiya-ku, Saitama 330-8503, Japan

<sup>c</sup> Department of Pathology, Yokosuka general hospital Uwamachi, 2-36 Uwamachi, Yokosuka, Kanagawa 238-8567, Japan

## ARTICLE INFO

## Article history:

Received 6 September 2016

Received in revised form 16 September 2016

Accepted 16 September 2016

## Keywords:

Schistosomiasis

*Schistosoma japonicum*

Colorectal cancer

Non-endemic area

## ABSTRACT

Some reports suggest the positive correlation between *Schistosoma japonicum* infection and colorectal cancer, however the sufficient evidence that supports a causal relationship between them has not been established. Japan used to be an endemic area of *S. japonicum* infection for 40 years ago. But now all of Japan is a non-endemic area of *S. japonicum* infection. We report a case of ascending colon cancer associated with deposited ova of *S. japonicum* in non-endemic area.

© 2016 The Authors. Published by Elsevier Ltd. This is an open access article under the CC BY-NC-ND license (<http://creativecommons.org/licenses/by-nc-nd/4.0/>).

## Introduction

Colorectal cancer (CRC) is one of the most prevalent malignancies in developed countries [1]. The incidence of CRC has increased over the last few decades by two to four-fold in Asian countries [2–4]. *Schistosoma japonicum* which is common in Southeast Asia [5], is regarded as a risk factor of CRC development, but the causal relation between *S. japonicum* and CRC has not been established. We present a case who developed ascending colon cancer with deposited ova of *S. japonicum*. All of Japan is a non-endemic area of *S. japonicum* infection, but used to be an endemic area for 40 years ago.

## Presentation of case

A 90-year-old woman had noticed abdominal pain and appetite loss for several months. She consulted a local clinic and anemia was

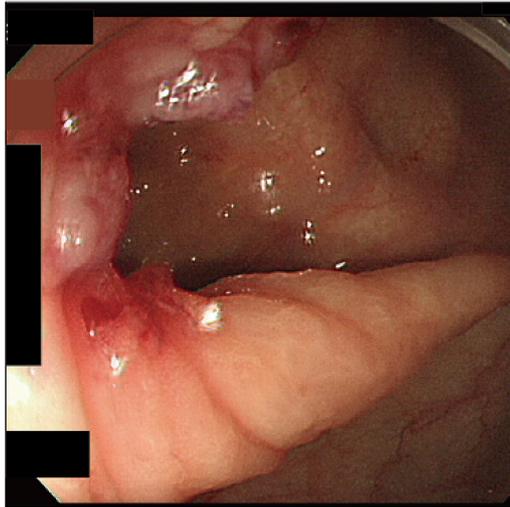
detected. She was referred to Yokosuka general hospital Uwamachi for further examination and was diagnosed with ascending colon cancer by colonoscopy, but she did not want to receive any treatment. She did not have any familial history of liver diseases or malignant diseases.

One year later, she felt severe appetite loss and vomited several times she came to our hospital again. Her heart rate was 82 per minute, blood pressure was 170/87 mmHg, respiratory rate was 16 per minute on arrival. A tumor was palpable in her right lower quadrant but she did not have rebound tenderness in her abdomen. No lymph nodes were palpable. The CEA serum tumor marker level was elevated to 89.3 ng/dl but CA19-9 was within the normal range. Colonoscopic examination revealed a type 2 tumor in the ascending colon (Fig. 1). Abdominal computed tomography (CT) revealed a 5 cm tumor enhanced heterogeneously located in the ascending colon (Fig. 2A). The small intestine was dilated. There was no evidence of ascites, but were swollen lymph nodes around the colon tumor and liver metastases in the segment III (Fig. 2B). The patient was diagnosed with ascending colon cancer T2 N1 M1 stage IV according to the tumor, node, and metastasis (TNM) classification, of the American Joint Committee on Cancer (7th edition) [6]. She underwent ileocecal resection as a palliative surgery.

Macroscopically, the size of the tumor was 40 × 63 mm. (Fig. 3A). Histological findings showed that the tumor was moderately differentiated adenocarcinoma (Fig. 3B). Additionally,

\* Corresponding author at: Department of Surgery, Saitama Medical Center, Jichi Medical University, 1-847 Amanuma-cho, Omiya-ku, Saitama, Saitama 330-8503, Japan.

E-mail addresses: [kensuken@jadecom.jp](mailto:kensuken@jadecom.jp) (K. Nakatani), [tkato@jichi.ac.jp](mailto:tkato@jichi.ac.jp) (T. Kato), [okada@jichi.ac.jp](mailto:okada@jichi.ac.jp) (S. Okada), [risamatsumoto1217@gmail.com](mailto:risamatsumoto1217@gmail.com) (R. Matsumoto), [pandadebanda4@gmail.com](mailto:pandadebanda4@gmail.com) (K. Nishida), [hiroyasu\\_101819@yahoo.co.jp](mailto:hiroyasu_101819@yahoo.co.jp) (H. Komuro), [maki14f@ybb.ne.jp](mailto:maki14f@ybb.ne.jp) (M. Iida), [tujimts@jadecom.jp](mailto:tujimts@jadecom.jp) (S. Tsujimoto), [toshiyukis@jadecom.jp](mailto:toshiyukis@jadecom.jp) (T. Suganuma).



**Fig. 1.** Colonoscopy revealed type 2 tumor in the ascending colon, biopsy specimen revealed tubular adenocarcinoma.

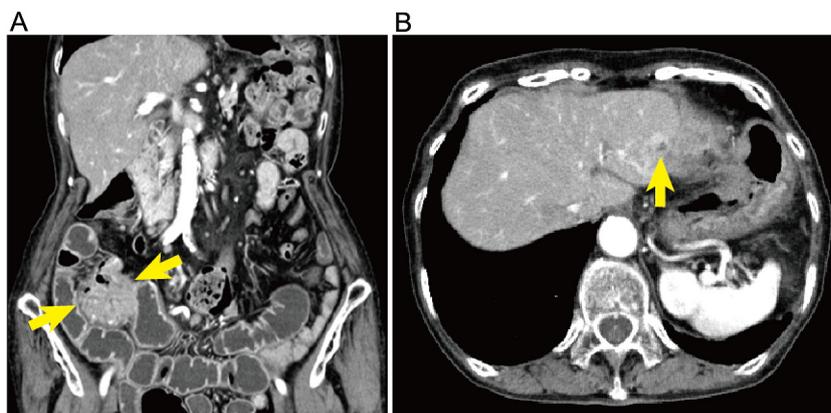
there were numbers of schistosomal ova in the muscular layer surround tumor (Fig. 3C) and also in the submucosa of adjacent mucosa (Fig. 3D). Postoperative course of the patient was complicated by a surgical site infection, and she was discharged from the hospital 30 days after the surgery. The patient has survived 2 years after surgery with liver metastasis without receiving any chemotherapy.

## Discussion

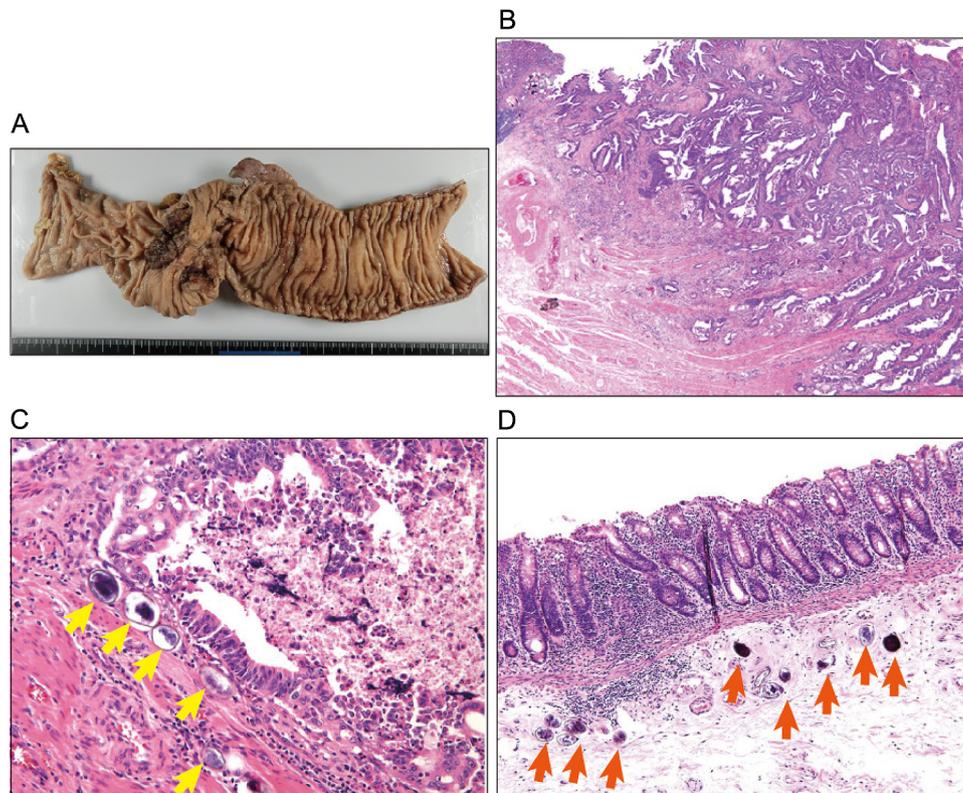
We reported a patient who underwent surgery for ascending colorectal cancer associated with deposited ova of *S. japonicum* in the resected colon. The last patient with deposited ova of schistosoma japonicum was reported in 1978 in Japan. The presented patient had lived Yamanashi prefecture in Japan for 70 years ago where schistosomiasis was an endemic parasitic disease in 1940s [7]. All of Japan is a non-endemic area of *S. japonicum* infection. Even in patients living in non-endemic area, a history of travel as well as where the patient has previously lived is important.

In IARC classification, *S. japonicum* is regarded as a carcinogenic risk to humans as attributed to group 2B (possibly carcinogenic) [8]. The consensus of available pathological data implicates an association between *S. japonicum* infestation and induction of colorectal cancer. Several reports in Japan and China have been published about the positive correlation between *S. japonicum* infection and colorectal carcinoma [9–13]. Recent molecular analyses have suggested the association between *S. japonicum* and colorectal cancer. Zhang et al. studied the mutation pattern in the TP53 gene in *S. japonicum*-associated rectal cancer that showed the majority of mutation in TP53 gene were detected in exon 7 in schistosomal group compared to exon 5 in non-schistosomal group [14]. Another study showed that *S. japonicum* ova-induced colorectal epithelial proliferative polyps have a high percentage of atypical hyperplasia (64.9%) and elevated CEA (90%) [15].

The relationship, however, between schistosomiasis and colorectal cancer has been debated for decades. If there is an increase in the risk of colorectal cancer, it is small [16]. Further epidemiological and molecular analyses are needed to clarify the cause and effect relationship between *S. japonicum* and colorectal cancer carcinogenesis.



**Fig. 2.** Abdominal CT revealed an irregular mass enhanced heterogeneously in the right side colon with size of 5 × 4 cm (A) and a liver tumor in the segment III (B).



**Fig. 3.** Resected specimen showed a type 2 tumor with the size of 40 × 63 mm in the ascending colon (A). Histological analysis of the resected specimen showed moderately differentiated adenocarcinoma invaded into sub serous layer (×100 H.E.) (B). The several ova of *S. japonicum* was detected in the muscular layer surround the tumor and sub mucosa in the adjacent colon (arrows in C and D).

## References

- [1] Jemal A, Bray F, Center MM, Ferlay J, Ward E, Forman D. Global cancer statistics. *CA Cancer J Clin* 2011;61:69–90.
- [2] Sung JJ, Lau JY, Goh KL, Leung WK, and Asia Pacific Working Group on Colorectal C: Increasing incidence of colorectal cancer in Asia: implications for screening. *Lancet Oncol* 2005;6:871–6.
- [3] Yee YK, Tan VP, Chan P, Hung IF, Pang R, Wong BC. Epidemiology of colorectal cancer in Asia. *J Gastroenterol Hepatol* 2009;24:1810–6.
- [4] Yiu HY, Whittemore AS, Shibata A. Increasing colorectal cancer incidence rates in Japan. *Int J Cancer* 2004;109:777–81.
- [5] Gryseels B, Polman K, Clerinx J, Kestens L. Human schistosomiasis. *Lancet* 2006;368:1106–18.
- [6] Sobin LH, Gospodarowicz M, Wittekind C. International Union Against Cancer (UICC) TNM classification of malignant tumours, vol. 7. New York: Wiley-Liss; 2010.
- [7] Inaba Y. A cohort study on the causes of death in an endemic area of Schistosomiasis japonica in Japan. *Ann Acad Med Singapore* 1984;13:142–8.
- [8] International Agency for Research on Cancer WHO: IARC working group on the evaluation of carcinogenic risks to humans. *IARC Monogr Eval Carcinog Risks Hum* 1994;61:45–119.
- [9] Shindo K. Significance of Schistosomiasis japonica in the development of cancer of the large intestine: report of a case and review of the literature. *Dis Colon Rectum* 1976;19:460–9.
- [10] Zhao ES. Cancer of the colon and schistosomiasis. *J R Soc Med* 1981;74:645.
- [11] Xu Z, Su DL. Schistosoma japonicum and colorectal cancer: an epidemiological study in the People's Republic of China. *Int J Cancer* 1984;34:315–8.
- [12] Matsuda K, Masaki T, Ishii S, Yamashita H, Watanabe T, Nagawa H, et al. Possible associations of rectal carcinoma with Schistosoma japonicum infection and membranous nephropathy: a case report with a review. *Jpn J Clin Oncol* 1999;29:576–81.
- [13] Qiu DC, Hubbard AE, Zhong B, Zhang Y, Spear RC. A matched, case-control study of the association between Schistosoma japonicum and liver and colon cancers, in rural China. *Ann Trop Med Parasitol* 2005;99:47–52.
- [14] Zhang R, Takahashi S, Orita S, Yoshida A, Maruyama H, Shirai T, et al. p53 gene mutations in rectal cancer associated with schistosomiasis japonica in Chinese patients. *Cancer Lett* 1998;131:215–21.
- [15] Yu XR, Chen PH, Xu JY, Xiao S, Shan ZJ, Zhu SJ. Histological classification of schistosomal egg induced polyps of colon and their clinical significance. An analysis of 272 cases. *Chin Med J (Engl)* 1991;104:64–70.
- [16] Ross AG, Bartley PB, Sleight AC, Olds GR, Li Y, Williams GM, et al. Schistosomiasis. *N Engl J Med* 2002;346:1212–20.