

Frequency of histological subtypes and patients' survival with tumors originating from thymus

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Abstract

Background: Thymoma is a rare tumor in the mediastinum. Due to the fact that a complete study has not been conducted on thymoma in Khuzestan region, this study was designed as part of a comprehensive study on thymoma in Ahvaz Golestan Hospital.

Materials and Methods: This retrospective study was conducted on patients' medical records with Thymoma attending Department of Radiotherapy and Oncology, Ahvaz Golestan Hospital, in a 10-year period from 2001 to 2011. The study also aimed to investigate the epidemiological data; diagnosis and treatment of patients for the duration of the same time.

Results: Among the 26 patients with a mean age of 45.38±SE, ratio of male to female was 5.5 to 1. The majority of the patients were in the range of 30-50 years old which jointly was at 61.8±SE. However, the minimum rate of tumor was seen in decades of 51-60 and 71-80 (7.7 %). In total, 72.7% (n=19) patients had invasive thymoma, while 27.3% (n=8) had noninvasive tumor. In addition, there was a significant relationship between invasive thymoma rate and developing myasthenia gravis. The 3-year survival rate for all patients was equivalent to 83.3%. However; 5-year survival was equal to 71%.

Conclusion: According to the study, most patients with invasive thymoma had a 5-year survival rate and also most of them were with myasthenia gravis.

Keywords: Thymoma, Khuzestan, Metastase, Myasthenia gravis, Invasive thymoma

Background

Thymoma is one of the rarest tumors in the mediastinum. The thymus is an epithelial lymph organ, shaped in trapezoidal form located anterior mediastinum in thoracic cavity which in adult replaced by adipose tissue. Thymic neoplasms include neurogenic tumors, germ cell lymphoma and mesenchymal. However, Thymoma and thymic carcinoma and thymolipomas, are originating from thymus components (1). Thymoma and thymic carcinomas are rare tumors and include totally 1.5-2 % of all malignant diseases and thymic carcinoma includes only 0.06% of total neoplasms in body (1). Thymoma is a tumor that originates from the thymus which is invasive locally and in 1/2 cases is noninvasive form. In 1/2 patients Thymoma is asymptomatic and by randomize is found in chest x-ray (2). Myasthenia gravis is an autoimmune disease that is a neuromuscular transport junction disorder due to AB against acetylcholine receptors in muscles (3). Approximately 10-67% of 100 patients with thymoma have myasthenia gravis; however, 10-15% of patients with myasthenia gravis have thymoma even which in cases with normal thymus, thymectomy improves symptoms (4). Thymoma is usually indolent but if it spreads, it is implanted in the pleural space which causes pleural plaques, diaphragmatic tumor and malignant pleural effusion (1). Thymoma is very sensitive to chemotherapy and clinical response was seen in

2/3 of the patients. Best used is as neoadjuvant that prepare the patient for complete resection. In addition, the use of osterotide and high-dose corti- costeroids is effective. Currently, the best treatment is MULTI MODAL that means chemotherapy before surgery and then radiation after surgery (5). In a study conducted by Eric A. Engels et al., of 849 patients, 455 men were and 394 were women (6, 7). In the study conducted by Andre J. et al., the 5-year survival rate was 94%, while 10-year survival was 90 % (8). As well as, in the study conducted by Kristen Bass Wilkins et al., the 5-year survival rate was equivalent to 87%, 10-year survival rate equivalent to 84% and 20-year survival rate was equivalent to 80% (9). In Stefano Margaritora et al.'s study 5, 10, 20, 30 and 35 year survival rates were equivalent to 89.9%, 84.1%, 73%, 58.6% and 58.6%, respectively (10). In another study conducted by Kazou Nakagawa et al., 5-year survival was 92% and 10-year survival was equivalent to 91% (11). In a further study conducted by BS Singhal et al., out of a total of 836 patients, 611 were men and 225 women. As well as, in this study, the age decade of patients with myasthenia gravis was documented for 60-70 years (12).

Materials and Methods

A retrospective, cross-sectional study was designed to investigate the patients with thymoma tumor attending the Department of Radiotherapy

and Oncology, Ahvaz Golestan Hospital for the duration of 2001-2011 years. For this study, all mediastinal tumor cases were considered and among them there were 26 patients with thymoma, whose data were extracted and studied. Based on age, sex, metastasis, invasive or non-invasive, Myasthenia Gravis and last visit a checklist was provided for each patient. The details of each patient were recorded on checklists after thorough examination of each case. As well as obtaining information on the latest status of the patient (the patient's survival) a phone call was given to the patients' family. To investigate and describe the desired variables, the obtained data were analyzed using the methods of descriptive statistics including frequency tables, graphs, and numerical parameters. Subsequently, the chi-square test was used to evaluate the relationship between variables. SPSS was used for data analysis and level of statistical significance was considered as 0.05 for the above tests.

Results

Out of a total of 26 patients, 22 (84.6%) were males and 4 (15.4%) women (Figure 1). There was a maximum age of 76 and a minimum age of 21 years with a mean age of 45.38. The highest rate of tumor was indicated in 31-40 age and 41-50 years (30.8%). However, the lowest tumor rate was docu-

mented for the age decades of 71-80 years old with 7.7% (Figure 2). Out of 22 male patients, 16 (72.7%) had invasive thymoma and six cases (27.3%) had noninvasive thymoma. One out of four women (25%) had invasive thymoma and three patients (75%) had noninvasive thymoma (Figure 5). The majority rate of thymoma was observed in the range of 31-40 and 41-50 years old which was allocated a total number of 16 people. With 61.8% (Figure 2) for 26 patients, seven cases (26.9%) had a myasthenia gravis that of this number, five cases (71.4%) had invasive thymoma and two people (28.6%) were noninvasive thymoma (Figure 3). Out of a total of seven patients with myasthenia gravis, five had invasive thymoma. In total, of the 26 patients, 21 cases completed the study and stood alive; however, five patients had died due to invasive thymoma.

The rate of 3-year survival for all patients was equal to 87% and 95% confidence interval was equal to 1 and 0.695. Furthermore, 5-year survival rate for all patients was 71% and the 95% confidence interval was equivalent to 0.95 and 0.47 (Figure 4). However, 3-year survival rate for invasive tumors was achieved by 83.3% with confidence interval (1 and 0.62). Nonetheless, for 5-year survival in the same tumor this rate was equivalent to a 70% with confidence interval (0.995 and 0.505) (Figure 4).

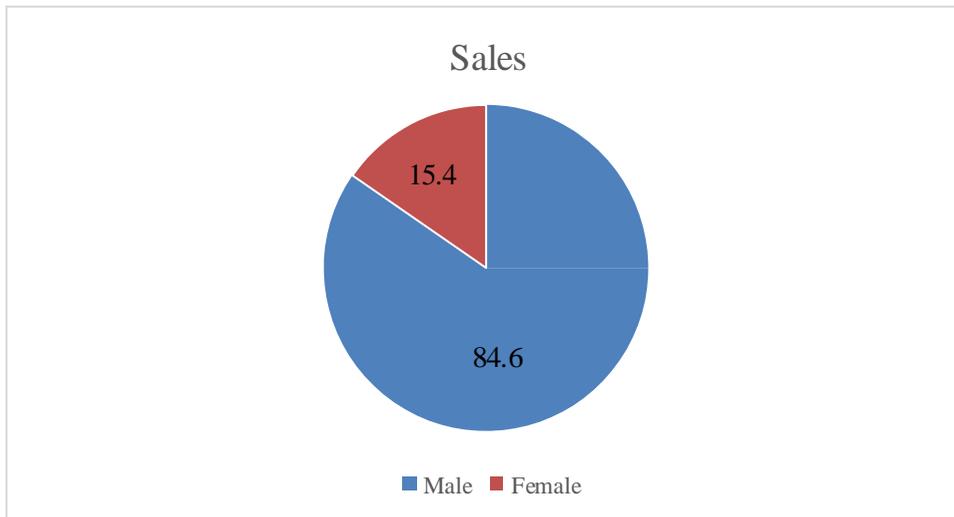


Figure 1: Sex distribution

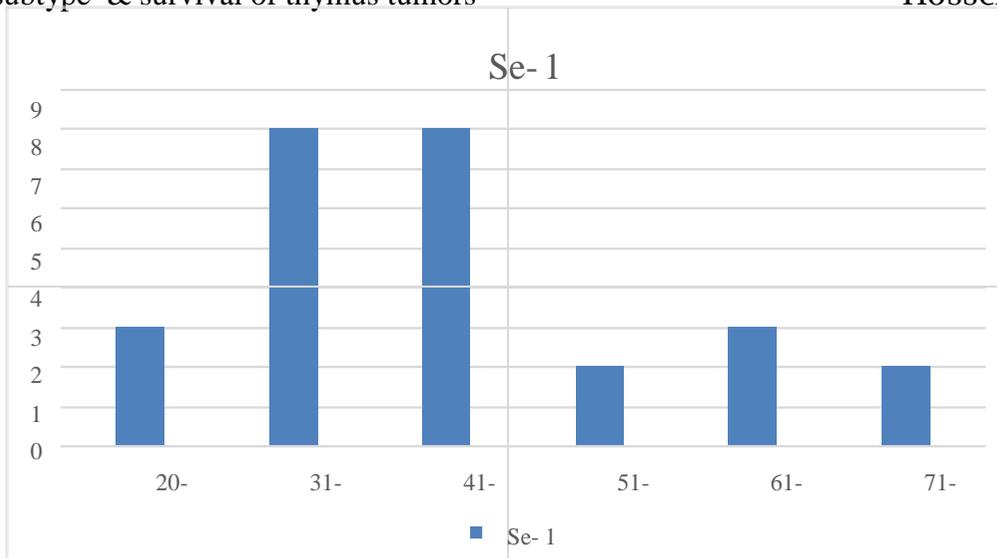


Figure 2: Frequency of patients in each age interval

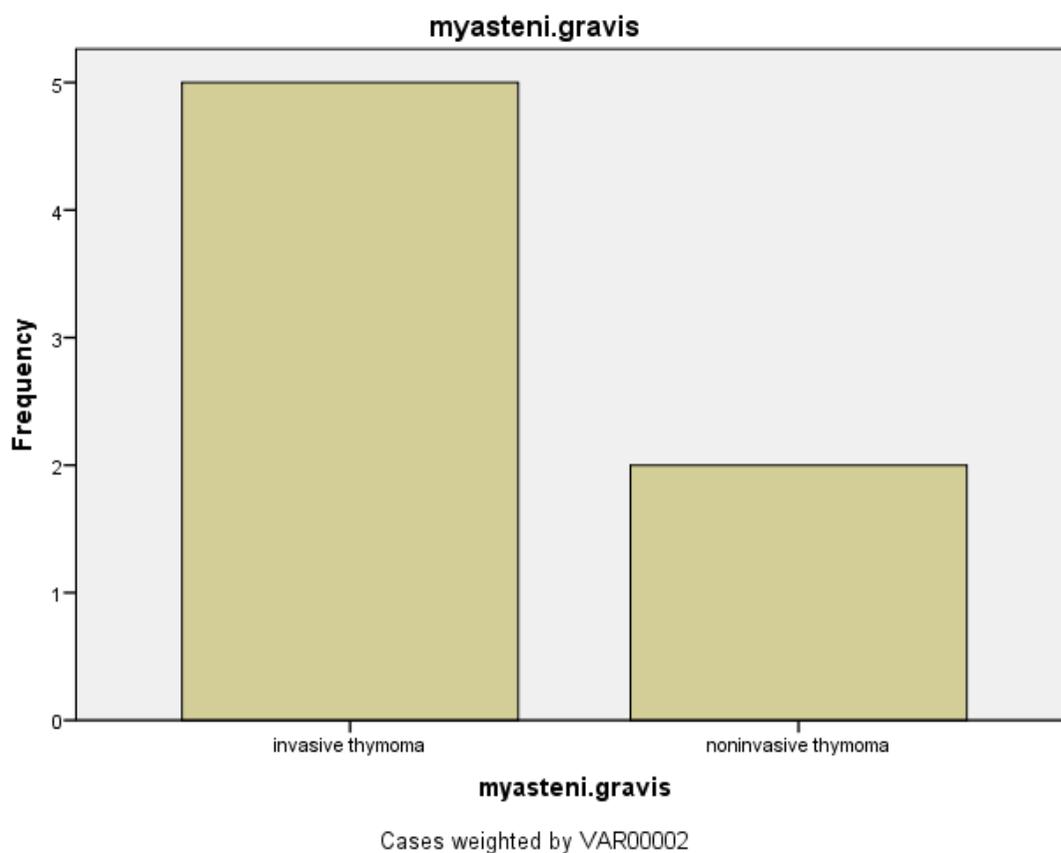


Figure 3: Incidence of invasive and non-invasive tumor in patients with myasthenia gravis

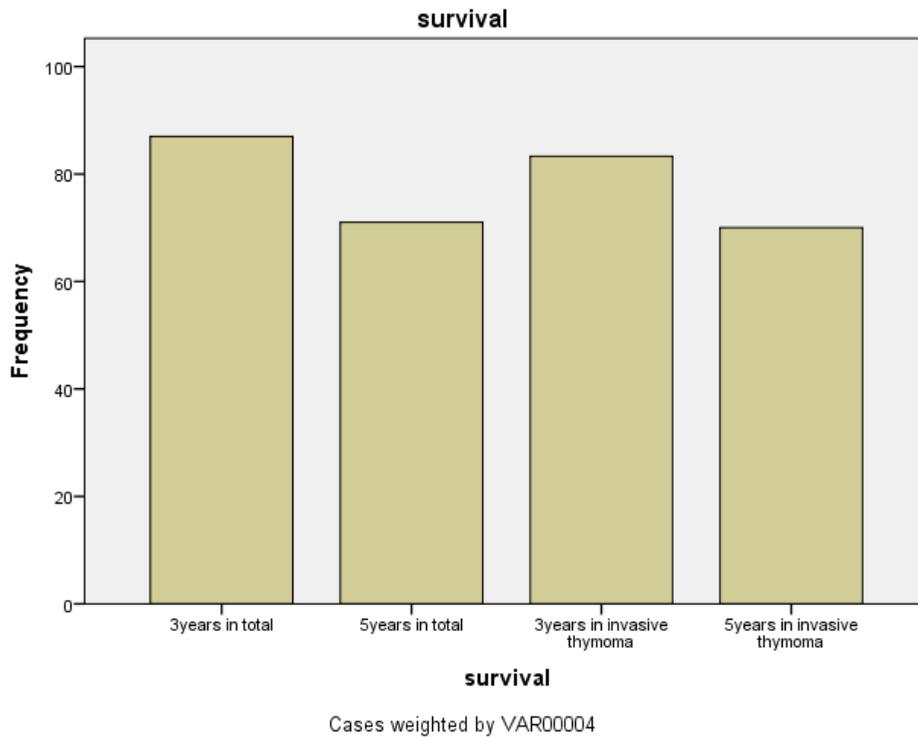


Figure 4: 3 and 5-year survival in all patients and survival of 3 and 5 years old with invasive tumors

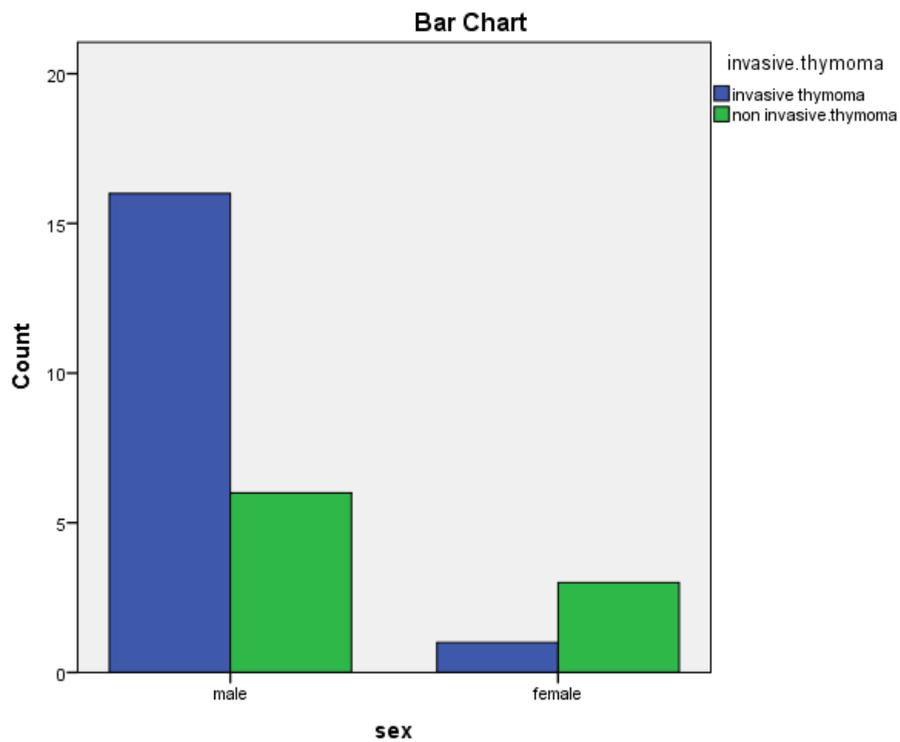


Figure 5: Frequency of male and female in terms of invasive and non-invasive tumor

Discussion

In this study, it was concluded that men with thymoma were more than women (5.5 to 1). Most patients were documented for the age range of 30- 50 and the mean age was 45.38 years. The age of patients was extended from 21 to 76 years old. In an article published by Eric A. Engels et al., a total number of 849 patients were studied and 455 men and 394 women (Male to female ratio of 1.15 to 1) reported in which from this point of view there was a significant difference in comparison with our study (6). In a study by BsSinghal et al., of 836 patients, 611 were men and 225 were women, which was similar to our study in terms of the number of men than women (12). The most accompanied rate of Myasthenia Gravis with type of invasive thymoma was seen in the fourth decade of age; however, this rate was observed for ages of 50 to 60 in a study by Zhei Feng Mao et al. (7). In a study by Bs singhal et al., this rate also was seen in decades of 60-70 ages, which was compatible with our study (12). According to the present study, of total seven patients with myasthenia gravis, 5 had invasive thymoma. On the whole, of 26 patients studied, 21 patients were alive and completed the study; however, five patients had died due to invasive thymoma. The rates of 3-year survival for all patients confirmed at 87% and 95% confidence intervals were equal to 1, 0.695. Overall, the 5-year survival rates achieved at 71% and 95% confidence interval were equivalent to 0.95 and 0.47. In addition, the 3-year survival rate in invasive tumors with confidence interval (1, 0.62) was 83.3% as well as a 5-year survival rate in the same tumor was equivalent to a 70% and confidence interval (0.995 ± 0.505), respectively. In the research by J. Andrea et al. (8), the 5 year survival rate was equivalent to 94%, which compared with our study (71%) there was a significant difference ($p < 0.001$). Furthermore, in the research by Kristen Bass Wilkins et al. (9), the 5 year survival rate of 87% was achieved, which compared with the current study (71%) there was a significant difference ($p = 0.02$). There was a significant relationship ($p < 0.001$) between the current study (71%) and Stefano Margaritora et al.'s study (89.9%) for the 5-year survival rate (10). In a study by Kazou Nakagawa et al. (11), 5-year survival rate was documented in 92%, which in comparison with our study (71%) there was a significant difference ($p < 0.001$). From a total number of 26 studied patients in the current study, seven patients had myasthenia gravis with 26.9%, which in comparison with F. Ysl et al.'s study with a total number of 39 patients who had myasthenia gravis and 15.7% there was a significant relationship (13).

Conclusion

The results of our study showed a significant relationship between invasive thymoma rate and developing myasthenia gravis. The 3-year and 5-year survival rate for all patients was equivalent to 83.3% and 71%, respectively. According to the study, most patients with invasive thymoma had a 5-year survival rate and also most of them had concomitant myasthenia gravis.

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Conflict of Interest

There is no conflict of interest to be declared.

Authors' contributions

All authors contributed to this project and article equally. All authors read and approved the final manuscript.

References

1. Hung AY, Eng TY, Scarbrough TJ, Dave Fuller C, Perez and Brady's Principles and Practice of Radiation Oncology. 5th ed. Philadelphia Hippincott, Williams and Wilkins. 2008.
2. Kondo K, Kinoshita H, Ishikura H, Miyoshi T, Hirose T, Matsumori Y, et al. Activation of matrix metalloproteinase-2 is correlated with invasiveness in thymic epithelial tumors. *J Surg Oncol*. 2001 Mar;76(3):169-75.
3. Willcox N, Schlupe M, Ritter A, Schuan JH, Newsom-Davis J, Christenson B. Myasthenic and non myasthenic thymoma. An expansion of minor cortical epithelial cell. *Am J Pathol. Japanese and cross medical center*. 2008; 58(8):489-493.
4. Keditsu KK, Karimundackal G, Jambhekar NA, Pramesh CS. Unusual clinical behavior of thymoma with recurrent myasthenia gravis. *Interact Cardiovasc Thorac Surg*. 2012; 14(6): 909-911.
5. Hung AY, Eng TY, Scarbrough TJ, Dave Fuller C, Perez and Brady's Principles and Practice of Radiation Oncology. 6th ed. Philadelphia Hippincott, Williams and Wilkins. 2013.
6. Engels EA, Pfeiffer RM. Malignant thymoma in the United States: demographic patterns in incidence and associations with subsequent malignancies. *Int J Cancer*. 2003; 105(4): 546-551.
7. Mao ZF, Mo XA, Qin C, Lai YR, Hackett ML. Incidence of thymoma in myasthenia gravis: a systematic review. *Journal of Clinical Neurology*. 2012; 8(3): 161-169.
8. McCart JA, Gaspar L, Incelet R, Casson AG. Predictors of survival following surgical resection of thymoma. *Eur J Surg Oncol*. 1993; 54(4):233-238.
9. Wilkins KB, Sheikh E, Green R, Patel M, George S, Takan M, et al. Clinical and pathologic predictors of survival in patients with thymoma. *Ann Surg*. 1999; 230(4): 56-9.
10. Margaritora S, Cesario A, Cusumano G, Meacci E, D'Angelillo R, Bonassi S, et al. Thirty-five-year follow-up analysis of clinical and pathologic outcomes of thymoma surgery. *Ann Thorac Surg*. 2010; 89(1):245-252.
11. Nakagawa K, Asamura H, Matsuno Y, Suzuki K, Kondo H, Maeshima A, et al. Thymoma: a clinicopathologic study based on the new World Health Organization classification. *J Thorac Cardiovasc Surg*. 2003; 126(4): 1134-1140.
12. Singhal BS, Bhatia NS, Umesh T, Menon S. A study from India in department of neurology Bombay Hospital Institute of Medical Science. *Neurol India*. 2008; 56(3):352-5
13. Aysal F, Baybas S, Selçuk HH, Sozmen V, Ozturk M, Kucukoglu H, et al. Paraneoplastic extralimbic encephalitis associated with thymoma and myasthenia gravis: Three years follow up. *Clin Neurol Neurosurg*. 2013; 115(5): 628-631.

