

### Should brucellosis be a part of differential diagnosis when tuberculous meningitis is predicted with Thwaites or Lancet scoring systems?

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**BACKGROUND:** Brucellar meningoencephalitis is one of the leading causes of chronic central nervous system (CNS) infections together with tuberculous meningitis (TBM). Since culture is very slow and non-culture diagnosis lacks sensitivity, Thwaites and Lancet scoring systems have been used in the rapid diagnosis of TBM. According to our clinical experience these two methods may misdiagnose brucellar CNS disease as TBM. Thus, we designed this multicenter study to test this hypothesis.

**METHODS:** 294 patients with brucellar meningoencephalitis treated in 35 tertiary hospitals were included. Neurobrucellosis inclusion criteria were all of the following: (i) the presence of clinical symptoms consistent with either meningitis or meningoencephalitis (ii) the presence of typical cerebrospinal fluid (CSF) findings consistent with meningitis, (iii) the presence of positive culture or serological tests for brucellosis in the blood or in the CSF, (iv) the absence of an alternative neurological diagnosis. The control group comprised 190 patients with TBM. Inclusion criteria were clinical evidence of meningitis and microbiological confirmation of TBM. The microbiological confirmation included culture, PCR analysis and Ehrlich-Ziehl-Neelsen staining from the CSF. Both patient groups were compared by using *Mann-Whitney U test*. A score of 12 was assigned as "probable", 6-11 as "possible", and less than 5 were noted as negative for tuberculous meningitis. According to the categories of both scores two patient groups were compared with *chi-square tests*. In testing the correlation of both systems "*Pearson test*" was used. The value of  $p < 0.05$  was accepted as significant.

**RESULTS:** The mean and median values of Thwaites and Lancet scoring systems for both groups are presented in table 1. The distribution of the scores in two groups were significantly different ( $p = 0.001$  and  $p < 0.001$  respectively). Added to that, there was a significant difference between the diagnostic categories for both prediction systems (Table).

According to Lancet scoring system, 88.1% of neurobrucellosis cases were classified as possible and 8.8% were categorized as probable while 51.6% of TBM cases were reported as possible and 45.3% were as probable. On the other hand, 99.3% of neurobrucellosis patients and 95.8% of microbiologically confirmed TBM patients met the criteria for TBM.

**CONCLUSION:** Brucellar meningoencephalitis can be easily confused with TBM when Thwaites and Lancet prediction systems are used. Thus, brucellosis should be considered in the differential diagnosis of TBM in endemic countries.

**Table. Mean, median values and categorical classification of Thwaites and Lancet scores**

Neurobrucellosis	TBM	p
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	n=294	n=190	
<u>Thwaites scoring system</u>			
Median score	-3	-3	<b>0.001*</b>
Mean±SD score	-3.11±2.30	-2.09±3.11	
Diagnostic classification			<b>0.017#</b>
TBM	292 (%99.3)	182 (%95.8)	
Bacterial meningitis	2 (0.7%)	8 (4.2%)	
<u>Lancet scoring system</u>			
Median score	10	11	<b>&lt;0.001*</b>
Mean±SD score	9.43±1.71	11.45±3.01	
Diagnostic categories			<b>&lt;0.001#</b>
Probable	26 (8.8%)	86 (45.3%)	
Possible	259 (88.1%)	98 (51.6%)	
Negative	9 (3.1%)	6 (3.2%)	

\**Mann-Whitney U test*

#*Chi-square test*

TBM: Tuberculous meningitis