Indian J Physiol Pharmacol 2001; 45 (1):119-121

LETTER TO THE EDITOR

RELEVANCE OF SERUM ALKALINE PHOSPHATASE AS A DIAGNOSTIC AID IN LUNG PATHOLOGY

Sir,

(Received on September 14, 2000)

Alkaline phosphatase (ALP, 3.1.3.1) is a dimeric glycoprotein present in humans that catalyses the hydrolysis of orthophosphoric monoesters at alkaline $P_{\mu}(1)$. It has been reported to occur in abundance in human bile canaliculi, bone, intestine, placenta and kidney (1). Hence serum ALP levels has been extensively used as a diagnostic tool for hepatobilliary disorders particularly in patients with extrahepatic cholestasis (2). It has also been suggested to be useful in the diagnosis of malignancies and diseases affecting the liver (3,4), bone (5) and kidney (6). However, in the last few years a number of workers have reported serum ALP as a useful diagnostic parameter in certain types of lung disorders (7-9). In the light of this information available in scientific literature the aim of the present work was to evaluate and confirm the extent of utility of serum ALP as a diagnostic marker for various lung pathology.

Serum was separated from 5 ml of venous blood collected in aseptic syringe without anticoagulants from patients with different types of lung diseases admitted to Wenlock Government Hospital and Attavar Cancer Hospital located in Mangalore, Karnataka. The diagnosis of these patients were confirmed by the physicians of respective hospitals with the aid of results of various tests such as sputum and blood analysis, chest X-ray, ultrasound and CT scan. Samples were taken only from patients with confirmed diagnosis admitted to the respective hospitals who were yet to be put on regular treatment regimen. Normal controls constituted adults who were students and staff of the department. Serum was analysed on the day of collection without storage. ALP was measured as reported (10) using disodium phenylphosphate as the substrate. The phenol liberated by ALP action was estimated using 4-amino antipyrine and the colour developed analysed at 520 mm and calculation done using appropriate phenol standard. All chemicals used were of reagent grade. Statistical analysis was done based on students 't' test (11) and values expressed as mean ± standard deviation (SD).

The average mean serum ALP values when calculated separately for each of the lung disease showed values higher than that obtained for normal controls (Table I). However, serum ALP values were statistically significant when compared to controls only in adenocarcinoma, adenosquamous carcinoma and pulmonary tuberculosis (Table I). Moreover, when each

120 Letter to the Editor

	NT	$Mean \pm SD$	
	No. of cases	King Armstrong units/100 ml	
Normal controls	15	10.2±1.6	
Chronic obstructive pulmonary disease	20	11.5±8.8	
Squamous carcinoma	25	12.2 ± 4.5	
Adenocarcinoma	15	28.1±17.8*	
Adenosquamous carcinoma	10	13.4±13.8**	
Pulmonary tuberculosis	20	19.4±9.3**	
Bronchial asthma	20	11.6±2.1	

TABLE I: Serum ALP level in healthy normal controls and patients with different lung pathology.

*P≤0.001; **P≤0.01

patients ALP values was segregated and categorised individually based on range of different serum ALP activity it was observed that some patients had enzyme values in the normal control range (Table II). This observation casts doubt on the utility of serum ALP as a marker for diagnosis of lung diseases.

Indian J Physiol Pharmacol 2001; 45(1)

Our study indicates that in some patients with confirmed lung pathology serum ALP values can still be found in the normal control range. The reason for this observation can be explained on the basis of information published that ALP in the lung is only found in plasma membrane and lemellar bodies of Type II alveolar epithelial cells (12, 13). Based on this information it would be logical to assume that an increase in serum ALP values above the normal control value could occur only in cases where these cells are involved in the disease process in the lung. Consequently it follows that if the lung pathology originates and develops in some other area of the lung devoid of Type II alveolar cells, there is no reason to expect perturbation in serum ALP level.

Further, we observed that the mean serum ALP values was higher in patients with lung disease when compared to mean normal control values. This observation was similar to that reported by others for status asthmaticus (14), adenocarcinoma of the lung

 TABLE II: Segregation and categorization of patients on the basis of their individual serum ALP values.

Range of ALP in KA units1%	0-12	13-24	25-36	37-48	49-60
	Percentage of cases				
Chronic obstructive pulmonary disease	70	30	-	-	-
Squamous carcinoma	64	32	4	1	-
Adenocarcinoma	20	20	26.7	26.7	6.6
Adenosquamous carcinoma	40	60	-	-	
Pulmonary tuberculosis	25	55	15	5	-
Bronchial asthma	70	30	Storig and	m hatel	-

KA units = King Armstrong units

Indian J Physiol Pharmacol 2001; 45(1)

(15) and active tuberculosis (16). However, the above observation does not in anyway authenticate the validity of serum ALP as a diagnostic marker for lung disease in the light of what we have discussed earlier. Moreover, it was also observed that in certain cases of active lung disease when Letter to the Editor 121

there was elevation of serum ALP, these values returned to normal levels consequent to complete cure of the lung pathology (data not shown). This observation validates the suggestion that serum ALP levels in lung diseases has more prognostic relevance than any diagnostic utility.

MALATHI M. AND SHRINIVAS B. RAO*

Departments of Biochemistry, Father Muller's Medical College, Mangalore – 575 002

and

*Kasturba Medical College, Mangalore – 575 001

REFERENCES

- Moss DW. Pespectives in alkaline phosphatase research. Clin Chem 1992; 12: 2486-2492.
- Kryszewski AJ, Neale G, Whitfield JB, Moss DW. Enzyme changes in experimental biliary obstruction. *Clin Chim Acta* 1973; 47: 175-182.
- Fishman WH, Inglis NI, Krant MJ. Serum alkaline phosphatase of intestinal origin in patients with cancer and cirrhosis of liver. *Clin Chim Acta* 1965; 12: 298-303.
- Price CP, Sammons HG. The nature of serum alkaline phosphatases in liver diseases. J Clin Pathol 1974; 27: 392-398.
- Whitaker KB, Whitby LG, Moss DW. Activities of bone and liver alkaline phosphatase in serum in health and disease. *Clin Chim Acta* 1977; 80: 209-220.
- Whitaker KB, Eckland D, Hodgson HJF, Saverymuttu S, Williams G, Moss DW. A Variant alkaline phosphatase in renal cell carcinoma. *Clin Chem* 1982; 28: 374-377.
- Capelli A, Lusuardi M, Cerutti G, Donner GF. Lung alkaline phosphatase as a marker of fibrosis in chronic interstitial lung disorders. Am J Resp Crit Care Med 1997; 155: 249-253.
- Wells All, DU Bois RM. Prediction of disease progression in idiopathic pulmonary fibrosis. *Eur Respir J* 1994; 7: 637-639.

 Hyde DM, King TE, McDermott T, Waldron JA, Colby TV, Thurlbeck WM, Flint A, Ackerson L, Chermack RM. Idiopathic pulmonary fibrosis: quantitative assessment of lung pathology. Am Rev Respir Dis 1992; 146: 1042-1047.

- Varley H. Calcium, phosphrous and phosphatases in "Practical Clinical Biochemistry" 4th ed., CBS Publ. Delhi 1988; 431-467.
- Snedecor GW. Statistical methods, Ames Iova State College Press 1948; 182-184.
- Meban C. The structural localization of alkaline phosphatase in the granular pneumocytes of hamster lung. *Histochemistry* 1975; 43: 367-372.
- Fisher AB, Furia L. Isolation and metabolism of granulate pneumocytes from rat lungs. Lung 1977; 154: 155-165.
- 14. Oosaki K, Mizushima Y, Hoshinok T. A case of bronchial asthma whose disease activity was associated with changes in serum ALP linked immunoglobulin levels. Kyobu Shikkan Gakki Zasshi 1992; 8: 1579-1582.
- Gault C. Meakins M. Eelin GH. Serum enzymes in patients with carcinoma of lung. Can Med Assoc J 1967; 96: 87-93.
- Gupta N, Garg UC, Ganguly NK. Enzyme levels in bronchoalveolar lavage fluid and serum of active pulmonary tuberculosis patients. *Enzyme* 1989; 41: 108-111.

*Corresponding Author