CASE REPORT

THORACIC RADIOTHERAPY FOR MEDIASTINAL NODAL RECURRENCE

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Summary: Radiotherapy has been used to treat loco-regional recurrences located at various intra-thoracic sites, but long-term survival of these patients has been rarely observed. We report herein a lung adenocarcinoma patient with loco-regional recurrence, who was successfully treated with high-dose radiotherapy. The patient could survive with no evidence of recurrence 5 years after thoracic irradiation. It is probably safe to administrate high-dose radiotherapy for some loco-regional recurrent patients with favorable prognostic factors such as good PS, no body weight loss. Further studies will be required to define a favorable subset of patients most likely to benefit from an aggressive approach.

Key words: Non-small cell lung cancer; Loco-regional recurrence; Radiotherapy

Introduction

Although surgery is the standard treatment for non-small cell lung cancer (NSCLC), unfortunately loco-regional recurrence of NSCLC occurs in 13 % to 50 % of patients (3, 4, 6, 7, 8, 10, 14). Loco-regionally recurrent NSCLC has been treated with radiotherapy, with reported median overall survival of nearly one year (3, 4, 6, 7, 8, 10, 14). We herein describe a successful treatment for NSCLC with postsurgical mediastinal lymph node recurrence by thoracic irradiation.

Case report

A 71-year-old man complicated of cough for over two months. Chest radiograph and CT scan revealed a tumor in the lower lobe of the left lung. Hilar and mediastinal lymph node adenopathy was not observed. He was referred to our hospital for further examination and treatment. Physical examination was unremarkable and his performance status (PS) was evaluated as one. Laboratory data including tumor markers were normal. As the tumor had heterogeneous density with irregular-shaped margin on CT scan, lung adenocarcinoma was suspected. But a definitive diagnosis could not be made by transbronchial biopsy. Although his minimal impairment of his lung function, video-assisted thoracic surgery was selected intra-operative diagnosis was adenocarcinoma. Subsequent left lower lobectomy with nodal dissection was performed based on the intra-operative histologic diagnosis. The final pathological diagnosis was adenocarcinoma, measuring 13 x 12 x 8 mm, and metastatic ipsilateral hilar node was histologically identified. Pathological tumor stage was proved to be T1N1M0. The patient had an uneventful postoperative course and was discharged. He was followed up at our outpatient department without any additional adjuvant therapy. Although he had no signs and symptoms, follow up chest CT scan, which was performed without contrast enhancement of his allergic reaction at the first time of chest CT scan, revealed left pre-tracheal lymph node recurrence developed six months after the surgery (Fig. 1). The patient had no body weight loss and his PS was zero at this time. Brain MRI, bone scan and CT of the abdomen showed no other distant metastasis. The patient was recommended additional therapy such as surgery, chemotherapy, or chemo-radiotherapy, but he wanted to receive radiotherapy. Clinical target volume (CTV) was included both supraclavicular fossa, both hilum region and subcarinal region to 5 cm below carina. Planning target volume (PTV) added 1 cm to CTV all around directions. PTV was irradiated using anterioposterior and posterioranterior port dose of 46 Gy. After then PTV was shrieked to pre-tracheal node area using oblique port. The total dose of radiotherapy was delivered 70 Gy using 10 MV photon. Disappearance of the recurrent mediastinal node was confirmed on subsequent CT scan, and the patient was followed up again at our outpatient department. He is still alive and well with no evidence of recurrence 5 years after thoracic irradiation (Fig. 2).

Discussion

Surgical resection is the treatment of choice for early stages of NSCLC, including sometimes patients with selected stage IIIA disease. Analysis of the patterns of failure

shows that loco-regional recurrence of NSCLC after surgical resection occurs in 13% to 50% of patients (3, 4, 6, 7, 8, 10, 14). Loco-regional recurrences usually divide into chest wall/pleural, parenchymal, bronchial stump, and mediastinal lymph node recurrences. Several previous authors evaluated favorable prognostic factors in loco-regional recurrent NSCLC patients after curative surgery (4, 7, 14). Emami et al reported that the best indicators for long-term survival were the interval from initial surgery to first recurrence and tumor response to radiation therapy (4). Jeremic et al evaluated favorable prognostic factors in NSCLC patients with loco-regional recurrence after curative surgical resection (7). In their univariate analysis, there was a significant difference in survival between high-dose and lowdose radiotherapy groups. They showed that age, extent of initial surgery, time from initial surgery to documented recurrence were not found to influence survival in the highdose radiotherapy group, but good PS, female gender, and no body weight loss were also favorable prognostic factors in patients treated with high-dose radiotherapy (7). In their patients, initial staging significantly influenced survival and patients with initial stage I did significantly survive longer than those with either initial stage II or initial stage IIIA. In addition, recurrent staging also influenced survival and in a similar manner as initial staging did (7). On the contrary, Shaw et al found no influence of either initial or recurrent staging on treatment outcome (14). In our patient, initial staging was stage IIA and recurrent stage was evaluated as stage IIIA, but the recurrence was observed only in ipsilateral pre-tracheal lymph node. He had good PS and no signs and symptoms, and no body weight loss at the time of initial diagnosis as well as recurrence. Early detection of single station mediastinal lymph node metastasis might have certain clinical significance.

Recently, Kelsey et al described patterns of failure after resection of NSCLC and implications for postoperative radiation therapy volumes (9). Even if postoperative thoracic radiotherapy has been widely used as adjuvant treatment, the clinical benefit in this treatment after surgical resection in NSCLC has poorly evaluated (1). Moreover, few studies have reported technical aspects of irradiation for locoregional recurrence of postoperative NSCLC. As above mentioned, clinical benefit of high-dose irradiation, 60Gy (8) or more (11), has been evaluated. Some authors have indicated that the omission of elective nodal treatment has no relationship with significant amount of failure in lymph node regions (2, 5, 11, 12, 13, 15). Tada et al reported that the prescribed dose was 60 Gy in 30 fractions over 6 weeks (16). The radiation field contained the ipsilateral hilar lymph nodes and the mediastinal lymph nodes, respectively, and the radiation field contained the recurrent tumor and margins of more than 20 mm (16). Kagami described that radiotherapy for loco-regional recurrences of NSCLC after complete surgery was an effective treatment modality in terms of improved survival and palliation (8). A randomized study from Italy demonstrated that carefully delivered postoperative radiation therapy by use of three-dimensional treatment planning improved local control and overall survival (17). In this study, the average field size was 6.5 X 7 cm. In our patient, CTV was included both supraclavicular fossa, both hilum region and subcarinal region to 5 cm below carina. PTV added 1 cm to CTV all around directions. Although our patient recurred 6 months after the surgical resection, the patient had above-mentioned several favorable factors as previously indicated (4, 7, 14). Therefore, the patient could survive with no evidence of recurrence 5 years after thoracic irradiation.

In summary we showed the effectiveness of curative, high-dose radiotherapy in the treatment of loco-regional recurrent NSCLC after curative resection. It is probably safe to administrate high-dose radiotherapy for some loco-regional recurrent patients with favorable prognostic factors such as good PS, no body weight loss. It warrants further studies that may help define a favorable subset of patients most likely to benefit from an aggressive approach.



Fig. 1: Follow up chest CT scan revealed left pre-tracheal lymph node recurrence due to tumor cell implantation developed 6 months after the surgery.



Fig. 2: A chest CT scan 5 years after the thoracic irradiation, which shows no evidence of local recurrence.

References

- Arriagada R, Le Péchoux C, Pignon JP. Resected non-small cell lung cancer: need for adjuvant lymph node treatment? From hope to reality. Lung Cancer 2003;42 Suppl 1:S57-64.
- Bradley J, Graham MV, Winter K, et al. Toxicity and outcome results of RTOG 9311: a phase I-II dose-escalation study using three-dimensional conformal radiotherapy in patients with inoperable non-small-cell lung carcinoma. Int J Radiat Oncol Biol Phys 2005;61:318-28.
- Curran WJ, Herbert SH, Stafford PM, et al. Should patients with post-resection locoregional recurrence of lung cancer receive aggressive therapy? Int J Radiat Oncol Biol Phys 1992:24:25-30.
- Emami B, Graham MV, Deedy M, Shapiro S, Kucik N. Radiation therapy for intrathoracic recurrence of non-small cell lung cancer. Am J Clin Oncol 1997; 20:46-50.
- Emami B, Mirkovic N, Scott C, et al. The impact of regional nodal radiotherapy (dose/volume) on regional progression and survival in unresectable non-small cell lung cancer: an analysis of RTOG data. Lung Cancer 2003;41:207-14.
- Green N, Kern W. The clinical course and treatment results of patients with postresection locally recurrent lung cancer. Cancer 1978;42:2478-82.
- Jeremic B, Shibamoto Y, Milicic B, et al. External beam radiation therapy alone for loco-regional recurrence of non-small-cell lung cancer after complete resection. Lung Cancer 1999;23:135-42.
- Kagami Y, Nishio M, Narimatsu N, et al. Radiotherapy for locoregional recurrent tumors after resection of non-small cell lung cancer. Lung Cancer 1998;20:31-5.

- Kelsey CR, Light KL, Marks LB. Patterns of failure after resection of non-smallcell lung cancer: implications for postoperative radiation therapy volumes. Int J Radiat Oncol Biol Phys 2006;65:1097-105.
- Kopelson G, Choi NC. Radiation therapy for postoperative local-regionally recurrent lung cancer. Int J Radiat Oncol Biol Phys 1980;6:1503-6.
- Martel MK, Ten Haken RK, Hazuka MB, et al. Estimation of tumor control probability model parameters from 3-D dose distributions of non-small cell lung cancer patients. Lung Cancer 1999;24:31-7.
- Rosenzweig KE, Sim SE, Mychalczak B, et al. Elective nodal irradiation in the treatment of non-small-cell lung cancer with three-dimensional conformal radiation therapy. Int J Radiat Oncol Biol Phys 2001;50:681-5.
- Senan S, van Sornsen de Koste J, Samson M, et al. Evaluation of a target contouring protocol for 3D conformal radiotherapy in non-small cell lung cancer. Radiother Oncol 1999;53:247-55.
- Shaw EG, Brindle JS, Creagan ET, et al. Locally recurrent non-small-cell lung cancer after complete surgical resection. Mayo Clin Proc 1992;67:1129-33.
- 15. Sim S, Rosenzweig KE, Schindelheim R, et al. Induction chemotherapy plus three-dimensional conformal radiation therapy in the definitive treatment of locally advanced non-small-cell lung cancer. Int J Radiat Oncol Biol Phys 2001; 51:660-5.
- Tada T, Fukuda H, Nakagawa K, et al. Non-small cell lung cancer: radiation therapy for locoregional recurrence after complete resection. Int J Clin Oncol 2005;10:425-8.
- Trodella L, Granone P, Valente S, et al. Adjuvant radiotherapy in non-small cell lung cancer with pathological stage I: definitive results of a phase III randomized trial. Radiother Oncol 2002;62:11-9.

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