Pathological variations of lung masses

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Abstract
Introduction: Lung is a common place for the spread of tumors from within or other parts of the body. Lung shadows with a diameter less than or larger than 3 cm are termed lung nodule or mass respectively.

Aims: To study the histopathological spectrum of lung masses and efficacy of computerized tomography guided FNAC/biopsy in diagnosis of lung masses.

Materials and Methods: Forty (40) cases picked up as lung masses on conventional screening & CT scan were subjected to CT guided FNAC/Biopsy and their pathological variations were studied.

Results: A total of 40 cases were studied. The mean age of the patients was 60.57 years with 25 (62.5%) males and 15 (37.5%) females. 65% of lesions were in right lung. Malignant lesions were more common and accounted for 27 (67.5 %) cases and benign in 8 (20%) cases. Amongst the malignant lesions, adenocarcinoma was seen in 16 (59.25%), squamous cell carcinoma in 10 (37.0%) and small cell carcinoma in 1 (3.70%) case.

Conclusion: FNAC is a simple, relatively safe, rapid, reliable technique for the diagnosis of pulmonary mass lesions, particularly with the aid of computed tomography (CT) scan. FNAC not only distinguishes between benign and malignant lesions but also helps in tumor typing of lung cancer.

Keywords: Lung mass, Computed tomography, Needle biopsy.

Introduction
Lungs, the most vital organs are exposed to both external and internal environment of the human body. It is thus prone to react to any adverse effects of the aetiological factors which may be infective, inflammatory, parasitic, vascular, neoplastic and other causes.

The detection & diagnosis of any lung lesion usually begins with a chest radiograph, either in a symptomatic patient or a patient undergoing chest radiograph for an unrelated reason. Lung cancer was not even recognized as a distinct disease until 1761. Different aspects of lung cancer were described further in 1810. Malignant lung lesions made up only 1% of all cancers seen at autopsy in 1878, but had risen to 10-15% by the early 1900.¹ Worldwide in 2012, lung cancer occurred in 1.8 million people and resulted in 1.6 million deaths representing 19.4% of all deaths from cancer.² The American cancer society’s estimate for lung cancer in the United states for 2018 are: about 234,030 new cases (121,680 in men & 112,350 in women), and about 154,050 deaths (83,550 in men 70,550 in women). Overall, the chance that a man will develop lung cancer in his life time is about 1 in 15; & for a woman the risk is about 1 in 17.³

CT of the chest introduced in the late 1970’s, today remains the most widely used imaging test to stage patients with lung cancer. Definitive diagnosis however relies on the histopathological diagnosis of the tissue obtained by a CT guided FNAC, core biopsy or any other mode of biopsy.

A study of wide variety of pathology of lung masses based on cytology/histopathological diagnosis was undertaken at a tertiary care hospital.

Materials and Methods
A cross-sectional study of 40 patients who had reported with a suspicious pulmonary lesion on a chest radiograph was done at a tertiary care centre over a period of two years. These patients subsequently underwent CT scan imaging and CT guided FNAC/Core needle biopsy to arrive at a final diagnosis.

FNAC/Core Needle Biopsy: Informed consent of the patients was taken for CT guided FNAC/biopsy of the lung mass. The puncture entry point was infiltrated with 2% lignocaine. FNAC was performed using 22 G / 23 G spinal needle on 10 patients, CORE needle biopsy using a 18 G automated core-biopsy needle on 29 patients and on 1 patient both FNAC and CORE needle biopsy had to be done for want of adequate and representative tissue. Air-dried smears were stained with May-Grünwald-Giemsa stain whereas alcohol-fixed smears were stained with Papanicolaou stains for rapid cytopathological evaluation of the lesions. Core biopsy tissue was sent in (10%) formalin solution to pathological lab and sections were stained by haematoxylin and eosin (H&E) stain.

A follow-up CT scan was done on every patient immediately after the procedure to rule out pneumothorax or any haemorrhage.
Observations and Results

The study was undertaken with the aim of studying the pathological variations of lesions presenting as suspicious lung masses. The computed tomographic observations of the lung masses were made with respect to location, size, margin, any associated calcification, cavitation, pleural effusion, mediastinal lymphadenopathy, character of contrast enhancement and bony involvement.

General Considerations

**Age & Sex:** The mean age of the patients was 60.57 years with the standard deviation of 12.2284. There were 25 (62.5%) males and 15 (37.5%) females.

Majority of the lesions 26/40 (65%) were in the right lung and 14/40 (35%) in the left lung. Their size ranged from 3 to 10 cm with largest group comprising of 17 (43%) having lung masses of the size 5-7 cm. Fig. 1.

![Distribution of lesions based on size](image)

Fig. 1: Distribution of lesions based on size

**Table 1: Distribution of benign lesions**

| S. No | Histopathological diagnosis          | Number of cases | Percentage (%) |
|-------|--------------------------------------|-----------------|----------------|
| 1.    | Granulomatous lesion lung             | 02              | 25%            |
| 2.    | Wegener's granulomatosis              | 01              | 12.5%          |
| 3.    | Mucormycosis                         | 01              | 12.5%          |
| 4.    | Pneumonitis                           | 01              | 12.5%          |
| 5.    | Pulmonary infarct                     | 01              | 12.5%          |
| 6.    | Pulmonary hamartoma                   | 01              | 12.5%          |
| 7.    | Fibrocollagenous tissue               | 01              | 12.5%          |
| **Total** |                                       | **08**         | **100%**       |

70 year old male, chronic smoker presented with anorexia and weight loss since 2 months

![Distribution of cases based on type of malignant lung lesions](image)

Fig. 2: Cyto-pathological diagnosis of malignant lesions

The distribution of benign lesions is shown in table 1.

Based on radiological characteristics of the lesions alone, 35 (87.5%) were provisionally diagnosed malignant and 5 (12.5%) as benign.

The final cyto-pathological diagnosis was however malignant in 27 (67.5%) cases and benign in 8 (20%) cases. The remaining 5 (12.5%) were inconclusive due to inadequate sampling.

Amongst the 27 (67.5%), malignant lesions, adenocarcinoma was seen in 16 (59.25%), of which mucin secreting adenocarcinoma is depicted in Fig. 4A, 4B and 4C. Squamous cell carcinoma was seen in 10 (37.03%) patients. Poorly differentiated squamous cell carcinoma is as shown in Fig. 3A, 3B and 3C. Small cell carcinoma was seen in 1 (3.70%) case. (Fig. 2).

![Fig. 3A: Chest XRay - Lobulated peripherally based mass lesion right middle zone. Associated incidentally old united fractures of right 4th and 9th ribs is seen posteriorly](image)
Fig 3B: CECT Chest mediastinal window axial section - Spiculated mass lesion apical segment of right upper lobe with irregular areas of breakdown within, suggestive of carcinoma right lung; C): CECT Chest lung window sagittal section – Spiculated mass lesion is seen in apical segment of right upper lobe

Fig 3D: Biopsy/7434/13 - HPE shows malignant cells with high N:C ratio with abundant cytoplasm. Opinion:- Non small cell carcinoma- poorly differentiated squamous cell carcinoma.

45 year old female, chronic smoker, presented with right sided chest pain and hemoptysis.

Fig 4 A&B: Chest XRay - Mass lesion right upper zone. Fig. 14B: CT guided biopsy - Multilobulated mass lesion with biopsy needle tip seen in the lesion in posterior segment of right upper lobe, suggestive of carcinoma right lung. Right sided pleural effusion is also noted.
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Fig. 4C: Biopsy/232/14 - HPE shows malignant epithelial cells arranged in glands with abundant pools of mucin amidst fibrocollagen

Opinion: Mucin secreting adenocarcinoma

In our study, cytology and HPE reports on FNAC/Core biopsy were considered as final confirmatory report. From a total of 35 cases in which adequate sampling was received; 30 were considered positive and 5 were considered negative for malignancy radiologically. False positive cases for malignancy based on radiology were 04 and false negative was 01. Amongst the false positive cases, 01 was found to be pneumonia as biopsy showed acute inflammatory cells with necrotic material, 02 cases were detected to be granulomatous lesions, (possibly of tubercular origin) and 01 was diagnosed as granulomatous polyangitis (Wegener’s granulomatosis), as confirmed by HPE and immunohistochemistry (IHC).

The one (01) false negative case suspected to be fungal pneumonia radiologically turned out to be adenocarcinoma.

Chi square value (with Yates correction) was 7.35, and p value 0.0066972 was statistically significant.

Complications: Pneumothorax occurred in 1 (2.5%) case. It was minimal and got resolved without any treatment. Mild pulmonary haemorrhage occurred in 1 (2.5%) case and 3 (7.5%) cases complained of chest pain which got settled after one day.

Discussion

Diagnosis of lung mass lesions has always remained a challenging task. Any shadow more than 3 cm in size on chest X-ray or CT scan is suspected of malignancy unless proved otherwise. However, many benign disease processes such as infections, granulomatous disease, and cryogenic organizing pneumonia may also present as a lung mass. Certain features can be depicted clearly with CT scan, which helps to suggest whether the lesion is benign or malignant. Transthoracic needle biopsy (FNAC/Core biopsy) using image guidance with CT has great diagnostic value for the definite characterization of lung lesions.

Numerous authors have expressed that the sensitivity and diagnostic accuracy is somehow slightly more in big lesions (>3cm) and in lesions located peripherally. Studies have also reported that cytological differentiation could be made properly in small cell and non-small cell lung cancers and there is good overlap in cytology and histopathological reports. Several factors can lead to non-specific biopsy results. The most common factors are tumor necrosis, inappropriate biopsy and insufficient sampling.

The most common complication of biopsy is pneumothorax. Kazerooni et al. showed that an increased lesion depth and small lesion size were strongly correlated with pneumothorax development. In our series, the parenchyma depth traversed for sampling was deeper in cases that developed pneumothorax.

In our study of 40 cases, on the basis of CT scan findings alone, a provisional radiological diagnosis about the nature of the lesion was made as malignant in 35 (87.5%) and benign in 5 (12.5%) cases. Histopathologically, out of 35, however 27 (76.5%) proved to be malignant and 8 (20%) cases were benign in nature. In 5 cases (12.5 %) results remained inconclusive due to insufficient sampling. Adenocarcinoma was the commonest comprising 16 (59.25%) cases, squamous cell carcinoma 10 (37.03%) cases, and small cell carcinoma only 1 (3.70%) case. Benign lesions were 8(20%) with their diagnosis as granulomatous lesion (2 cases), and Wegener’s granulomatosis, mucormycosis, pneumonitis, pulmonary infarct, pulmonary hamartoma, fibrocollagenous tissue (1 case) each. We thus found a good overlap between the CT scan diagnosis and the final diagnosis as confirmed on cytology/histopathology. The most common age group was 51-70yrs with M:F ratio as 1.6:1.
In a similar study of forty cases by Manas et al.,
the most common age group was 41-70yrs with male
preponderance in the ratio of M:F as 5:1. The
predominant lesion found in the study was malignancy
in 24 (60%); followed by inflammatory lesions in 12
(30%). One case each was reported as suspicious of
malignancy (2.5%) and granulomatous (2.5%), the
material was inadequate for interpretation in two cases
(5%). Out of 24 FNAC proven cases of malignancy,
adenoacarcinoma was found in 12 (30%) followed by
squamous cell carcinoma in 9 (22.5%) cases and undifferentiated carcinoma in 3 (7.5%) cases. The
diagnostic accuracy was 95% considering cytological
criteria as the standard.

Bahadir T et al. in their study of 134 cases found
the lesion at the mean age of 58.9 yrs. Maximum
lesions were in the upper zone of right lung. The final
diagnosis of FNAC were malignant in 107 (80%) cases,
benign in 5 (4%) & 22 (16%) were non-specific. The
specificity for diagnosing malignancy was 100%,
adenoacarcinoma was seen in 27 (25%) cases, squamous
cell carcinoma in 20 (19%) cases.

Mukherjee, S et al. in their study of 94 cases found
malignancy in 90(%) cases. The most common
age group was 40-70 yrs with male preponderance
(85%). There was no false positive for malignancy.
Hence there was 97.7% sensitivity and 100% specificity
of CT guided FNAC as a diagnostic procedure. Over all
accuracy for cytological typing was available in
88(95%) cases. Non-small cell carcinoma predominated, similar to the experience of other
workers.

Tan et al. found the incidence of adenoacarcinoma
significantly higher than that of squamous cell
carcinoma.

Jayashankar E et al. in their study of sixty cases
found that majority of cases were in 5th and 6th decade
and there was male preponderance. Provisional
diagnosis based on radiologic findings was: 48
malignant and 12 cases benign. The cytological
diagnosis however revealed 41 malignant and 19
benign. The diagnostic accuracy of CT guided FNAC
was more than 90%. The percentage distribution
of different types of bronchogenic carcinomas was
squamous cell carcinoma (51%), adenoacarcinoma
(43%), small cell carcinoma (3%) and poorly
differentiated carcinoma(3%).

Mini Gangopadhyay et al. in their study of 127
cases found FNAC sensitivity as 96% and 100%
specificity in diagnosing malignant lung tumours.
Among the benign lesions, specific diagnosis was
obtained in 48.1% cases. The males were 76(59.8%)
and females 51(40.15%). The mean age was 58.71 yrs.
27 cases(21.2%) were benign, 96 cases(75.6%)
malignant, 2 cases were suspected of malignancy and
in another 2, the material was inadequate for
cytopathological opinion. The most common carcinoma
diagnosed by cytology was adenoacarcinoma (54.2%),
followed by squamous cell carcinoma (25%) and small
cell carcinoma (8.3%). In the benign lesions, 12 cases
yielded tubercular granulomas, 12 cases as chronic non-
specific inflammation, 1 case of fungal infection
(Aspergillus fumigatus) and 2 cases were of lung
abscess.

Mondal SK, et al. in their prospective study of
130 cases analysed the radio-pathological correlation
and found that that the peak incidence was in the fifth
and sixth decade. The benign lesions were 10(8.07%) and
malignant lesions were 114(91.93%) as shown by
cytology. The most common tumour was
adenocarcinoma (51.72%) followed by squamous cell
 carcinoma (6.89%). Diagnostic accuracy of CT guided
FNAC was 95%. Post procedural complications like
haemorrhage and chest pain were noted in only 3 cases.

Haresh chand et al. in their series of retrospective
study of 119 malignant cases found squamous cell
carcinoma in 55(46.2%), adenoacarcinoma in 18
(15.1%), Small cell carcinoma in 18(15.1%), Poorly
differentiated carcinoma in 12(10.1%), Others 4(3.4%) and
in 12(10.1%), the metastatic carcinoma. Male to
female ratio was almost 1:1. The mean age group was
54.46 yrs. Upper zone of the right lung was the most
common area involved in all the series. Various studies
cite the incidence of malignant tumours on lung FNAC
as 58-88%. 

Conclusion

The prevalence of lung mass is an extremely
common finding that most radiologists will encounter
on a frequent basis. Familiarity with the various
manifestations of lung mass on chest radiography may
help suggest the initial diagnosis. Once a suspicious
abnormality is detected, CT guided FNAC/Core biopsy
is the next step in the diagnostic work-up to arrive at
a definitive diagnosis.

Literature reveals that 70% of the primary lung
cancers are diagnosed as non-small cell carcinomas,
whereas small cell carcinomas are rare and comprise
around 20% of all primary lung tumours. In our study,
the same pattern was observed. In the non-small cell
group of carcinomas, adenoacarcinoma remains the
commonest tumor. Though squamous cell carcinoma
used to be the commonest, studies indicate that
adenocarcinomas may have overtaken it in incidence.

FNAC which is not only a safe and rapid technique
obviates the need for invasive techniques like
mediastinoscopy, video-assisted thoracoscopy, and
diagnostic thoracotomy in the diagnosis of pulmonary
lesions. More so, initiation of specific therapy like
chemotherapy or surgery is possible without
unnecessary delay. When lung cancer is found at an
early stage, the chance of cure is much higher than at
later stages of the disease.

Conflicts of Interest: The author had none to declare.
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