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## EVALUATION FOR CT CRITERIA OF GRANULOMATOUS INFECTION AND TUMOUR OF SPINE

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### ABSTRACT

To develop criteria to distinguish between granulomatous infection and neoplastic process in the spine by mean of the computed tomography scan (CT scan), the authors retrospectively analyzed 31 cases of granulomatous infection and 13 cases of neoplastic disease. The result was that the reliable criteria for granulomatous infection were diffuse bony destruction, contiguous levels of spinal involvement, absence of posterior element involvement, presence of "complete" pattern of prevertebral soft tissue, and presence of disc involvement. Neoplastic diseases were characterized by focal bony destruction, separated levels of spinal involvement, presence of posterior element involvement, "partial" pattern of prevertebral soft tissue, and absence of disc involvement. Blinded testing of these criteria is potential for improving diagnostic accuracy in clinical practice.

### INTRODUCTION

Computed tomographic examination of the spine is an available method in Thailand for diagnosis and characterization of vertebral lesions. The importance of CT is underscored by the fact that clinical assessment and conventional radiographic analysis are occasionally inconclusive in distinguishing between granulomatous infection and neoplastic process of the spine. This retrospective study was designed to identify reliable CT criteria for distinction between these two major categories of disease involving the spine.

### MATERIALS AND METHOD

Transaxial CT image from a total of 44 cases of granulomatous infection and neoplastic disease affecting the spine were retrospectively evaluated. The pathologic identity of each lesion was determined with surgical specimen (36/44) and assumed

on the basis of known widespread metastasis in the remainders (8/44).

There are 31 cases of granulomatous infections and 13 cases of neoplastic diseases examined. The types of tumor included adenocarcinoma from unknown primary site, breast carcinoma, lung carcinoma, hepato-cellular carcinoma, multiple myeloma, and hemangioma. The authors defined the CT findings for which each site of involvement would be evaluated in an attempt to identify distinguishing features between the two pathologic categories.

Each CT study was assessed for [1]

- a. Presence or absence of prevertebral (anterior to lamina) soft tissue involvement, and when present, whether this was partial or complete (lamina to lamina) (Fig. 1,3).
- b. Presence or absence of bony destruction and whether this was focal, multifocal or diffuse (Fig 2,4,5).
- c. Presence or absence of posterior element involvement (Fig.1,3)



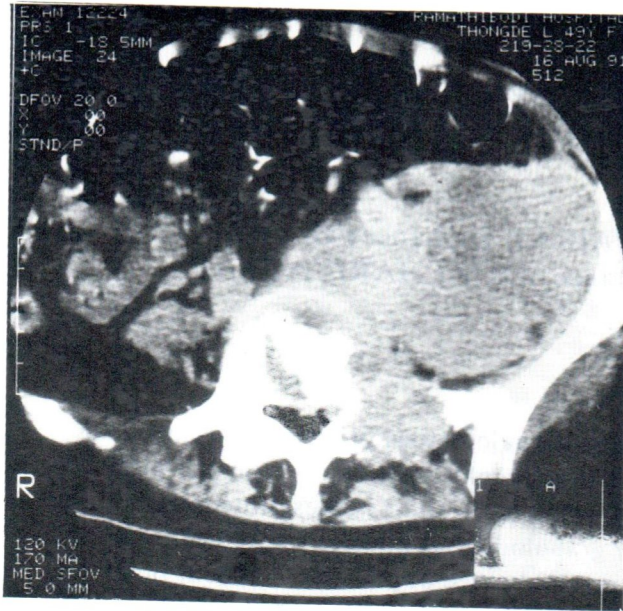


Fig 1. Breast carcinoma metastasized to L<sub>5</sub>. Characteristic features are posterior element involvement and partial prevertebral soft tissue involvement.



Fig 2. The same case as fig 1. The image viewed in bone window also showed focal bony destruction

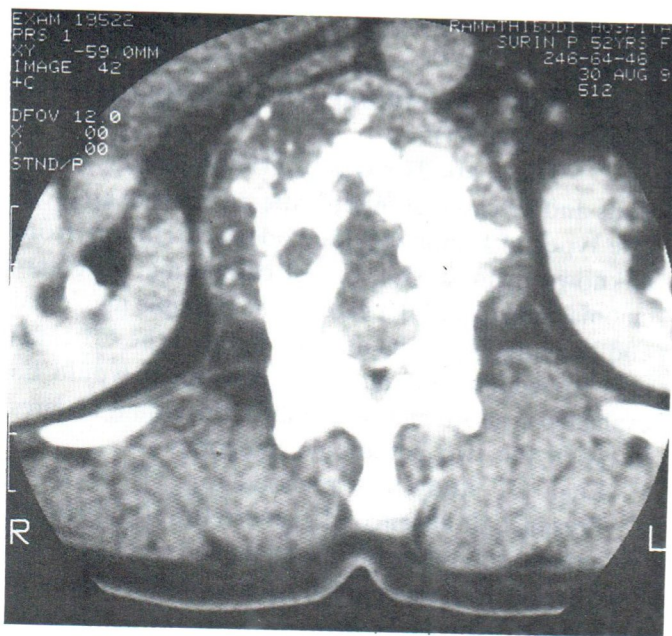


Fig 3. Pathologic tissue proved to be caseous granulomatous infection at L<sub>1</sub>. Complete pre-vertebral soft tissue involvement and extension into the spinal canal are noted.



Fig 4. Pathologic tissue proved to be granulomatous infection at T<sub>11</sub>. The image revealed diffuse bony destruction

- d. Presence or absence of multilevel disease and whether this was contiguous (across an intervertebral disc space) or separated.
- e. Presence or absence of disc involvement.

A statistical analysis of the results were performed to determine which CT findings were most valuable in distinguishing between granulomatous infection and tumoral process.



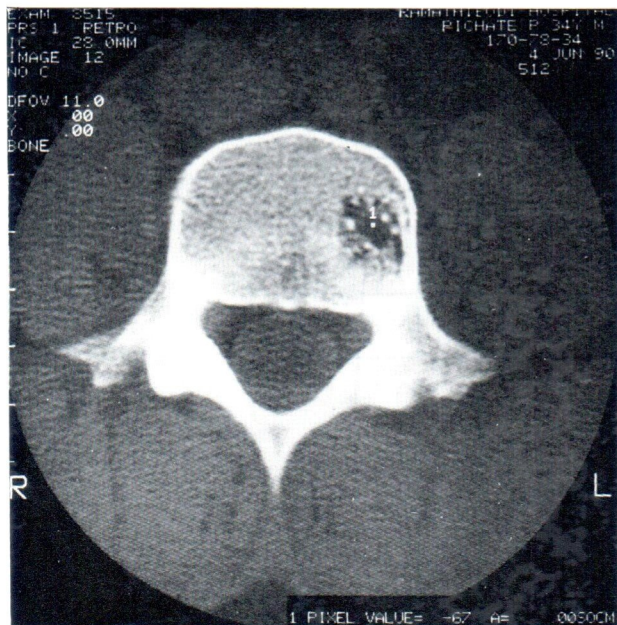


Fig 5. Pathologic tissue proved to be haemangioma of L4. The image showed focal bony destruction

**DISCUSSION**

Granulomatous infection are grouped with similarities in clinical presentation and histologic feature of granulomatous identification on frozen section. They may caused by fungi, certain bacteria and spirochetes. The most common granulomatous spinal

infection in the world by far is tuberculosis. As in our study, all granulomatous infection (31 cases) are tuberculosis. These infections spread hematogenously and subligamentously [2,3,4,5]. The latter route of spreading causes more prevertebral soft tissue (sensitivity 1.0, specificity 0.53, p = 0.00004) and contiguous levels of spinal involvement (sensitivity 1.0, specificity 0.25, p = 0.042) than those in neoplastic processes. Criteria of diffuse bony destruction in granulomatous diseases in this study (sensitivity 0.86, specificity 0.67, p = 0.004) differs from the previous study of Van Lom et.al [1]. They reported criteria of focal bony destruction in this disease (sensitivity 1.0, specificity 1.0). This was likely due to late presentation of many cases of granulomatous infection in Thailand. Absence of posterior element involvement is also a reliable criteria for granulomatous infection (sensitivity 0.61, specificity 0.84, p = 0.014) due to in this disease there are three major primary foci of spinal involvement : peridiscal, central body and anterior body locations. In one series of 914 cases, the disease was at peridiscal location 33% , central body 11.6% , and anterior body in 2.1%. In 52.8% of cases, the disease was widespread at presentation [5].

Neoplastic process of the spine arises from focal lesions or from distant malignancy. Local involvement of the spine may result from primary lesion arising in the spinal cord, its coverings, or contiguous spread of tumour from the paraspinous

**Results**

**Table 1**

|                          |          | Granulomatous infection | Tumor |    |
|--------------------------|----------|-------------------------|-------|----|
| Prevertebral soft tissue | Presence | 31                      | 6     | 37 |
|                          | Absence  | 0                       | 7     | 7  |
|                          |          | 31                      | 13    | 44 |

Analysis of Single Table Odd ratio : Undefined

**Chi-Squares**

**P-values**

|                 |                  |           |
|-----------------|------------------|-----------|
| Yates corrected | 16.03            | 0.0000624 |
| Fisher exact    | 1-tailed P-value | 0.0000448 |
|                 | 2-tailed P-value | 0.0000448 |
|                 | Sensitivity      | 1.0       |
|                 | Specificity      | 0.53      |

**Table 2**

|                          |          | Granulomatous infection | Tumor |    |
|--------------------------|----------|-------------------------|-------|----|
| Prevertebral soft tissue | complete | 24                      | 0     | 24 |
|                          | partial  | 7                       | 6     | 13 |
|                          |          | 31                      | 6     | 37 |

Analysis of Single Table Odd ratio : Undefined

**Chi-Squares**

**P-values**

|                               |       |           |
|-------------------------------|-------|-----------|
| Yates corrected               | 10.04 | 0.0015301 |
| Fisher exact 1-tailed P-value |       | 0.0007381 |
| 2-tailed P-value              |       | 0.007331  |
| Sensitivity                   | 0.77  |           |
| Specificity                   | 1.0   |           |

**Table 3**

|                             |         | Granulomatous infection | Tumor |    |
|-----------------------------|---------|-------------------------|-------|----|
| Pattern of bony destruction | diffuse | 26                      | 3     | 29 |
|                             | focal   | 4                       | 6     | 10 |
|                             |         | 30                      | 9     | 39 |

Analysis of Single Table Odd ratio = 13.00

**Chi-Squares**

**P-values**

|                               |      |           |
|-------------------------------|------|-----------|
| Yates corrected               | 7.72 | 0.0054599 |
| Fisher exact 1-tailed P-value |      | 0.0038571 |
| 2-tailed P-value              |      | 0.0038571 |
| Sensitivity                   | 0.86 |           |
| Specificity                   | 0.67 |           |

soft tissue and lymphatics. Regional and distant spread of metastatic disease to spine may occur with almost any of the solid tumour of the body, with osseous malignancy of appendicular skeleton, and with systemic lymphoreticular malignancy such as multiple myeloma and lymphoma. Metastases are the most common

skeletal tumour , and the spine is the most common site of skeletal involvement. They are most common secondary to carcinoma of the breast, lung and prostate gland. [5,6]

Absence and partial prevertebral soft tissue involvement in neoplastic processes of the spine is

**Table 4**

|                               |          | Granulomatous infection | Tumor |    |
|-------------------------------|----------|-------------------------|-------|----|
| Posterior element involvement | absence  | 19                      | 2     | 21 |
|                               | presence | 12                      | 11    | 23 |
|                               |          | 31                      | 13    | 44 |

Analysis of Single Table Odd ratio : 8.71

| <b>Chi-Squares</b>   | <b>P-values</b> |
|----------------------|-----------------|
| Yates corrected 6.01 | 0.0142584       |
| Sensitivity 0.61     |                 |
| Specificity 0.84     |                 |

**Table 5**

|                                       |            | Granulomatous infection | Tumor |    |
|---------------------------------------|------------|-------------------------|-------|----|
| Multiple levels of spinal involvement | contiguous | 29                      | 6     | 35 |
|                                       | seperated  | 0                       | 2     | 2  |
|                                       |            | 29                      | 8     | 37 |

Analysis of Single Table Odd ratio = Undefined

| <b>Chi-Squares</b>            | <b>P-values</b> |
|-------------------------------|-----------------|
| Yates corrected 3.55          | 0.0593750       |
| Fisher exact 1-tailed P-value | 0.042042        |
| 2-tailed P-value              | 0.042042        |
| Sensitivity 1.0               |                 |
| Specificity 0.25              |                 |

a reliable criteria (sensitivity 0.53, specificity 1.0,  $p = 0.00004$  and sensitivity 1.0, specificity 0.77,  $p = 0.0007$  respectively) because most neoplasm spread by hematogenous and lymphatic routes, but less likely by subligamentous route. These also caused more presence of multiple seperated levels of spinal involvement (sensitivity 0.25, specificity 1.0,  $p = 0.042$ ) than in granulomatous infections. Criteria of focal pattern of bony destruction (sensitivity 0.67, specificity 0.86,  $p = 0.004$ ) is reliable due to marrow

replacement in any degree by the tumour cells but granulomatous infection usually causes diffuse fragmentation of the vertebral bodies [8]. The intervertebral discs are spared in tumoural process (sensitivity 1.0, specificity 1.0) owing to their resistance to tumour invasion and is a reliable criteria for tumoural process, while presence of disc involvement (sensitivity 1.0, specificity 1.0) is observed in granulomatous infection.



**Table 6**

|                  |          |                         |       |    |
|------------------|----------|-------------------------|-------|----|
|                  |          | Granulomatous infection | Tumor |    |
| Disc involvement | presence | 31                      | 0     | 31 |
|                  | absence  | 0                       | 13    | 13 |
|                  |          | 31                      | 13    | 44 |

Analysis of Single Table Odd ratio : Undefined

**Chi-Squares**

**P-values**

Yates corrected 39.33  
 Sensitivity 1.0  
 Specificity 1.0

0.0000

**Table 7**

|                          |             |       |                         |    |
|--------------------------|-------------|-------|-------------------------|----|
|                          |             | Tumor | Granulomatous infection |    |
| Prevertebral soft tissue | absence     | 7     | 0                       | 7  |
|                          | soft tissue | 6     | 31                      | 37 |
|                          |             | 13    | 31                      | 44 |

Analysis of Single Table Odd ratio = Undefined

**Chi-Squares**

**P-values**

Yates corrected 16.03  
 Fisher exact 1-tailed P-value  
 2-tailed P-value  
 Sensitivity 0.53  
 Specificity 1.0

0.0000624  
 0.0000442  
 0.000448

The limitation of this study is its purely retrospective nature, and all granulomatous infections in this study were tuberculosis so this study may mainly represent the spinal tuberculosis rather than other granulomatous infections.

In conclusion, under these criteria for CT interpretation of spinal pathology, it can be practical benefit in the patients with clinical undiagnostic

between granulomatous infection (particularly tuberculosis) and neoplastic process [7].

**ACKNOWLEDGEMENT :**

We would like to thank Ms.Amarin Taksinsathian for the statistical analysis

**Table 8**

|                          |          |                         |    |
|--------------------------|----------|-------------------------|----|
|                          | Tumor    | Granulomatous infection |    |
| Prevertebral soft tissue | partial  | 7                       | 13 |
|                          | complete | 24                      | 24 |
|                          |          | 31                      | 37 |

Analysis of Single Table Odd ratio : Undefined

**Chi-Squares**

**P-values**

|                               |       |           |
|-------------------------------|-------|-----------|
| Yates corrected               | 10.04 | 0.0014301 |
| Fisher exact 1-tailed P-value |       | 0.0007381 |
| 2-tailed P-value              |       | 0.0007381 |
| Sensitivity                   | 1.0   |           |
| Specificity                   | 0.77  |           |

**Table 9**

|                  |         |                         |    |
|------------------|---------|-------------------------|----|
|                  | Tumor   | Granulomatous infection |    |
| Bony destruction | focal   | 4                       | 10 |
|                  | diffuse | 26                      | 29 |
|                  |         | 30                      | 39 |

Analysis of Single Table Odd ratio = Undefined

**Chi-Squares**

**P-values**

|                               |      |           |
|-------------------------------|------|-----------|
| Yates corrected               | 7.72 | 0.0054599 |
| Fisher exact 1-tailed P-value |      | 0.0038571 |
| 2-tailed P-value              |      | 0.0038571 |
| Sensitivity                   | 0.67 |           |
| Specificity                   | 0.86 |           |

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**Table 10**

|                               |          |                         |    |
|-------------------------------|----------|-------------------------|----|
|                               | Tumor    | Granulomatous infection |    |
| Posterior element involvement | presence | 11                      | 12 |
|                               | absence  | 2                       | 19 |
|                               |          | 13                      | 31 |
|                               |          |                         | 44 |

Analysis of Single Table Odd ratio = 8.71

**Chi-Squares**

**P-values**

Yates corrected 6.01  
 Sensitivity 0.84  
 Specificity 0.61

0.0142584

**Table 11**

|                                       |            |                         |    |
|---------------------------------------|------------|-------------------------|----|
|                                       | Tumor      | Granulomatous infection |    |
| Multiple levels of spinal involvement | seperated  | 2                       | 0  |
|                                       | contiguous | 6                       | 29 |
|                                       |            | 8                       | 29 |
|                                       |            |                         | 37 |

Analysis of Single Table Odd ratio = Undefined

**Chi-Squares**

**P-values**

Yates corrected 3.55  
 Fisher exact 1-tailed P-value  
 2-tailed P-value  
 Sensitivity 0.25  
 Specificity 1.0

0.0593750  
 0.0420420  
 0.0420420

|   |   |
|---|---|
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|---|---|



**Table 12**

|                  |          |       |                         |    |
|------------------|----------|-------|-------------------------|----|
|                  |          | Tumor | Granulomatous infection |    |
| Disc involvement | absence  | 13    | 0                       | 13 |
|                  | presence | 0     | 31                      | 31 |
|                  |          | 13    | 31                      | 44 |

Analysis of Single Table Odd ratio = Undefined

| <b>Chi-Squares</b>    | <b>P-values</b> |
|-----------------------|-----------------|
| Yates corrected 39.33 | 0.00000         |
| Sensitivity 1.0       |                 |
| Specificity 1.0       |                 |

**Table 13**

Retrospective analysis of CT examinations reveals the reliable criteria as followed :

|  | Sensitivity | Specificity | P-values |
|--|-------------|-------------|----------|
| <u>Reliable criteria for granulomatous infection of spines</u> |             |             |          |
| - presence of prevertebral soft tissue                         | 1.0         | 0.53        | 0.00004  |
| - complete pattern of prevertebral soft tissue                 | 0.77        | 1.0         | 0.0007   |
| - diffuse bony destruction                                     | 0.86        | 0.67        | 0.004    |
| - absence of posterior element involvement                     | 0.61        | 0.84        | 0.014    |
| - contiguous levels of spinal involvement                      | 1.0         | 0.25        | 0.042    |
| - presence of disc involvement                                 | 1.0         | 1.0         | -        |
| <u>Reliable criteria for neoplastic process of spines</u>      |             |             |          |
| - absence of prevertebral soft tissue                          | 0.53        | 1           | 0.00004  |
| - partial pattern of prevertebral soft tissue                  | 1.0         | 0.77        | 0.0007   |
| - focal bony destruction                                       | 0.67        | 0.86        | 0.004    |
| - presence of posterior element involvement                    | 0.84        | 0.61        | 0.014    |
| - separated levels of spinal involvement                       | 0.25        | 1.0         | 0.042    |
| - absence of disc involvement                                  | 1.0         | 1.0         | -        |