

Research Paper

Comparison between CT and MRI in the Diagnostic Accuracy of Thymic Masses

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Abstract

Purpose: The aim of this study was to compare diagnostic accuracy between CT and MRI for thymic masses.

Methods: We searched literature and collected information on first author, publication year, cases of different types of thymic lesions, correct diagnostic cases of CT and MRI and results of quantitative analysis of CT and MRI. The ROC curve was applied to compare the diagnostic performance of different imaging modalities.

Results: Eight literatures were finally included and analyzed in this study. There were 253 cases examined by CT and 340 cases by MRI in total. We showed outcomes of quantitative analysis of each study in this article. The sensitivity of CT and MRI was both 100%, while the specificity was 75% and 80%, respectively. AUC of CT was 0.875 [95%CI: 0.473, 0.997] and that of MRI was 0.880 [95%CI: 0.531, 0.995].

Conclusion: The diagnostic accuracy of MRI is superior to CT in detecting thymomas, thymic cysts or thymic hyperplasia but that of CT and MRI is still unclear in differentiating thymic carcinomas and lymphomas/germ cell tumors.

Key words: thymic mass, thymomas, CT, MRI, diagnostic accuracy

Introduction

Thymic masses are comprised of hyperplasia, cysts, thymic epithelial tumors (TETs), lymphomas, malignant germ cell tumors or metastatic cancers and so on [1, 2]. True thymic hyperplasia is usually regarded as a rebound phenomenon and characterized by an increase in mass of the gland after a stressor, such as chemotherapy, radiation, steroid treatment, burns or surgery [3]. Thymic cysts are relatively uncommon lesions that can be found at any age and can be congenital or acquired [2, 3]. Although TETs are rare neoplasms as to the whole tumors, they are the most common mediastinal tumors in adults, including thymomas and thymic carcinomas [4, 5]. The overall incidence of thymoma is 1.5 cases per million (www.nccn.org/). Thymoma is a slow-growing neoplasm that can invade to adjacent

structures, including the pericardium and pleura, whereas distant metastases are rare [6]. Thymic carcinomas are far rarer but more aggressive than thymomas, thus thymic carcinomas is often found having invaded to adjacent structures and metastasized to distant organ [7].

Thymectomy is the main therapeutic method of thymic masses, while radiotherapy and chemotherapy are often as (neo)adjuvant and palliative procedures [8, 9]. Generally, surgery is often performed under patients with paraneoplastic syndromes like myasthenia gravis [6, 10] or with thymic malignancy except for lymphoma. Therefore, patients would be overtreated or be in lack of essential therapy if thoracic surgeons could not differentiate the properties of thymic lesions. Imaging evaluation

has been employed to initially diagnose and properly staging thymoma, especially, on the detection of direct invasion and distant organ metastasis [6].

Computed tomography (CT) is currently the first choice to identify and characterize thymic masses due to the high spatial, temporal resolution and convenience [11-13]. Moreover, CT does offer a higher sensitivity and specificity for detecting of thymic lesions regardless of high radioactivity [12]. Some CT imaging parameters were significantly correlated with stage, classification and the completeness of resection of thymomas, but none can reliably predict overall survival or disease-free survival duration [11]. Magnetic Resonance imaging (MRI) is not routinely used to detect mediastinal masses like TETs except for patients with iodine allergy or with renal failure [14, 15]. However, MRI is better in detecting cystic lesion like thymic cysts or distinguishing cystic from solid masses than CT [15]. The typical appearance of thymoma in MRI is low to intermediate signal intensity on T1-weighted sequences and high signal intensity on T2-weighted sequences [15]. MRI has different functional modes like Chemical shift MRI and diffusion-weighted magnetic resonance imaging, thereby it can show better image contrast [16]. Additionally, MRI does have an excellent contrast resolution [15] and can be a better tool when patients are suspected to have invasive thymoma with adjacent structures invasion [17, 18].

However, it remains unclear which diagnostic modality is the most accurate diagnostic tool for thymic lesions. Herein, we try to unveil which imaging tool is better in accuracy of diagnosis of thymic masses.

Material and methods

Search strategy

A literature search was conducted (through 1st, January, 2009 to 1st, April, 2018) to identify all articles that have been published about accuracy of pre-operative CT and MRI examination. The terms "computed tomography", "CT", "magnetic resonance imaging", "MRI", "diagnosis", "accuracy", "detection", "thymic masses", "thymomas", "thymic cysts", "anterior mediastinal mass" and "thymic carcinomas" were searched in the title, abstract or key words on PubMed/MEDLINE, Web of science and Cochrane Library.

Inclusion and exclusion criteria

The inclusion criteria were as follows: 1) cohort studies; 2) diagnostic comparison between MRI and CT (these two tools combined or single modality); 3) staging and classification of thymomas were according to World Health Organization Histologic

Classification [19] or Masaoka-Koga stage system [20]; 4) all thymic masses were confirmed by histopathological examination. Case reports, reviews, editorials, abstracts, comments were all excluded.

Information collection

We collected information from each study on first author, publication year, cases of different types of thymic lesions and results of quantitative analysis of CT and MRI. Additionally, we found out correct and erroneous cases of each imaging in detecting thymic lesions.

Statistical analysis

Statistical analyses were performed with SPSS 23.0 for Windows software. ROC (receiver operating characteristics) analysis was applied to compare the diagnostic capability of different imaging modalities of thymic masses. The differences between groups were analyzed using a Student's t-test. P value < 0.05 was considered as significant.

Results

Publication screening

Sixteen studies were obtained after applying search strategy and screening abstracts, and eight studies were included further applying inclusion criteria [2, 21-27] (Table 1). There were 253 cases on CT examination and 340 cases on MRI in total. On CT, these included 99 thymomas, 12 thymic carcinomas, 22 thymic cysts, 75 thymic hyperplasia, 25 lymphomas or germ cell tumors and 20 other masses like seminoma. On MRI, these included 139 thymomas, 15 thymic carcinomas, 12 thymic cysts, 95 thymic hyperplasia, 47 lymphomas or germ cell tumors and 32 other masses.

Table 1. Basic information of included studies

Nation	Authors	Year	Cases	Mean age	Imaging modality
United States	B. Ackman et al.	2014	160	51(15-87)	CT
	T. Araki et al.	2014	18	56(37-79)	CT
Korea	Chang et al.	2017	37	54	CT
Italy	Priola et al.	2016	83	38.4(14-71)	CT/MRI
Japan	Seki et al.	2014	35	51.6(22-75)	CT/MRI
Japan	Tomiyama et al.	2009	127	45.9(10-82)	CT/MRI
Japan	Usuda et al.	2015	16	*	MRI
Japan	H. Yabuuchi et al.	2015	48	50.7(21-83)	MRI

*: unmentioned

Quantitative assessment of CT and MRI

Quantitative analysis was performed to compare percentage of correct diagnosis of thymic masses between CT and MRI. Percentage of correct diagnosis on CT is 77.78% in thymoma, 37.5% in thymic carcinoma, 61.36% in thymic cysts, 84.00% in thymic

hyperplasia, 66.00% in lymphoma/germ cell tumor, and 57.50% in other mass (Table 2). While percentage of correct diagnosis on MRI is 88.13% in thymoma, 30.00% in thymic carcinoma, 70.83% in thymic cysts, 97.89% in thymic hyperplasia, 48.94% in lymphoma/germ cell tumor and 54.69% in other mass (Table 3).

Table 2. Percentage of correct diagnosis by CT for all thymic mass.

	cases	correct	incorrect	No.(%)correct
Thymoma	99	77	22	77.78%
Thymic carcinoma	12	4.5	7.5	37.50%
Thymic cyst	22	13.5	8.5	61.36%
Thymic hyperplasia	75	63	12	84.00%
Lymphoma/germ cell tumor	25	16.5	18.5	66.00%
Other mass	20	11.5	8.5	57.50%

Table 3. Percentage of correct diagnosis by MRI for all thymic mass.

	cases	correct	incorrect	No.(%)correct
Thymoma	139	122.5	16.5	88.13%
Thymic carcinoma	15	4.5	10.5	30.00%
Thymic cyst	12	8.5	3.5	70.83%
Thymic hyperplasia	95	93	2	97.89%
Lymphoma/germ cell tumor	47	23	24	48.94%
Other mass	32	17.5	14.5	54.69%

Thymic cysts were generally 38 HU (Hounsfield unit, range 6-62 HU) on contrast-enhanced CT and are 45 HU (range 26-64 HU) on unenhanced CT. Table 4 showed quantitative assessment of CT in thymoma and thymic carcinoma by using HU and contrast HU.

Table 4. Quantitative assessment of CT in thymoma and thymic carcinoma*.

	mean Hounsfield unit	contrast Hounsfield unit
Thymoma		
low-risk thymoma	29.78	65.8(59.55-86.83)
high-risk thymoma	14.55	49.35(48.85-60.45)
thymic carcinoma	19.95	57.45(55.75-65.50)

*Data from Chang et al. 2017

Table 5. Quantitative assessment of MRI in different thymic masses.

	Yabuuchi et al. 2015	Usuda et al. 2015	Abdel Razek et al. 2014	Priola et al. 2015	Priola et al. 2015
Thymic mass	ADC($\times 10^{-3} \text{mm}^2/\text{s}$)				SII (%)
Low-risk thymoma	1.03 \pm 0.38	1.71 \pm 0.48	1.30 \pm 0.08	1.53 \pm 0.25	-0.81 \pm 3.56
High-risk thymoma	1.04 \pm 0.27		1.16 \pm 0.07	1.09 \pm 0.24	1.18 \pm 4.20
Thymic carcinoma	1.26 \pm 0.33	1.26 \pm 0.74	1.18 \pm 0.23	*	*
Thymic hyperplasia	*	*	*	1.86 \pm 0.17	35.51 \pm 12.51
Lymphoma	1.11 \pm 0.21	2.35 \pm 0.05	*	*	*

*: unmentioned

Apparent diffusion coefficient (ADC) and SII (which is obtained from the dual-echo technique without the need of an internal control and reflects the lipid content of the tissue) are important quantitative

parameter of MRI [28, 29]. Table 5 showed quantitative assessment of MRI in different thymic masses. Quantitative evaluation of MRI and CT in different types of thymoma was showed in Table 6. Of note, Priola et al. reported SII of thymic hyperplasia was 0.38 \pm 24.73% [25].

Table 6. Quantitative assessment of MRI and CT in different types of thymoma.

	Priola et al.2015	Priola et al. 2016	Priola et al. 2016
Thymoma	ADC*($\times 10^{-3} \text{mm}^2/\text{s}$)	SII ** (%)	Radiodensity (HU)***
Early thymoma	1.43 \pm 0.26	-0.95 \pm 3.65	33.64 \pm 10.82
Advanced thymoma	1.31 \pm 0.31	0.91 \pm 4.4	32.31 \pm 11.43

*: Apparent Diffusion Coefficient

**: SII is obtained from the dual-echo technique without the need of an internal control and reflects the lipid content of the tissue.

***: Radiodensity in Hounsfield unit

Comparison of diagnostic accuracy between CT and MRI

Diagnostic performance of CT and MRI in ROC curve was shown in Figure 1. At quantitative measurement, the sensitivity of CT and MRI were both 100%, while the specificities were 75% and 80%, respectively. Additionally, area under curve (AUC) of CT is 0.875 [95%CI: 0.473, 0.997] and that of MRI is 0.880 [95%CI: 0.531, 0.995].

Discussion

Thymoma is the most common type of neoplasms in the mediastinum and those with low risk and in early stage usually have good prognosis [9, 30]. Thymectomy is the mainstay of therapy of thymic masses and radiotherapy and chemotherapy are often as (neo)adjuvant and palliative procedures[8, 9]. Meanwhile, if thymoma or thymic carcinoma was discovered earlier, the rate of whole resection is high. Therefore, it is crucially important for thoracic surgeons to obtain accurate pre-operative identifications of staging and histological types of thymic masses.

Indeed, thymic lesions are uncommon and relative studies are rare compared with diseases involving other organs like lung. Therefore, only eight studies were finally included in our study in recent five years. Intriguingly, we found that the lower degree in which thymic masses invaded, and the higher CT attenuation were demonstrated. The performance of diagnosis of MRI was a little superior than that of CT in thymomas, thymic cysts and thymic hyperplasia, whereas differentiating thymic carcinomas, lymphomas/germ cell tumors and other masses were inferior to that of CT. In our study, ROC analysis was used to evaluate the diagnostic capability of CT and MRI, and they were both powerful modalities for thymic lesions. Of note, we

found that CT and MRI both have high sensitivity. AUC of CT and MRI was 0.875 and 0.880, respectively. Thus, we can safely conclude that the diagnostic capability of MRI is superior to that of CT in differentiating certain types of thymic masses combining with the results of statistical correctness of included studies.

CT is generally considered to be superior to MRI in the evaluation and characterization of most anterior mediastinal masses [15]. However, the versatility of MRI is superior to that of CT in certain scenarios, especially in the differentiation of thymic cysts [2]. Indeed, MRI is a promising modality to differentiate thymic masses because of high contrast resolution, no

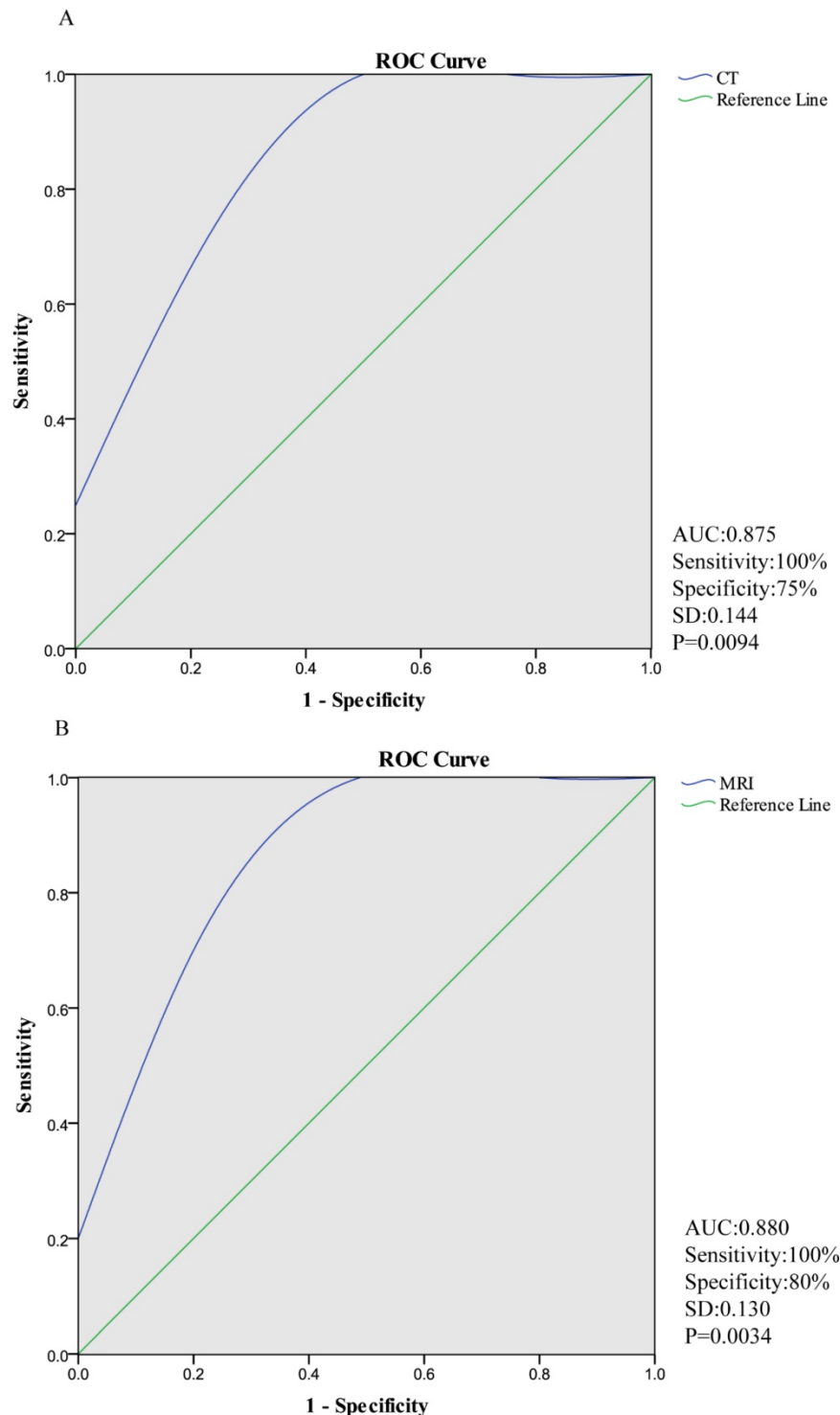


Figure 1. Diagnostic performance of CT and MRI in ROC curve. (The sensitivity of CT and MRI were both 100%, while the specificity were 75% and 80%, respectively. AUC (area under curve) of CT is 0.875 [95%CI: 0.473, 0.997] and that of MRI is 0.880 [95%CI: 0.531, 0.995].)

radiation exposure [28]. Chemical shift MRI (CS-MRI) is highly accurate in distinguishing thymomas from thymic lymphoid hyperplasia or normal thymus. However, it is not useful in differentiating between thymic lymphoid hyperplasia and normal thymus [28, 29]. Diffusion-weighted magnetic resonance imaging (DWI) could not only distinguish benign mediastinal tumors from malignant ones, but also distinguish among low-risk thymoma, high-risk thymoma and other malignant mediastinal tumors [16, 24]. DWI did not show excellent predictive ability of distinguishing early diseases from advanced ones based on Masaoka-Koga system [16, 31].

Priola et al. demonstrated that CS-MRI was more reliable than CT for distinguishing thymomas from thymic lymphoid hyperplasia and normal thymus through both qualitative and quantitative analyses, especially in evaluating patients with myasthenia gravis [25]. In qualitative assessment, MRI had higher accuracy than CT, and signal intensity of MRI was significantly different between groups [25]. Seki et al. [27] and Tomiyama et al. [2] both found that DWI is equal to CT in the diagnosis of thymomas. Additionally, Benveniste et al. concluded that the performance of MRI is similar to that of CT in distinguishing patients with advanced stage TETs on the 6th International Thymic Malignancy Interest Group Annual Meeting in October 2015 (www.itmig2015.org).

Our study demonstrated that the diagnostic utility of CT was inferior to that of MRI in detecting thymomas, thymic cysts or thymic hyperplasia, which was consistent with outcomes of some researchers [16, 24, 28, 29]. CT possesses advantages in the spatial, temporal resolution convenience and time-short as the first choice of detection of thymic masses, while MRI does well in detecting the relation between masses and their surrounding tissues especially connective tissues invasion such as vessels. In addition, we still cannot ignore that confounding diagnosis may occur when different masses performed same images. For instance, when rebound thymic hyperplasia occurs in patients who previously underwent chemotherapy for malignancy, distinguishing thymic rebound from recurrent neoplasm may be difficult [18]. Thymoma and thymic carcinomas may exhibit cystic changes both on CT and MRI [2]. Moreover, different diseases can co-exist such as hyperplasia and cysts, and sometimes this mixed lesion looks simply like a malignant tumor. Practically, differentiating thymomas from lymphomas or thymic hyperplasia was often difficult [26, 32]. Some researchers even demonstrated that CT could predict subtypes of TETs [33], but more evidence is required.

To assess whether pre-operative CT could identify those patients who should or not receive surgical therapies, Hayes et al. designed a study to evaluate the ability of CT in predicting the resectability of thymomas according to imaging characteristics. There were several preoperative imaging features including lobulated tumor contour and tumor sizes which showed significant association with an incomplete surgical resection or late stage thymomas on univariate analysis [34]. On multivariate analysis, greater than or equal to 50% abutment of the circumference of an adjacent vessel on CT was statistically associated with an incomplete resection or late stage thymomas [34].

Still, our study has some limitations. For example, the number of included studies was small due to the rarity of thymic masses. Next, further differentiation between contrast CT and non-contrast CT cannot be realized because of lack of relevant studies. Also, comparisons in different functional modes of MRI are unavailable. Therefore, more researches are warranted to further elucidate the capabilities of MRI to predict whether thymic lesions need to be resected.

Conclusion and future expectation

The diagnostic accuracy of MRI is superior to CT in detecting thymomas, thymic cysts or thymic hyperplasia and that of CT and MRI is still unclear in differentiating thymic carcinomas and lymphomas/germ cell tumors and mere image analysis is insufficient for accurate diagnosis of thymic masses. Imaging plays a crucial role in diagnosis, staging and even treatment of patients with thymic masses, but both CT and MRI have their own shortcomings and cannot provide accurate pre-operative diagnosis. Therefore, it may be the future direction of diagnosing thymic diseases that trying to find more reliable biomarkers.

Abbreviations

TETs, thymic epithelial tumors; AUC, area under curve; ADC, apparent diffusion coefficient; MRI, magnetic resonance imaging; CT, computed tomography.

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Competing Interests

The authors have declared that no competing interest exists.

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