Partial frequency of radiation pneumonitis and its association with the energy and treatment technique in patients with breast cancer, Isfahan, Iran

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Background: Breast cancer is the most common type of cancer among women, and the second cause of cancer mortality after pulmonary cancer in this gender. Radiotherapy is one of the major treatments, which locally controls the disease and prohibits in recurrence. Radiation pneumonitis is one of the radiotherapy complications, which usually occurs within 1.5-3 months after radiotherapy. As there is no precise estimation concerning this complication in Isfahan, partial frequency of radiation pneumonitis and its association with the energy and treatment technique in patients with breast cancer were evaluated. Materials and Methods: This was an analytic cross-sectional study performed in 2010 in university referral center. A total of 382 patients with breast cancer, undergone surgery and referred for radiotherapy entered the study. A posterior and anterior and lateral X-Rays were taken as control images before starting radiotherapy and all X-Rays were repeated after 3-4 months post radiotherapy. The occurrence of radiotherapy pneumonitis was evaluated by the same radiologist. Data were analyzed through SPSS version 20. Results: Out of 382 patients undergone breast conservative surgery (BCS) or modified radical mastectomy (MRM), and radiotherapy, 60 patients had pneumonitis of whom 6 patients underwent BCS and were treated by tangential field (three cases by Co 60 and three cases by PH 9). The rest of radiotherapy pneumonitis patients (n = 54) underwent MRM of whom, 42 cases were treated by one-field and 12 by two-field treatment techniques. Conclusion: Incidence of radiotherapy pneumonitis was different with respect to the adopted technique (one-field, two-field and tangential) (P = 0.023), with the highest association with two-field radiotherapy.

Key words: Breast cancer, breast conservative surgery, chest X-ray, energy field, modified radical mastectomy, radiotherapy pneumonitis

INTRODUCTION

Breast cancer is the most common type of cancer among women,[1,7,33] and the second cause of cancer mortality after pulmonary cancer in this gender.[2,33] The age of its occurrence has diminished in the world and Iran.

Now, there are numerous treatments for this type of cancer, which have notably decreased its mortality. Post-surgery radiotherapy is the main part of treatment to locally control the disease and prohibit its recurrence.[3,7,6,21,26,32] When a treated patient’s life gets longer, the treatment complications are more likely to occur. Complications resulted from radiotherapy are not an exception and should be noticed besides numerous treatment benefits of radiation.[13,23,24,29,30,32] One of these complications is radiotherapy pneumonitis usually occurring 1.5-3 months post radiotherapy and is characterized by some clinical signs and radiographic images.[4,14,15,19,2-4] Its incidence has been reported 1.2-35% in North America and west Europe.[5,6,27,29,31]

Breast cancer is the most common cancer in female in Isfahan, Iran.[30] Although there is some reports regarding its frequency, no studies has been carried out concerning frequency of pneumonitis resulted from radiotherapy in Isfahan. Seyed Alshohada Treatment Center is one of the major radiotherapy centers in Iran. Therefore, this study was conducted to investigate partial frequency of radiation pneumonitis and its association with the energy and treatment technique in patients with breast cancer in Seyed Alshohada Hospital in Isfahan during September 2010 to September 2011.

MATERIALS AND METHODS

This was an analytic cross-sectional study. Data were collected during September 2010 to September 2011 in Isfahan Seyed Alshohada Education and Treatment Center. According to our statistics specialists, Chi-square test was adopted to analyze the data.

A total of 382 female patients with breast cancer who had undergone surgery modified radical mastectomy or breast conservative surgery (MRM or BCS) and had been referred for radiotherapy were recruited. The only diagnostic methods for radiotherapy pneumonitis were
supposed to be patient’s clinical condition and Possterior Anterior and Laterl chest X-Rays, which were requested once as a baseline before starting radiotherapy and once as control 3-4 months after the last course of radiotherapy had finished. These baseline and control X-rays were investigated by a fixed radiologist in Seyed Alshohada Hospital for each patient and the occurrence of radiotherapy pneumonitis were evaluated according to a check-list of radiotherapy pneumonitis signs consists of ground glass pattern, opacification, diffuse haziness, and patchy consolidation. The radiologist just knew about the type of surgery, but not the treatment technique and the used energy for radiotherapy. Then, the needed data including patients’ age, the type of surgery (MRM and BCS), type of energy^{[10-11]} (Electron, Cobalt 60, Photon), treatment technique (one-field, two-field, tangential), sub-groups of energy^{[10,12]} (E6, E8, E9, E10, E13, Co 60, PH6, PH9, PH12) and occurrence or not occurrence of radiotherapy pneumonitis were collected. Descriptive results were presented in tables and diagrams.

The patients with a history of taking bleomycin, dactinomycin, and nitrous compounds of urea or with a history of chest X-ray due to any reason were excluded from the study. The existence of a background pulmonary disease before starting radiotherapy and smoking cigarettes or water pipe for a long time were other exclusion criteria.

The SPSS 20 was used for statistical analysis.

RESULTS

A total of 382 patients entered the study of whom three were male and 382 female patients with breast cancer undergoing surgery. Mean (±SD) age of the subjects was 48.5 ± 10.6 years. The youngest subject was 22 and the oldest was 80 years old. Out of 382 patients, 332 and 50 subjects underwent MRM and BCS respectively. From a total of 382 patients, undergone radiotherapy, 60 developed pneumonitis as a result of radiotherapy with the frequency distribution of 15.7% [Table 1].

From 60 cases of radiotherapy pneumonitis, 6 subjects underwent BCS and were treated by tangential field (three cases by Co 60 and three cases by PH9).

The rest of radiotherapy pneumonitis patients (n = 54) underwent MRM of whom 42 were treated by one-field and 12 by two-field techniques respectively. The association between two types of conducted surgeries (BCS or MRM) was not significant (P = 0.454).

The association between type of energy (electron, Co, PH) and radiotherapy pneumonitis was not significant (P = 0.268). The association between treatment technique (one-field, two-field, tangential) and radiotherapy pneumonitis was significant (P = 0.023). [Table 2 and 3].

This association was more notable by two-field technique, than one-field and finally tangential technique respectively. Although the association between sub-groups of energy and pneumonitis was not significant, it was significant for the sub-groups of one-field technique (E6, E13, E8, E9, E10) (P = 0.876). The association was not significant for sub-groups of two-field technique (E8, E10, E13+ E10, E10+ E9, E9+ E13, E9+ E6, E9+ PH9, E13+ PH12, PH6, PH9, E6+ PH9) (P = 0.665).

DISCUSSION

With regard to the obtained results, it was concluded that partial frequency of radiotherapy pneumonitis was 15.6% during September 2010 to September 2011 in the present study, which is laid in the observed statistics in other countries (1.2-35%)^{[16,19]} and even is lower than their mean.

Among all studied factors, there was just a significant association between treatment technique (anterior one-field, anterior two-field, tangential) and radiotherapy pneumonitis.

The patients with a history of taking bleomycin, dactinomycin, and nitrous compounds of urea or with a history of chest X-ray due to any reason were excluded from the study. The existence of a background pulmonary disease before starting radiotherapy and smoking cigarettes or water pipe for a long time were other exclusion criteria. The SPSS 20 was used for statistical analysis.

### Table 1: Frequency distribution of radiotherapy pneumonitis

<table>
<thead>
<tr>
<th>Radiotherapy pneumonitis</th>
<th>Number</th>
<th>Percent</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Positive</td>
<td>60</td>
<td>15.7</td>
<td>0.268</td>
</tr>
<tr>
<td>Negative</td>
<td>322</td>
<td>84.3</td>
<td></td>
</tr>
<tr>
<td>Total</td>
<td>382</td>
<td>100</td>
<td></td>
</tr>
</tbody>
</table>

### Table 2: Association between type of energy and radiotherapy pneumonitis

<table>
<thead>
<tr>
<th>Type of energy</th>
<th>Positive</th>
<th>Negative</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Electron</td>
<td>54</td>
<td>16.9</td>
<td>264</td>
</tr>
<tr>
<td>Cobalt</td>
<td>3</td>
<td>12</td>
<td>0.665</td>
</tr>
<tr>
<td>Photon</td>
<td>3</td>
<td>12</td>
<td>22</td>
</tr>
<tr>
<td>Total</td>
<td>60</td>
<td>15.7</td>
<td>322</td>
</tr>
</tbody>
</table>

P=0.268

### Table 3: Association between the type of field and radiotherapy pneumonitis complication

<table>
<thead>
<tr>
<th>Type of field</th>
<th>Positive</th>
<th>Negative</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>One-field</td>
<td>42</td>
<td>15.2</td>
<td>234</td>
</tr>
<tr>
<td>Two-field</td>
<td>12</td>
<td>28.6</td>
<td>29</td>
</tr>
<tr>
<td>Tangential</td>
<td>6</td>
<td>9.1</td>
<td>59</td>
</tr>
<tr>
<td>Total</td>
<td>60</td>
<td>15.6</td>
<td>322</td>
</tr>
</tbody>
</table>

P=0.023
There was no significant association between surgical techniques, types of energy, and the type of energy sub-groups. Therefore, selection of the an appropriate technique seems to be the most important factor, which can lead to positively or negatively affect radiotherapy pneumonitis incidence in addition to other important factors such as age, pulmonary, cardiac diseases, smoking and etc… influencing its incidence.\textsuperscript{[15,17]} Some researchers have reported a higher incidence of pneumonitis as a result of concurrent radiotherapy and chemotherapy\textsuperscript{[10,21,28,31]} so that this factor has been reported as the most effective element in the increase of pneumonitis incidence.

**CONCLUSION**

No significant association was observed between radiotherapy pneumonitis and the type of surgery and used energy, but there was a significant association between the applied technique (one-field, two-field, tangential) and radiotherapy pneumonitis ($P = 0.023$). As the number of subjects was low in some two-field sub-group, the precise association between the type of two-field energy and radiotherapy pneumonitis could not be detected. An independent study should be conducted on two-field cases and the energy used on them together with adequate sample size to investigate the association between two-field type of energy and radiotherapy pneumonitis.

**REFERENCES**

