Coexistent Extra Adrenal Pheochromocytoma and Extrapulmonary Tuberculosis: A Case Report and Review of Literature

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ABSTRACT
Tuberculosis is a major cause of morbidity and mortality in the developing countries, accounting for two billion infected cases and 1.7 million deaths which were attributed to it in 2006, as per the WHO. Cancer was the leading cause of death, next only to coronary artery disease, with 11.3 million new cases and 7.9 million deaths in 2007. The combination of these two deadly diseases poses significant diagnostic and therapeutic challenges to the clinicians and the pathologists. The relationship of tuberculosis and malignancy can be viewed in two ways: tuberculosis can be a risk factor for cancer or tuberculosis and malignancy can coexist in the same biopsy or in the same patient. An early and an accurate diagnosis is a must for planning the appropriate management. We are describing a very rare case of coexistent extra adrenal pheochromocytoma and extrapulmonary tuberculosis in a single patient, which was diagnosed by fine needle aspiration cytology, with a brief review of the literature.

INTRODUCTION
Pheochromocytoma is a rare tumour which arises from the chromaffin tissue in the adrenal medulla. Extra-adrenal pheochromocytomas (EAPs) account for 10% of the cases, which present as thoracic or retroperitoneal masses in paravertebral locations [1,2]. It is extremely desirable for a clinician to make an accurate preoperative diagnosis, to prevent any untoward reaction or a fatality at surgery. Pharmacological tests can be used in the cases when they occur in the adrenal gland with a classic history of a paroxysmal attack of hypertension and headache [3]. Though it has been traditionally contraindicated, fine needle aspiration cytology (FNAC) has been used safely in cases of EAP for a preoperative diagnosis, especially in the unsuspected cases [4,5].

Very rarely, pheochromocytoma occurs concurrently with tuberculosis. A clinical overlap between these two entities may mislead the clinician, resulting in a delayed diagnosis [4]. Tuberculosis (TB) may retard the response to the treatment of pheochromocytoma. Likewise, the institution of chemotherapy may lead to the worsening of the underlying TB. The awareness on the coexistence of these two conditions is highly significant, since the treatment of both needs to be rationalized.

CASE REPORT
A 55 year female presented with a mass per abdomen of three months duration, with a history of fever, tiredness and palpitation. She was detected to have hypertension six months back and was on antihypertensive drugs since then. On general examination, her pulse was found to be 102 beats/minute and her blood pressure was 200/100 mm of Hg. She had marked pallor and a palpable, single, 2 x 2 cm posterior cervical lymph node. A mass which was present in the right lumbar region was palpable. The laboratory investigations revealed that the haemoglobin was 3 gm/dl and that the ESR was 30 mm at end of one hour. The serum electrolytes and the kidney and the liver function tests were normal. Ultrasonography showed a heterogeneous mass lesion in the right lumbar region.

A computed tomography (CT) scan showed a well-defined hyper dense mass lesion which measured 4 x 4 cm in the right lumbar region, above the bifurcation of the aorta. [Table/Fig-1].

An informed written consent was obtained from the patient after explaining to her, the dangers of performing the FNAC in that setting, in her language. An FNAC was done under ultrasound guidance by using a 21 gauge needle. The slides were fixed in 90% alcohol for hematoxylin and eosin staining and they were air dried.
was made. The urinary catecholamine level was 16 ng/dl. The cyto-
tology of the cervical lymph nodes showed the features of tuber-
culous lymphadenitis [Table/Fig-4]. Acid fast staining was positive for 
the tubercle bacilli. The culture of the FNAC specimen showed that 
the morphology of the isolates resembled that of Mycobacterium 
tuberculosis. Chest X ray did not show any evidence of pulmonary 
tuberculosis. The sputum for acid fast bacilli test as well as the 
culture was negative. After the control of the hypertension, the sur-
gical excision of the tumour was done, which confirmed the diag-
nosis of benign pheochromocytoma on histopathology. The patient 
was started on antituberculous therapy, category 1, of the Revised 
National Tuberculosis Control Programme. The patient is on regular 
followup and is doing well after 6 months.

**DISCUSSION**

Pheochromocytoma is a rare tumour which accounts for 0.1% of 
the hypertensive cases [1]. EAP accounts for 10% of the pheo-
chromocytoma cases and it presents as soft tissue mass lesion 
which runs parallel along the entire length of the abdominal aorta 
[2]. The most common sites for EAP include the paraadrenal area, 
followed by the organ of Zuckerkandel at the aortic bifurcation, 
the chest, the inferior mesenteric artery, over left common iliac 
artery, the bladder wall, and the celiac plexus, the superior mes-
enteric artery, the renal vessels, over right iliac artery, beneath the 
diaphragm and around the left ureter, in the decreasing order of 
frequency [6].

The patients are usually adults who present with the early and the 
classic symptoms of headache, palpitation and sweating. It is more 
likely for the clinician to elicit a history of an attack of hypertension 
than to observe it. At times, even the history is not available, espe-
cially in cases of a sustained hypertension [3]. An abdominal mass 
and pain, which are commonly seen in the late stages, however, 
may at times be the first presentations and in such cases, usually, 
the diagnosis is delayed by up to three years [7].

Traditionally, FNAC has been contraindicated in the diagnosis of 
pheochromocytoma due to the untoward reactions and its fatal-
ity. However, when it is used in the unsuspected cases of EAP 
which present as abdominal masses, no complication is noted 
[4,5]. The cytological diagnosis of pheochromocytoma has certain 
limitations. Gong et.al., in a report on two cases of EAP which 
were diagnosed by FNAC, while describing the pitfalls, have elo-
quently elucidated the clues for the correct diagnosis at cytology 
[4]. A microacinar formation may at times be misinterpreted as 
an adenocarcinoma. The dense eosinophilic cytoplasm may be 
suggestive of a squamous differentiation. However, like any oth-
er endocrine neoplasm, a marked pleomorphism and prominent 
nucleoli, binucleation, multinucleation and an intracytoplasmic and 
an intranuclear inclusion with a background which shows the pink 
eosinophilic Nissl's substance may aid in arriving at a correct diag-
nosis [4,5].

The different modalities for the diagnosis of pheochromocytoma in-
clude pharmacological tests, radiology which includes ultrasound, 
CT, metaiodobenzylguanidine (MIBG) scan, FNAC and biochemical 
tests. CT scan is useful for localizing and assessing the extension of 
the tumour. Urine catecholamines and their metabolites are sen-
tive tests both in the diagnosis and the follow up of pheochromocy-
tomas [8].

Bayle, in 1810, first stated that the coexistence of tuberculosis and 
malignancy in the lung was definitely occurring more commonly than it would occur by chance [9]. Since then, the association of
tuberculosis and malignancy has been investigated extensively. In a large review which was written by Falgas et al., the authors have put forth three different types of associations between TB and malignancy: i) the development of cancer on the background of a previous tuberculosis infection, ii) the concurrent existence of TB and malignancy in the same patient(s) or clinical specimen(s), and iii) the diagnostic challenges which arise from the multifaceted presentations of these two disorders [10].

The common malignancies which coexist with TB include lung cancer, haematological malignancies which include non-Hodgkin's and Hodgkin's lymphomas, chronic myeloid leukaemia, breast cancer, gastrointestinal tract malignancies, endometrial carcinomas, fallopian tube carcinomas and rarely, thymic carcinomas [10].

There are only two case reports on the concomitant occurrence of TB and pheochromocytoma. In one case which was reviewed by Grace and Walter, an intrathoracic pheochromocytoma was found to be associated with pulmonary TB [3]. In another case which was reported by Toure et al., a multifocal extrapulmonary TB was associated with an adrenal pheochromocytoma [11]. As has been depicted in the above cases, a clinical overlap and the misleading manifestations of the two diseases could render the diagnosis challenging and either could be missed.

Though many speculations have been made regarding the concurrent occurrence of TB in malignancy, the exact mechanism has yet to be elucidated. Much weight has been given to the postulate that the immunosuppression which is caused by a local or a systemic effect of malignancy might reactivate the preexisting tuberculous lesion or that it may result in the acquisition of a new primary infection [10].

The diagnosis of a coexisting pheochromocytoma and TB is not only of academic interest, but it has several clinical implications. The pheochromocytoma may hinder the treatment of the tuberculosis. In the case which was reported by Grace and Walter, the patient responded to the antituberculous therapy only after the surgical removal of the pheochromocytoma [3].

The presence of a palpable lymph node which is caused by tuberculosis may lead to an over staging in the TNM system [12]. The symptoms of TB may be mistakenly attributed by the treating clinician to a complication of the chemotherapy or to a worsening of the underlying disease.

CONCLUSION:

The clinician and the pathologist must be aware that pheochromocytoma and tuberculosis may coexist and that this may pose a diagnostic challenge due to the overlapping clinical manifestation, especially in a country with a high prevalence of TB.

FNAC is an easy, safe and a cost effective test in the diagnosis of pheochromocytoma, in a well-equipped hospital, as this will help in the management of the hypertensive crisis if there is any. A high degree of suspicion, careful attention to the cytology and the appropriate biochemical markers aid in making a correct diagnosis.

REFERENCES


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FINANCIAL OR OTHER COMPETING INTERESTS:
None.