A cytological comparison of efficacy of bronchoalveolar lavage, transbronchial needle aspiration and image guided fine needle aspiration for diagnosis of pulmonary lesions in a rural tertiary care centre

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Abstract
Introduction: Pulmonary lesions include a wide variety of benign and malignant conditions of lung, pleura, mediastinum and vertebrae. Lung cancer is a frequently diagnosed cancer worldwide. Although clinical data, location and radiological findings can narrow down the diagnostic possibilities, a cytological diagnosis is warranted before initiating the specific treatment for the malignant disease.

Aims and Objectives: The present study was undertaken with the aim to compare the sensitivity and specificity of bronchoalveolar lavage (BAL), transbronchial needle aspiration cytology (TBNA) and image-guided fine needle aspiration cytology (FNAC) in the evaluation of patients with pulmonary lesions.

Materials and Methods: The present study was conducted on 100 patients with pulmonary lesions coming to the department of pulmonary medicine at MMIMSR, Mullana. For diagnostic purposes various procedures like BAL, TBNA and image guided FNAC were performed. Cytological examination was carried out in the department of Pathology.

Results: Among 100 cases, TBNA was done in 63 cases, BAL in 52 cases and image guided FNAC was performed in 37 cases. Using these cytological techniques, a diagnosis was obtained in 81% cases. The sensitivity and specificity of each technique was assessed. On cytological evaluation, 12% cases had a non-neoplastic diagnosis, whereas 69% cases, malignancy was diagnosed. However, in 19% cases no diagnosis was offered. The most common diagnosis amongst malignant lesions was non-small cell carcinomas. (Squamous cell carcinoma- 28%, Adenocarcinoma-23%) followed by small cell carcinoma (8%).

Conclusion: Image-guided FNAC of pulmonary lesions is a simple, safe, economically viable technique with low morbidity and offers quick and early diagnosis. Our study revealed that TBNA and FNAC were comparatively better in the diagnosis when compared to BAL.

Keywords: Image-guided Fine Needle Aspiration Cytology, Pulmonary Lesions, Squamous Cell Carcinoma.

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Introduction
Lung cancer is a frequently diagnosed cancer in the world and a common cause of death related to cancer. Lung cancer comprises 17% of the total newly diagnosed cancer cases in males and causes 23% of total cancer related deaths. The prognosis of lung cancer depends on the stage of the cancer at the initial diagnosis. Therefore, an early diagnosis of lung cancer improves the prognosis of lung cancer. Different modalities used for early diagnosis are radiological investigations, TBNA, bronchial biopsy, exfoliative cytology like bronchial washings, BAL and image guided FNAC.

Image-guided FNAC is well established diagnostic tool for diagnosis of radiological apparent lesions of the lung. The procedure has a low complication rate and is highly sensitive for the diagnosis of non-palpable deep-seated lung lesions. TBNA of the mediastinum is rapidly becoming a technique of choice for minimally invasive staging of lung cancer. BAL via a flexible bronchoscope was introduced by Reynolds and Newball in 1974 as a therapeutic procedure for clearing secretions in the alveolar spaces. Subsequently, the technique has been used as diagnostic procedure for confirmation of pulmonary infections particularly in immunocompromised individuals and interstitial lung diseases, thereby replacing the need of open biopsy.

These cytological procedures are easily performed and well tolerated for routine assessment of the patients with lung lesions. While correlating these cytological techniques false negative and false positive results can be reduced, thereby, improving the diagnostic efficacy. These cytological procedures have many advantages but more work requires to be done. The present study is undertaken with the aim to study the efficacy of BAL, TBNA and image
Materials and Method

The present study was conducted on 100 patients with pulmonary lesions coming to the department of respiratory medicine, MMIMSR at Mullana over the time period June 2014 to December 2016. Ethical clearance for the study was obtained from institute ethical committee. Informed consent was obtained from all the patients after briefly explaining the technique. The patient’s requisition forms including patient’s age, sex, smoking history and radiological investigations were filled by clinicians. For diagnostic purposes, procedures like BAL and TBNA were performed in the department of pulmonary medicine. Image-guided FNAC were done in the department of radio-diagnosis, with the help of lumbar puncture needle No. 20-22, attached to 20 ml disposable syringe with Franzen handle through percutaneous and transthoracic approach. The collected material was smeared on the slides. Half of slides were wet fixed whereas the rest of the slides were air-dried and sent to the department of pathology. Wet fixed slides were stained with hematoxylin & eosin stain and papanicolaou (PAP) stains, while air-dried smears were stained with Leishman-Giemsa. Final diagnosis was made with the help of bronchoscopic findings, clinico-radiologic impressions, cytological findings and response to treatment.

Observations and Results

Hundred cases were included in the present study. Among these 100 cases, both BAL and TBNA were performed in 52 cases whereas in 11 cases TBNA alone was done. Among 37 cases of image-guided FNAC, 33 cases were ultrasound-guided and 4 cases were computed tomography (CT) guided (Table1). Age of the patients was between 14 years to 80 years (mean 53.14 years). Male female ratio was 2.7:1 having a male preponderance. Majority of patients had history of smoking. In our study, the most common presenting complaint was cough (73%) followed by shortness of breath (63%) and chest pain (53%). The lung lesions were central in 63% and the peripheral in 37% cases.

Of the 100 cases, a firm diagnosis was offered in 81 cases using various cytological techniques. In 62 patients where TBNA was performed, diagnosis was possible in 46 cases whereas in 17 cases no diagnosis could be arrived at. The sensitivity and specificity of TBNA was 86.3% and 83.3%. BAL was done in 52 cases and correct diagnosis was achieved in 14 cases. The sensitivity and specificity of BAL procedure in our study was 34.2% and 90.9%, respectively. In cases of image-guided FNAC, the correct diagnosis was obtained in 35 cases. Thus, the sensitivity and specificity of image guided procedure was 97.2% and 100% respectively.

The lesions were categorized as non-neoplastic, neoplastic and non-diagnostic. In non-neoplastic category, there were 6 cases each of granulomatous inflammation (Fig. 1A and 1B) and non-specific inflammatory pathology. Of the six cases of granulomatous inflammation, two cases were Ziehl-Neelson positive and thus, these were labelled as tuberculosis. In the neoplastic category, no benign lesion was encountered in the study. Study showed 69 malignant cases when 8 cases were diagnosed as small cell carcinoma, 28 cases squamous cell carcinoma, 23 cases adenocarcinoma and 4 cases poorly differentiated carcinoma (Table 2).

![Photomicrograph of cytological smear showing epithelioid cell granuloma. (Leishman-Giemsa 400X)](image1)

![Photomicrograph revealing clusters of atypical squamous cells in background of keratinous debris. (Leishman-Giemsa 400X)](image2)
Fig. 3a, b: a- Photomicrograph of adenocarcinoma exhibiting gland formation. (Leishman-Giemsa 400X), b- Smear showing dispersed tumor cells of adenocarcinoma exhibiting nuclear pleomorphism with vague gland formation. (Hematoxylin & Eosin 400X)

Fig. 4a, b: a- Small cell carcinoma- tumor cells revealing nuclear moulding and nuclear streaking. (Leishman-Giemsa 400X), b- Photomicrograph revealing tumor cells with hyperchromatic nucleus exhibiting nuclear moulding. (Hematoxylin & Eosin 400X)

Table 1: Distribution of cases according to the procedure performed

<table>
<thead>
<tr>
<th>Type of procedure</th>
<th>No. of cases</th>
</tr>
</thead>
<tbody>
<tr>
<td>TBNA and BAL</td>
<td>52</td>
</tr>
<tr>
<td>TBNA</td>
<td>11</td>
</tr>
<tr>
<td>Image guided FNAC</td>
<td>37</td>
</tr>
<tr>
<td>Total</td>
<td>100</td>
</tr>
</tbody>
</table>

Table 2: Cytological diagnosis of 100 cases of Pulmonary lesions

<table>
<thead>
<tr>
<th>Cytologic diagnosis</th>
<th>No. of cases</th>
</tr>
</thead>
<tbody>
<tr>
<td>Squamous cell carcinoma</td>
<td>28</td>
</tr>
<tr>
<td>Adenocarcinoma</td>
<td>23</td>
</tr>
<tr>
<td>Small cell carcinoma</td>
<td>08</td>
</tr>
<tr>
<td>Granulomatous inflammation</td>
<td>06</td>
</tr>
<tr>
<td>Non- specific inflammatory pathology</td>
<td>06</td>
</tr>
<tr>
<td>Poorly differentiated carcinoma</td>
<td>04</td>
</tr>
<tr>
<td>Adenosquamous carcinoma</td>
<td>01</td>
</tr>
<tr>
<td>Large cell carcinoma</td>
<td>01</td>
</tr>
<tr>
<td>Adenoid cystic carcinoma</td>
<td>01</td>
</tr>
<tr>
<td>Hodgkins disease with pulmonary involvement</td>
<td>01</td>
</tr>
<tr>
<td>Carcinoid tumor</td>
<td>01</td>
</tr>
<tr>
<td>Anaplastic large cell lymphoma</td>
<td>01</td>
</tr>
<tr>
<td>Non-diagnostic</td>
<td>19</td>
</tr>
<tr>
<td>Total</td>
<td>100</td>
</tr>
</tbody>
</table>

Discussion

The cytological evaluation in diagnosis of pulmonary lesions has been considered as one of its best application. The present study is one of a kind, as here, an attempt has been made to compare the various techniques in efficiently diagnosing various lesions of the lung.

The present study was done on 100 cases of pulmonary lesions using various cytological techniques. The mean age of the patients under study was 53.14 years which is comparable to studies done by Sareen et al., Bhat et al and Rohtagi et al. There was male preponderance of pulmonary lesions as mentioned by majority of the studies of the literature. The male: female ratio of present study was 2.7:1. Bodh et al found in their study that male: female ratio is 3.35:1 whereas Sareen et al showed male: female ratio of 8.4:1. Cough and shortness of breath was the most common complaints. Similar findings were observed in the study by Sareen et al.

Three cytological techniques i.e. TBNA, BAL and image guided FNAC were performed for the evaluation of the lesions. Both TBNA and BAL were performed in 52 cases of radiologically suspected lung lesions. The sensitivity and specificity of BAL procedure was 34.2% and 90.9%, respectively. The sensitivity of our study correlated however, the specificity was slightly higher than the study done by Bhat et al. In present study, 14 cases were correctly diagnosed, one case was false positive and twenty seven cases were false negative. The main reason for false negative cases was due to non-representative material or paucicellular smears. The cytological sampling by BAL procedure depends on exfoliated cells from the lesion. Further, it is dependent on technical skill of physician who is performing the lavage, differentiation of malignant growth, as poorly differentiated lesions are less cohesive and cytological preservation of material is obtained. While the exfoliative cells lying in bronchial cavity undergo degenerative changes and loose their morphological details making them difficult to differentiate from reactive bronchial epithelial cells. False positive cases can result from misinterpretation of cellular changes in inflammatory conditions such as pneumonia of long duration (atypical
macrophages) and squamous metaplasia. In the present study, there was one false positive case due to misinterpretation of squamous metaplasia as suspicious of malignancy. As there were fewer number of false positive cases, this was the strength of BAL cytology in the present study.

TBNA was performed in 63 cases and the correct diagnosis was achieved in 44 cases. There were two false positive cases due to misinterpretation of squamous metaplasia as dysplastic squamous cells suggestive of malignancy. In a study conducted by Ye T et al.,

the sensitivity and specificity of TBNA was 95.08% and 100%, respectively which was slightly higher than our study. The relatively lower sensitivity and specificity of TBNA in our study was due to paucicellular smears which could be due to low technical expertise of the pulmonologist to obtain adequate representative sample from the lesion.

Current diagnosis was possible in 35 of the 37 cases of image guided FNAC. No false positive case was reported on the cytology, therefore, giving specificity as close to 100% and the sensitivity of 97.2%. In a study done by Sareen et al.,

the sensitivity and specificity of CT guided FNAC were found to be 87.25% and 100%, respectively which correlated well with our study. No complication was noted during or after the procedure in any of the cases. There was one false negative case in present study which on repeated aspirations yielded necrotic material only. The diagnosis was later confirmed on biopsy as squamous cell carcinoma.

Squamous cell carcinoma (Fig. 2A and 2B; 28% cases) was the most common malignancy followed by adenocarcinoma (Fig. 3A and 3B; 23% cases) and small cell carcinoma (Fig. 4A and 4B; 8% cases). In western world, adenocarcinoma is the most common malignancy of the lung. However, most of the Indian studies mentions squamous cell carcinoma to be the most frequently occurring lung malignancy. In the present study, four cases were diagnosed as poorly differentiated carcinoma without further subtyping due to the presence few viable anaplastic cells in a necrotic background. The other less common malignancies diagnosed in our study were adenosquamous carcinoma (1 case), large cell carcinoma (1 case), adenoid cystic carcinoma (1 case), carcinoid (1 case), Hodgkin’s disease with pulmonary involvement (1 case) and anaplastic large cell lymphoma (1 case).

Thus, for pursuing diagnosis of pulmonary lesions especially lung cancer, there are number of cytological procedure to choose from. For central lesions, TBNA is the preferred technique whereas the peripheral lesions are best approached with image-guided FNAC. However, BAL is the safest procedure and by increasing the number of attempts while obtaining sample can increase its sensitivity and specificity especially in peripheral lung lesions.

Conclusion

Image-guided FNAC of pulmonary lesion is a simple, safe, economically prudent technique with low morbidity and leading to quick and early diagnosis. It was found in our study that TBNA and FNAC are comparatively more efficient than BAL even though BAL and TBNA can be done in one sitting.

References

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