Successful treatment of a giant ossified benign mesenteric schwannoma

Ying-Sheng Wu, Shao-Yan Xu, Jing Jin, Ke Sun, Zhen-Hua Hu, Wei-Lin Wang

Ying-Sheng Wu, Shao-Yan Xu, Jing Jin, Zhen-Hua Hu, Wei-Lin Wang, Division of Hepatobiliary and Pancreatic Surgery, Department of Surgery, First Affiliated Hospital, School of Medicine, Zhejiang University, Hangzhou 310003, Zhejiang Province, China

Ying-Sheng Wu, Shao-Yan Xu, Jing Jin, Zhen-Hua Hu, Wei-Lin Wang, Key Laboratory of Combined Multi-Organ Transplantation, Ministry of Public Health, Hangzhou 310003, Zhejiang Province, China

Ying-Sheng Wu, Shao-Yan Xu, Jing Jin, Zhen-Hua Hu, Wei-Lin Wang, Key Laboratory of Organ Transplantation, Hangzhou 310003, Zhejiang Province, China

Ying-Sheng Wu, Shao-Yan Xu, Jing Jin, Zhen-Hua Hu, Wei-Lin Wang, Key Laboratory of Precision Diagnosis and Treatment for Hepatobiliary and Pancreatic Tumor of Zhejiang Province, Hangzhou 310003, Zhejiang Province, China

Ying-Sheng Wu, Shao-Yan Xu, Jing Jin, Zhen-Hua Hu, Wei-Lin Wang, Collaborative Innovation Center for Diagnosis and Treatment of Infectious Diseases, First Affiliated Hospital, School of Medicine, Zhejiang University, Hangzhou 310003, Zhejiang Province, China

Ke Sun, Department of Pathology, First Affiliated Hospital, School of Medicine, Zhejiang University, Hangzhou 310003, Zhejiang Province, China

ORCID number: Ying-Sheng Wu (0000-0002-9406-9169); Shao-Yan Xu (0000-0001-8016-0917); Jing Jin (0000-0001-8706-4979); Ke Sun (0000-0002-3789-3143); Zhen-Hua Hu (0000-0002-8636-0757); Wei-Lin Wang (0000-0001-9432-2649).

Author contributions: Wu YS and Xu SY contributed equally to this work; Wu YS, Xu SY and Jing Jin collected case data, prepared the photos and wrote the manuscript; Sun K proofread the pathologic materials; Hu ZH and Wang WL proofread and revised the manuscript; all of the authors approved the final version to be published.

Supported by the National Natural Science Foundation of China, No. 81572307.

Informed consent statement: Informed consent was obtained from the patient.

Conflict-of-interest statement: The authors declare that there is no conflict of interest related to this report.

Open-Access: This article is an open-access article which was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/

Manuscript source: Unsolicited manuscript

Correspondence to: Wei-Lin Wang, PhD, MD, Division of Hepatobiliary and Pancreatic Surgery, Department of Surgery, First Affiliated Hospital, School of Medicine, Zhejiang University, 79# Qingchun road, Hangzhou 310003, Zhejiang Province, China. wam@zju.edu.cn

Telephone: +86-571-87236466
Fax: +86-571-87236466

Received: August 30, 2017
Peer-review started: August 31, 2017
First decision: September 20, 2017
Revised: October 3, 2017
Accepted: October 26, 2017
Article in press: October 26, 2017
Published online: January 14, 2018

Abstract

Primary benign schwannoma of the mesentery is extremely rare. To date, only 9 cases have been reported in the English literature, while mesenteric schwannoma with ossified degeneration has not been reported thus far. In the present study, we present the first giant ossified benign mesenteric schwannoma in a 58-year-
old female. Ultrasound, computed tomography and magnetic resonance imaging were used, but it was still difficult to determine the definitive location and diagnose the mass. By laparotomy, a 10.0 cm × 9.0 cm × 9.0 cm giant mass was found in the mesentery and was then completely resected. Microscopically, the tumour located in the mesentery mainly consisted of spindle-shaped cells with a palisading arrangement. Some areas of the tumour were ossified, and a true metaplastic bone formation was observed, with the presence of bone lamellae and osteoblasts. Immunohistochemical investigation of the tumour located in the mesentery showed that the staining for the S-100 protein was strongly positive, while the stainings of SMA, CD34, CD117 and DOG-1 were negative. The cell proliferation index, measured with Ki67 staining, was less than 3%. Finally, a giant ossified benign mesenteric schwannoma was diagnosed. After surgery, the patient was followed up for a period of 43 mo, during which she remained well, with no evidence of tumour recurrence.

**Key words:** Schwannoma; Mesentery; Ossification; Laparotomy; S-100

© The Author(s) 2018. Published by Baishideng Publishing Group Inc. All rights reserved.

**Core tip:** To date, only 9 cases of mesenteric schwannomas have been reported in the English literature; an ossified mesenteric schwannoma has not been reported. In the present study, we present the first giant ossified benign mesenteric schwannoma. It was challenging to determine the location and obtain a precise diagnosis of the mesenteric schwannoma prior to surgery. We completely resected the mesenteric schwannoma by laparotomy. In this paper, a literature review was conducted to deepen our understanding of mesenteric schwannomas.

Wu YS, Xu SY, Jin J, Sun K, Hu ZH, Wang WL. Successful treatment of a giant ossified benign mesenteric schwannoma. *World J Gastroenterol* 2018; 24(2): 303-309 Available from: URL: http://www.wjgnet.com/1007-9327/full/v24/i2/303.htm DOI: http://dx.doi.org/10.3748/wjg.v24.i2.303

**INTRODUCTION**

Schwannomas are mesenchymal neoplasms of the peripheral nerve sheath[1], which mainly develop in young to middle-aged patients with no obvious gender difference. Schwannomas are usually solitary sporadic lesions. About 3% occurred in patients with NF-2, 2% in those with schwannomatisis, and 5% in association with multiple meningiomas with or without NF2. Most of schwannomas, whether sporadic or inherited, display inactivating germine mutations of the tumour suppressor gene *NF2* located on chromosome 22 which encodes the protein Merlin or schwannomin[2], This protein, localized to regions of the cell membrane engaged in cell contact and mobility, is expressed in Schwann cells, meningeal cells and the lens of the eye. The mechanism by which the loss of this protein results in tumorigenesis is not well understood. Schwannomas are usually benign (> 90%) and occupy approximately 5% of benign soft-tissue neoplasms[3,4]. The tumour can sometimes show secondary degenerative changes, including cyst formation, hyalinization, haemorrhage and calcification[5]. Schwannomas usually occur in the head, neck and extremities[6], while schwannomas in the bowel mesentery are extremely rare. To our knowledge, only nine cases of schwannomas located in the bowel mesentery have been reported[7,15]. In addition, mesenteric schwannoma with ossified degeneration has not been reported thus far. In the present study, we present the first giant ossified benign mesenteric schwannoma.

**CASE REPORT**

On January 14, 2014, a 58-year-old woman visited our department due to a lesion detected incidentally in her abdominal cavity during a routine health examination in the local hospital. On physical examination, a lesion was palpable in her upper left abdomen, sized 12 cm × 6 cm, and she felt slight tenderness. Four years previously, she had undergone resection of a lipoma in her back. Blood test findings, including tumour markers, were unremarkable. Ultrasound (US) revealed a solid lesion in the upper left abdomen, with clear margin, and also showed some cystic and strong echo areas in the lesion. Color doppler flow imagings (CDFIs) showed blood signals in the lesion. In a native computed tomography (CT) scan, the lesion in the upper left abdomen appeared well-defined and was 9.6 cm in diameter; while regions of high density were visible, compatible with calcification and/or ossification (Figure 1A). Contrast-enhanced CT study revealed a lesion with slight and inhomogeneous enhancement (Figure 1B). On T1-weighted images, the upper left abdominal lesion appeared hypointense, while appeared inhomogeneous and hyperintense on T2-weighted images (Figure 2). According to these imaging results, the abdominal lesion was primarily considered to be a teratoma.

After sufficient pre-surgical preparation, an exploratory laparotomy was performed. We found a giant mass originating from the mesentery and surrounded by a fibrous capsule. The territory of tumour blood supply was the branch of superior mesenteric artery. We carefully separated tissues around the tumour, ligated the vessels supplying the tumour and resected the tumour completely from the mesentery. Intraoperative frozen pathology revealed an ossified mesenteric schwannoma.

Macroscopically, the mass was 10.0 cm × 9.0 cm × 9.0 cm in size and yellowish-white in colour. Microscopically, in the mesenteric tumour, both
hypercellular and hypocellular areas were visible. The tumour mainly consisted of spindle-shaped cells with a palisading arrangement; atypical cells or signs of malignancy were not observed (Figure 3A). Some areas of the tumour were ossified, and a true metaplastic bone formation could be seen, with the presence of bone lamellae and osteoblasts (Figure 3B). Immunohistochemical investigation of the tumour showed a strong positivity for S-100 protein (Figure 4), while SMA, CD34, CD117 and DOG-1 were negative. The cell proliferation index, measured with Ki67 staining, was less than 3%. Finally, a giant benign ossified mesenteric schwannoma was diagnosed. After surgery, the patient recovered smoothly and left the hospital 8 d later. She was followed up for a period of 43 mo, during which she was well, with no evidence of tumour recurrence during the follow-up time.

**DISCUSSION**

Schwannoma is an encapsulated tumour arising from Schwann cells[16]. Malignant peripheral sheath nerve tumours (MPNSTs) are uncommon and are always associated with von Recklinghausen’s disease[17]. Schwannomas mostly develop in young to middle-aged patients[18] and typically arise in the head, neck, and extremities[19]. The tumours in the ligament[20], mesentery[10] and intra-abdominal organs[21-23] are extremely rare. Secondary degenerative changes, including cyst formation, calcification, haemorrhage, and hyalinization, can sometimes be shown. However, mesenteric schwannoma with ossified degeneration has not been reported thus far, and to our knowledge, only nine cases of schwannomas in the bowel mesentery have been reported[7-15]. In the present study, we present a giant ossified benign mesenteric schwannoma. Brief clinical characteristics of ten cases of mesenteric schwannoma in the English-language literature, including our case, are outlined in Table 1 and the summarized clinical characteristics are present in Table 2. The mean age of these patients was 52.89 ± 12.89 years (range 38-80 years), and the male-female ratio was 4:5. Most patients were asymptomatic (70.00%). The mean tumour size was
8.86 ± 6.68 cm (range 2-22 cm). All of the tumours were benign.

Obtaining an accurate preoperative diagnosis of a mesenteric schwannoma prior to surgery is nearly impossible. The final diagnosis is based on the histo-pathological examinations of resected specimen[8]. Microscopically, these tumours present two morphologic features: Antoni type A areas and Antoni type B areas[24]. Antoni type A areas consist of closely packed spindle cells, with occasional nuclear palisading. Antoni type B areas are hypocellular and contain more mixoid tissue with high water content[18,25], which can be cystic, haemorrhagic, calcified and even ossified[5,26]. In the present study, some areas of the tumour were ossified, and a true metaplastic bone formation could be seen, with the presence of bone lamellae and osteoblasts, just as reported by Gurzu et al[20]. Schwannomas show strong immunoreactivity for S-100 protein, while CD34, CD117, DOG-1 and SMA are negative[27]. In all these cases, immunohistochemical staining is crucial to confirm the differentiation of the tumour cells. In the present study, the tumour located in the mesentery showed a strong positivity for S-100 protein, while SMA, CD34, CD117 and DOG-1 were negative. The cell proliferation index, measured with Ki67 staining, was less than 3%. Ki67 staining is potentially useful in distinguishing benign peripheral nerve sheath tumours from malignant peripheral nerve sheath tumours, particularly in differentiating problematic cases of cellular schwannoma and malignant peripheral nerve sheath tumours. Ki-67 labelling indices ≥ 20% are highly predictive of malignant peripheral nerve sheath tumours[28]. The main differential diagnosis for schwannomas is gastrointestinal stromal tumours (GISTs). GISTs are composed of spindled mesenchymal cells, which show a diffuse growth pattern and, frequently, positivity for CD34, CD117 and DOG-1[29]. Negative immunohistochemical staining for CD34, CD117 and DOG-1 can exclude the possibility of GIST.

Multiple imaging modalities, including US, CT, MRI, are helpful for establishing a probable diagnosis; however, an accurate preoperative diagnosis of the schwanna is still difficult to obtain[30]. On US, schwannomas are usually well-defined hypodense lesions[12]. Plain CT scan shows a well-defined hypodense mass. Cystic, haemorrhagic, calcified and even ossified degeneration sometimes can be seen. On dynamic CT, Antoni A areas of schwannomas are typically well enhanced. Because of the loose stroma and low cellularity, schwannomas with Antoni B areas generally show low density and can be cystic[5,13]. On T1-weighted images, schwannomas appeared hypointense, while appeared inhomogeneous and hyperintense on T2-weighted images[14,30]. Celiac angiography is important to determine the arteries supplying the tumour[11]. Fine-needle aspiration cytology, accompanied by immunohistochemical staining, may play an important role in defining the appropriate treatment procedure and prognosis[31].

Surgery is curative for mesenteric schwannomas. In the present case, by laparotomy, a giant mass was found that had originated from the mesentery and was surrounded by a fibrous capsule. The patient was followed up for a period of 43 mo, during which she remained well, with no evidence of tumour recurrence.
In conclusion, a schwannoma located in the mesentery is extremely rare. To our knowledge, our study presents the tenth reported mesenteric schwannoma. In addition, a mesenteric schwannoma with ossified degeneration has not been reported thus far. In the present study, we present the first giant benign mesenteric schwannoma with ossified degeneration. It was difficult to acquire an accurate diagnosis of the mesenteric schwannoma preoperatively, because of the lack of specific symptoms, radiological characteristics and tumour markers.

**ARTICLE HIGHLIGHTS**

**Case characteristics**
A 58-year-old woman visited our department due to a lesion detected incidentally in her abdominal cavity during a routine health examination in the local hospital.

**Clinical diagnosis**
A lesion, palpable in her upper left abdomen, was sized 12.0 cm × 6.0 cm, and she felt slight tenderness.

**Differential diagnosis**
Gastrointestinal stromal tumour, intra-abdominal teratoma, sarcoma and neurogenic tumour.

**Laboratory diagnosis**
Blood test findings, including tumour markers, were unremarkable.

**Imaging diagnosis**
Ultrasound revealed a solid lesion in the upper left abdomen, with clear margin, and also showed some cystic and strong echo areas in the lesion. Color doppler flow imaging (CDFIs) showed blood signals in the lesion. In a native computed tomography (CT) scan, the lesion in the upper left abdomen appeared well-defined and was 9.6 cm in diameter, while regions of high density were visible, compatible with calcification and/or ossification. Contrast-enhanced CT study revealed a lesion with slight and inhomogeneous enhancement. On T1-weighted images, the upper left abdominal lesion appeared hypointense, while appeared inhomogeneous and hyperintense on T2-weighted images (Figure 2). According to these imaging results, the abdominal lesion was primarily considered to be a teratoma.

**Pathological diagnosis**
Macroscopically, the mass was 10.0 cm × 9.0 cm × 9.0 cm in size and yellowish-

---

Table 2  Summary of clinical data from all 10 cases of mesenteric schwannomas

| Parameter                          | Value                  |
|------------------------------------|------------------------|
| Age (yr) (n = 8)                   | Mean: 52.89 ± 12.14 (38-80) |
| Sex (male/female), (male %) (n = 9) | 4/5 (44.44%)          |
| Symptoms (n = 10)                  | Asymptomatic: 7 (70.00) |
| Symptomatic                        | Abdominal pain: 3 (30.00) |
|                                  | Hematochezia: 1 (10.00) |
|                                  | Nausea: 1 (10.00)       |
|                                  | Vomiting: 1 (10.00)     |
|                                  | Constipation: 1 (10.00) |
| Mean size (cm) (n = 7)             | 8.86 ± 6.68 (2-22)     |
| Operation (n = 9)                  | Laparotomy: 6 (66.67)   |
|                                  | Laparoscopic operation: 2 (22.22) |
|                                  | Enucleation: 1 (11.11)  |
| Histology (n = 10)                 | Benign: 10 (100.00)    |
|                                  | Malignant: 0 (0.00)     |
| Follow-up (mo) (n = 8)             | 18.43 ± 12.00 (5-43)   |
| Survived                           | 7 (100.00)             |

---

Figure 3  Microscopic examination. A: Microscopically, in the mesenteric tumour, both hypercellular and hypocellular areas were visible. The tumour mainly consisted of spindle-shaped cells with a palisading arrangement; atypical cells or signs of malignancy were not observed (HE × 40); B: Some areas of the tumour were ossified, and a true metaphasic bone formation could be seen, with the presence of bone lamellae and osteoblasts (HE × 200). HE: Hematoxylin and eosin.

Figure 4  Immunohistochemical staining. The mesenteric tumour was strongly positive for S-100 (× 40).
white in colour. Microscopically, in the mesenteric tumour, both hypercellular and hypopcellular areas were visible. The tumour mainly consisted of spindle-shaped cells with a palisading arrangement; alkaptonic cells or signs of malignancy were not observed. Some areas of the tumour were ossified, and a true metaplastic bone formation could be seen, with the presence of bone lamellae and osteoblasts. Immunohistochemical investigation of the tumour showed a strong positivity for S-100 protein, while SMA, CD34, CD117 and DOG-1 were negative. The cell proliferation index, measured with Ki67 staining, was less than 3%. Finally, a giant benign ossified mesenteric schwannoma was diagnosed.

**Treatment**

The authors completely resected the mass located in the mesentery by laparotomy.

**Related reports**

Schwannomas in the bowel mesentery are extremely rare. To our knowledge, only 9 cases of schwannomas located in the bowel mesentery have been reported. In addition, mesenteric schwannoma with ossified degeneration has not been reported thus far.

**Experiences and lessons**

In the present study, the authors present the first giant ossified benign mesenteric schwannoma. It was difficult to obtain an accurate diagnosis of the mesenteric schwannoma preoperatively, because of the lack of specific symptoms, radiological characteristics and tumour markers.

**REFERENCES**

1. Le Guellec S. [Nerve sheath tumours]. Ann Pathol 2015; 35: 54-70 [PMID: 25541115 DOI: 10.1016/j.annpat.2014.11.008]
2. J D, R S, K C, Devi NR. Pancreatic schwannoma - a rare case report. J Clin Diagn Res 2014; 8: FD15-FD16 [PMID: 25177575 DOI: 10.7860/JCDR/2014/8465.4642]
3. Ariel IM. Tumors of the peripheral nervous system. CA Cancer J Clin 1983; 33: 282-299 [PMID: 6413007 DOI: 10.3322/canjclin.33.5.282]
4. Pilavaki M, Choumouzi D, Kiziridou A, Skordalaki A, Zampoukias T, Drevelgas A. Imaging of peripheral nerve sheath tumours with pathologic correlation: pictorial review. Eur J Radiol 2004; 52: 229-239 [PMID: 15544900 DOI: 10.1016/j.ejrad.2003.12.001]
5. Xu SY, Sun K, Xie HY, Zhou L, Zheng SS, Wang WL. Hemorrhagic, calcified, and ossified benign retropitoneal schwannoma: First case report. Medicine (Baltimore) 2016; 95: e4318 [PMID: 27472709 DOI: 10.1097/MD.0000000000004318]
6. Das Gupta TK, Brasfield RD. Tumors of peripheral nerve origin: benign and malignant solitary schwannomas. CA Cancer J Clin 1970; 20: 228-233 [PMID: 4316984 DOI: 10.3322/canclin.20.4.228]
7. Medina-Gallardo A, Curbelo-Peña Y, Molinero-Polo J, Saladich-Cubero M, De Castro-Gutiérrez X, Vallverdú-Cartie H. Mesentric intramural schwannoma: uncommon case of neurogenic benign tumor. J Surg Case Rep 2017; 2017: rjx008 [PMID: 28485819 DOI: 10.1093/jscr/rjx008]
8. Tepox Padrón A, Ramirez Márquez MR, Cordóva Ramón JC, Cosme-Labarthe J, Carrillo Pérez DL. Mesenteric schwannoma: an atypical Schwannoma in the hepatoduodenal ligament: A case report and literature review. World J Gastroenterol 2017; 23: 3744-3751 [PMID: 28611527 DOI: 10.3748/wjg.v24.i20.3744]
9. Ahel B, Hart WR, Olson JR. Tumors of the peripheral nerve system. Hum Pathol 1970; 1: 503-551 [PMID: 4330996]
10. Bayraktutan U, Kantarci M, Ozgokce M, Aydilin B, Atamanalp SS, Sipal S. Education and Imaging. Gastrointestinal: benign cystic schwannoma localized in the gastrooduodenal ligament; a rare case. J Gastroenterol Hepatol 2017; 32: 985 [PMID: 25515808 DOI: 10.1111/j.1440-1746.2012.06960.x]
11. Liu LN, Xu XH, Zheng SG, Sun LP, Guo LH, Wu J. Solitary schwannoma of the gallbladder: a case report and literature review. World J Gastroenterol 2014; 20: 6685-6690 [PMID: 24914396 DOI: 10.3748/wjg.v20.i21.6685]
12. Nishikawa T, Shimura K, Tsuyuguchi T, Kiyono S, Yokosuka O. Contrast-enhanced harmonic EUS of pancreatic schwannoma. Gastrointest Endosc 2016; 83: 463-464 [PMID: 26341855 DOI: 10.1016/j.gie.2015.08.041]
13. Xu SY, Guo H, Shen Y, Sun K, Xie HY, Zhou L, Zheng SS, Wang WL. Multiple schwannomas synchronously occurring in the porta hepatitis, liver, and gallbladder: first case report. Medicine (Baltimore) 2016; 95: e4578 [PMID: 27537565 DOI: 10.1097/MD.0000000000004378]
14. Tao L, Xu S, Ren Z, Lu Y, Kong X, Weng X, Xie Z, Hu Z. Laparoscopic resection of benign schwannoma in the hepatoduodenal ligament: A case report and review of the literature. Oncol Lett 2016; 11: 3349-3353 [PMID: 27123115 DOI: 10.3892/ol.2016.4410]
15. Xu SY, Sun K, Xie HY, Zhou L, Zheng SS, Wang WL. Schwannoma in the hepatoduodenal ligament: A case report and literature review. World Oncol 2016; 11: 308-311 [PMID: 27457049 DOI: 10.1016/j.wol.2016.02.004]
16. Guzru S, Bara T, Bara T Jr, Jung I. Metaplastic bone formation in the abdominal wall–an incidental finding in a patient with gastric cancer. Case report and hypothesis about its histogenesis. Am J Dermatopathol 2013; 35: 844-846 [PMID: 24257191 DOI: 10.1097/DAD.0b013e318285d5bc]
17. Weiss SW, Langloss JM, Enzinger FM. Value of S-100 protein in the diagnosis of soft tissue tumors with particular reference to benign and malignant Schwann cell tumors. Lab Invest 1983; 49: 299-308 [PMID: 631027]
18. Pekmezci M, Reuss DE, Hibe AC, Dahiya GT, Gutmann DH, von Deimling A, Horvai AE, Perry A. Morphologic and immunohistochemical features of malignant peripheral nerve sheath tumors and cellular schwannomas. Mod Pathol 2015; 28: 187-200 [PMID: 25189642 DOI: 10.1038/modpathol.2014.109]
19. Kovacs A, Jung I, Bara T, Bara T Jr, Azamirei L, Kovacs Z, Guzru S. First Case Report of a Sporadic Adrenocortical Carcinoma With Gastric Metastasis and a Synchronous Gastrointestinal Stomal Tumour.
Tumor of the Stomach. *Medicine* (Baltimore) 2015; 94: e1549 [PMID: 26376405 DOI: 10.1097/MD.0000000000001549]

30 Xu SY, Sun K, Owusu-Ansah KG, Xie HY, Zhou L, Zheng SS, Wang WL. Central pancreatectomy for pancreatic schwannoma: A case report and literature review. *World J Gastroenterol* 2016; 22: 8439-8446 [PMID: 27729750 DOI: 10.3748/wjg.v22.i37.8439]

31 Domanski HA, Akerman M, Engellau J, Gustafson P, Mertens F, Rydholm A. Fine-needle aspiration of neurilemoma (schwannoma). A clinicocytopathologic study of 116 patients. *Diagn Cytopathol* 2006; 34: 403-412 [PMID: 16680779 DOI: 10.1002/dc.20449]

**P- Reviewer:** Lee CL, Lee WJ, Maruyama H, Serio G  
**S- Editor:** Gong ZM  
**L- Editor:** A  
**E- Editor:** Ma YJ
