



Coexisting active pulmonary tuberculosis in tuberculous spondylitis: the prevalence and the role of chest CT

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1		31
2		32
3	In clinical practice, tuberculous (TB) spondylitis alone does	33
4	not warrant patient isolation (1). We experienced patients	34
5	with underlying TB spondylitis involved the appearance of	35
6	only subtle lung lesions on the chest radiograph, but chest	36
7	computed tomography (CT) scans showed active pulmonary	37
8	TB. If the coexisting lung lesion is present, it will be	38
9	helpful to early diagnosis and treatment of TB. However,	39
10	a coexisting lung lesion is easy to overlook, particularly on	40
11	plain chest radiography. And there is a lack of clear evidence	41
12	that TB spondylitis will require additional chest CT scans	42
13	in previous studies. Therefore, we hypothesized that pre-	
14	diagnostic consideration of the coexistence of an active lung	43
15	lesion in patients with suspicious TB spondylitis can help	44
16	diagnose TB.	45
17	We retrospectively reviewed consecutive 50 patients with	46
18	a histological or microbiologically confirmed diagnosis of	47
19	TB spondylitis among surgically confirmed TB spondylitis	48
20	between January 2005 and December 2015 (IRB No.	49
21	KHU 2017-08-030). Patients' demographic information	50
22	and medical history were reviewed using clinical charts.	51
23	Two radiologists (So Youn Shin and Eun Jung Shim)	52
24	retrospectively reviewed all images to reach consensus. We	53
25	analyzed spine CT scans and magnetic resonance (MR)	54
26	images for each affected level and also reviewed chest	55
27	plain radiographs and CTs to evaluate lung involvement	56
28	in TB. We defined active pulmonary TB as follows: (I)	57
29	centrilobular nodules, (II) branching linear opacities with	58
30	nodularity (tree-in-bud sign), and (III) lobular or patchy	59
	lesions of consolidation and cavitation (2). The presence	60
	of miliary TB and pleural effusion were also analyzed. We	
	also reviewed microbiological analyses to determine the	
	level of activity of TB. Statistical analysis was performed to	
	investigate the presence of concomitant active pulmonary	
	TB in TB spondylitis patients according to the level of TB	
	spondylitis using SPSS 23 (Statistical Package for Social	
	Science, version 23.0, IBM Corporation, Chicago, IL, USA)	
	and R 3.5.1 (http://cran.r-project.org), with P values below	
	0.05 considered statistically significant.	
	Among 50 patients with surgically confirmed TB	
	spondylitis, there was no significant difference in gender	
	(male: female =24:26) and the mean age was 51.82±18.79 years	
	old (range, 15–79 years). Only nine (18%) had a history	
	of pulmonary TB. <i>Table 1</i> shows the results of radiologic	
	image analysis. The most frequently involved region of the	
	spine was lower T (T7–12) regions, and the most frequently	
	affected region was T12. Twenty-one (42%) showed	
	concomitant active pulmonary TB on radiologic image	
	analysis. In microbiologic results, of the 21 patients with TB	
	spondylitis who had concomitant active pulmonary TB on	
	radiologic image analysis, only one (4.76%) tested positive	
	in a sputum AFB and was therefore regarded as potentially	
	contagious at initial clinical diagnosis. In statistical analysis,	
	TB spondylitis involving the upper (C or T) spinal region	
	had a significant correlation with coexisting active pulmonary	
	TB (P=0.0033), compared with lower spine involvement	
	(<i>Table 2</i>).	

Table 1 Results of radiologic image analysis of distribution of affected spinal levels of TB spondylitis and evidence of possibility of concomitant active pulmonary TB

Variables	Number (n)	Percentage (%)
Involved spine level [†]	137 of 50 patients	
C (C1–7)	6	4.38
Upper T (T1–6)	13	9.49
Lower T (T7–12)	55	40.15
L (L1–L5)	53	38.69
LS (S1–S5)	10	7.30
Evidence of possibility of concomitant active pulmonary TB		
Positive (+)	21	42
Results of performed radiologic image		
Chest radiograph	21	100
Chest CT	17	80.95
Radiologic image analysis ^{††}		
Centrilobular nodules	10	47.62
Branching linear opacities with nodularity (tree-in-bud)	2	9.52
Lobular or patchy lesions of consolidation and cavitation	3	14.29
Miliary TB	4	19.05
Pleural effusion or empyema	7	33.33
Negative (–)	29	58
Inactive pattern on radiologic image analysis	2	6.90
Without any lung lesion	27	93.1

[†], if the affected vertebrae is continuous or multiple levels, we separately counted the number of vertebrae body. ^{††}, if the images showed two or more findings, we described multiple counts of findings, respectively. TB, tuberculous.

Table 2 Univariate and multivariate logistic regression analysis for the possibility of TB with concomitant active pulmonary tuberculosis

Involved spine level ^{††}	Univariate			Multivariate [†]		
	P value	Adjusted OR	95% CI	P value	Adjusted OR	95% CI
C or T spine	0.0059	7.38	1.78–30.69	0.0033	12.75	2.33–69.70
T or L spine	0.3911	0.34	0.03–4.01	0.4614	0.37	0.03–5.23
L or S spine	0.0685	0.34	0.25–34.85	0.0579	0.29	0.08–1.04

[†], backward elimination; ^{††}, if the affected vertebral levels from C6 to T2, we counted the lesion as both C and T spine involvement. Note: P values of less than 0.05 are regarded as statistically significant. TB, tuberculous; OR, odds ratio; CI, confidence interval.

61 According to several studies of affected levels of TB
62 spondylitis, lower thoracic and upper lumbar regions are the
63 most commonly affected sites (3–5). In our study, consistent
64 with previous studies, the lower T (T7–12) regions were the
65 most commonly involved sites.

66 It is known that the probability of concomitant
67 pulmonary TB in TB spondylitis patient shows wide
68 variation among countries (4,6). In an article of literature
69 review by Schirmer *et al.* (4), the probability of concomitant
70 pulmonary TB in TB spondylitis patients varies from

8% to 100%. In recent large scale studies for spinal TB, the incidences of concomitant pulmonary TB show from 14.37% to 28% (7-9). The lower rates of concomitant pulmonary TB in those studies compared to our study (42%) may be related to differences in diagnosis of 'active pulmonary TB'—clinical versus imaging assessment. In clinical practice, sputum acid-fast bacilli (AFB), culture, or TB polymerase chain reaction (PCR) are not sensitive enough for screening for active pulmonary TB. We demonstrated that TB spondylitis involving the upper (C or T) spinal regions was significantly correlated with a coexisting active pulmonary TB. Therefore, we suggest that a patient with TB spondylitis involving the upper (C or T) spinal region would need to be assessed the concomitant active pulmonary TB. Moreover, because patients with TB spondylitis often complain of back pain and this condition can make it difficult to undergo a posteroanterior chest radiograph in the erect position and therefore to delineate lung nodules on a spine CT, we want to emphasize that radiologic imaging could be an additional approach for diagnosis of TB to avoid missing cases of potentially active pulmonary TB.

A negative AFB smear is commonly regarded as having a low infectivity and is common at initial diagnosis, which make it difficult to diagnosis and treat the disease early. However, respiratory transmission could also occur (17%) from person with sputum smear-negative TB (10,11). In our study, among concomitant active pulmonary TB on radiologic image analysis, only one patient (4.76%) tested positive in sputum AFB test and three patients (14.29%) showed positive in sputum TB PCR and regarded as potentially contagious at initial clinical approach. Chest radiographs are not specific and it can appear normal even when the disease is present (12). Neither military tuberculosis nor pleural effusion is usually not considered as infectious, however, these findings may help in the diagnosis of spinal TB. This suggests greater attention should be paid to the potential for TB transmission despite negative smear results.

In conclusion, we found 42% of TB spondylitis had a coexisting potentially active pulmonary TB lesion on radiologic image analysis. This literature showed the higher co-morbidity of active pulmonary TB and TB spondylitis, the higher potential risk of nosocomial infection of TB. Because the possibility of concomitant active pulmonary tuberculosis in TB spondylitis patient can be easily overlooked, there is a chance of the possibility of nosocomial infection of TB. And we recommend that chest

CT (at least low-dose chest CT) would be useful in the initial evaluation of TB spondylitis, especially with upper (C or T) spinal region involvement, in spite of subtle evidence of active pulmonary TB on plain chest radiography. It would be helpful in diagnosing TB earlier and preventing airborne dissemination.

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