Intensity modulated radiation therapy to treat primary female mediastinal seminoma and massive pericardial effusion: A case report

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Abstract. Primary mediastinal seminoma is a rare extragonadal germ cell tumour that mainly occurs in males. The present study reports the case of a 27-year-old woman that presented with superior vena cava syndrome and a large mass in the mediastinum, which was diagnosed as primary female mediastinal seminoma. The patient received 6 cycles of cisplatin-based chemotherapy (4 cycles BEP chemotherapy (120 mg cisplatin, 0.45 g etoposide and 60 mg bleomycin, once every 3 weeks); 2 cycles IEP chemotherapy (120 mg cisplatin, 100 mg epirubicin and 6 g ifosfamide, once every 3 weeks]) and the patient showed an increase in the size of the mediastinal mass and hydropericardium, indicating a resistance to chemotherapy. Radiotherapy to the mediastinum (50 Gy over 18 fractions) and pericardium (30 Gy over 18 fractions) was performed. Following radiotherapy, the patient was considered to have a complete response to the treatment, and subsequent to a 5-year follow-up, no recurrence or metastasis was identified. To the best of our knowledge, no similar cases have been reported in the literature at present.

Introduction

Extragonadal germ cell tumours (EGCTs) mostly originate from primordial germ cell or thymus cells which may transform to germ cells, accounting for 2-5% of all germ cell tumours (1). EGCTs are further differentiated into seminoma, non-seminoma and mixed-cell tumours. Seminoma is the most common type of EGCT, and all three types are treated using radiotherapy and chemotherapy-based comprehensive treatments. EGCTs mostly present in the medial side of the body, including the mediastinum and retroperitoneum (2). Primary mediastinal seminoma is a rare extragonadal germ cell tumour that mainly occurs in males (3). Primary mediastinal seminoma commonly occurs in the anterior mediastinum of men with an average age of 33 years, and it is rarely observed in women. The incidence rate of primary mediastinal seminoma accounts for 1-4% of mediastinal tumors (4). As primary mediastinal seminoma is a highly radiosensitive tumour, radiotherapy is currently considered to be the most important means of treatment, though chemotherapy may also be used (4).

Case report

In November 2008, a 27-year-old woman presented at The Second Xiangya Hospital of Central South University (Changsha, China) with the signs and symptoms of superior vena cava syndrome (swollen neck), which had been present for 10 days, and was admitted to the present department. The patient had given birth 2 months prior to presentation. CT (Siemens) revealed a 12x6-cm mass in the anterior mediastinum, and water density shadow was observed inside the thickened pericardium (Fig. 1). Examination of the mediastinal mass biopsy (prepared using a BARD biopsy gun; Bard Company, Murray Hill, NJ, USA) in December 2008 revealed distributed macro-nucleic cells under microscope (N-117M CKX41; Olympus Corporation, Tokyo, Japan), with the following immunohistochemical findings: Strong expression of placental alkaline phosphatase (PLAP); expression of cluster of differentiation (CD)3 (dilution, 1:100; GD-x0297M-AF03; Abcam, Cambridge, UK), CD45 (dilution, 1:100; GD-x0297M-AF20; Abcam) and CD20 (dilution, 1:100; GD-x0297M-AF20; Abcam); and no expression of CD79a (dilution, 1:100; GD-x0297M-AF79a), cytokeratin (dilution, 1:500; GD-cyt-001; Abcam), S100 (dilution, 1:100; GD-S-100; Abcam), terminal deoxynucleotidyl transferase (dilution, 1:200; GD-te-001; Abcam), CD5 (dilution, 1:100; GD-x0297M-AF05; Abcam), CD30 (dilution, 1:100; GD-x0297M-FAF30; Abcam), CD15 (dilution, 1:100; GD-x0297M-AF15; Abcam), anaplastic lymphoma kinase (dilution, 1:500; GD-ana-001; Abcam), B cell lymphoma-2 (dilution, 1:100; GD-bcl-005; Abcam) and CD10 (dilution, 1:100; GD-x097M-AF10; Abcam) (Fig. 2).
These findings were consistent with the diagnosis of primary mediastinal seminoma and stage IV disease. The patient was administered with 2 cycles of the bleomycin, etoposide and cisplatin (BEP) chemotherapy regimen (120 mg cisplatin, 0.45 g etoposide and 60 mg bleomycin, once every 3 weeks). CT re-examination showed a distinctly reduced mediastinum...
mass size, decreased pericardial effusion and disappearance of the right mediastinal exudates (Fig. 3). The BEP regimen was prolonged with the same dosages for an additional 2 cycles. CT re-examination showed mild shrinkage of the mediastinal mass, but pericardial effusion had increased (Fig. 4). Thus, the chemotherapy regimen was revised to the ifosphamide, epirubicin and cisplatin regimen. CT re-examination (Fig. 5) shows that the mediastinal mass had decreased in size again, with reduced pericardial effusion, considering the treatment efficacy the regimen was prolonged. Another CT re-examination was performed 3 weeks subsequently, which showed the advancement of disease with an enlarged mediastinal mass and increased volume of pericardial effusion (Fig. 6). The patient was administered with intensity-modulated radiotherapy therapy from June 2008. A dose of 50 Gy/18 fractions was administered to the upper mediastinum and a dose of 30 Gy/18 fractions was administered to the pericardial area. Pericardiocentesis was performed post-radiotherapy, which drained 700 ml of red fluid, together with the infusion of 1 g fluorouracil. Post-radiotherapy CT examination showed that the mediastinal mass had disappeared, with a mild amount of pericardial effusion (Fig. 7). For 5 years subsequent to treatment, to the present time, there has been no relapse of the tumour and the patient is leading a normal life.

Discussion

In 2004, the World Health Organization defined mediastinal seminoma as a type of extragonadal low-grade malignant germ cell tumour that commonly occurs in the anterior mediastinum of men, with the average age of 33 years, and it is rarely observed in women (4). The clinical data show that primary mediastinal seminoma accounts for 0.5-5% of all mediastinal tumours (5).

Clinical records show that the most common clinical symptoms of primary mediastinal seminoma include chest
pain, dyspnoea and hoarseness of voice (1). In the majority of cases, CT shows heterogeneously distributed density of the mediastinal mass (6). Monitoring of serum tumour markers is significantly important for the diagnosis and prognostic evaluation of patients with mediastinal seminoma. According to the 2010 American Society of Clinical Oncology clinical guidelines, increased levels of biomarkers such as HCG, LDH and AFP should determine the presence of relapse or metastasis of the disease (7). In addition, 80-90% of mediastinal seminoma cases are found to be PLAP-positive by immunohistochemistry (2). The diagnosis of primary mediastinal seminoma in the present patient was confirmed by the clinical presentation, CT and the morphological appearance of the tumour. In addition, immunohistochemical analysis was performed for PLAP, CD3, CD20 and CD45, all of which were found to be expressed.

Among mediastinal tumours, seminoma is a highly radiosensitive tumour, and therefore long-term radiotherapy is considered the most important means of treatment and increases the long-term survival rate of patients by 60-80% (8). Patients administered with platinum-based chemotherapy had an overall long-term survival rate of 90%. At present, the most commonly accepted treatment methods for mediastinal seminoma may be classified as two types. Radiotherapy is used if patients develop post-chemotherapeutic local recurrence and remnant lesions and surgery is used in patients with post-chemotherapeutic remnant lesions >3 cm in size.

In the present case, there was recurrent relapse of the disease and the case was unique as the patient was female. The chemotherapeutic effect varies between men with primary mediastinal seminoma (9). Another explanation is that the patient was diagnosed with mediastinal seminoma through biopsy. However, the pathological biopsy specimen was small, and the tumour may have contained non-seminomatous germ cell tumour type cells, which were not biopsied. In addition, mixed-type gonadal cell tumours account for 13-34% of mediastinal tumours and they are less sensitive to chemotherapy compared with cases of seminoma (2). The present patient had large amounts of pericardial effusion, which is a contraindication for radiotherapy. However, considering that the patient was resistant to chemotherapy and the challenging nature of surgery due to extensive invasion, with no significant cardiac compression symptoms, such as dyspnoea or shock, the patient was advised to undergo radiotherapy. Subsequent to the aforementioned treatments the patient did not experience any relapse or metastasis during the 5 years of follow-up.

Thus, the present study shows that various comprehensive treatment methods should be used for the effective treatment of mediastinal seminoma. Although radiotherapy is contraindicated in pericardial effusion, the present study indicates that it could be considered if there are no other means of treatment and with no evident signs of cardiac compression.

References