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ORIGINAL STUDY

Comparison of fine needle aspiration and core needle biopsy in the diagnosis of bone tumours

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Much literature exists regarding the diagnostic yield and accuracy of core needle biopsy (CNB) and fine needle aspiration (FNA) but none compares both in the same tumour. Ninety-four patients were prospectively studied using a FNA and CNB. With FNA 70 diagnoses were possible (74,5%). Accurate diagnosis rate was 97,1%. In 92 patients (97,9%) a diagnosis was obtained with CNB and 91 (98,9%) were accurate.. The diagnostic yield was 74,5% for FNA and 97,9% for CNB (p < 0.0001). The diagnostic accuracy was 97,1% for FNA and 98,9% for CNB (p = 0.5787). Regarding determining malignancy FNA and CNB had 98,3% and 98,5% sensibility, 100% and 100% specificity, 100% and 100% positive predictive value and 95,2% and 96,2% negative predictive value, respectively. In conclusion FNA is as accurate as CNB on all accounts. Despite the reliability of FNA, the number of inconclusive cases makes it an inferior technique when compared with CNB.

Keywords : Bone neoplasms ; Diagnosis ; Biospy ; fineneedle ; core needle.

INTRODUCTION

With the exception of the "usually leave me alone lesions", biopsy is mandatory to diagnose bone tumours and related lesions. In this procedure the goal is to obtain the maximum representative sample causing a minimum morbidity and tumour spread.

Open biopsy is no longer the gold standard for the diagnosis of these lesions (15) and percutaneous

No benefits or funds were received in support of this study. The authors report no conflict of interests. biopsies – core needle biopsy (CNB) and fine needle aspiration (FNA) – have emerged as the primarily diagnostic modalities (5,7,10,15).

Accuracy of FNA has been described to be at least 85% and to be even higher in discriminating between benign and malignant lesions (4). Söderlud and colleagues (14) have shown that FNA accuracy could reach 99% when radiographic findings aid the cytological interpretation (concordance). In fact, radiographic analysis of bone lesions gives valuable information about the tumour matrix and the reaction of surrounding host tissues. These elements are absent in soft tissue lesions.

Traditionally, CBN is favoured over FNA because its accuracy is higher (4,11) but mainly because insufficient FNA samples are a significant problem, ranging from 4% to 33% (4).

This study is a prospective evaluation of 94 bone lesions in which FNA was performed followed by CNB in order to compare the accuracy, the scarcity of samples and the possibility to initiate the treatment with each method. To our knowledge no previous study has evaluated these 2 techniques in the same bone lesion.

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PATIENTS AND METHODS

Between January 2011 and January 2014 all patients with undiagnosed bone lesions were invited to participate in the study. Indications for biopsy were the presence of a bone lesion whose diagnosis was not obtainable by anamnesis, physical examination, laboratory and imaging studies; another group of patients were those with a history of malignancy elsewhere and in which a secondary bone lesion could not be excluded by the aforementioned methods – in this group of patients the objective was to exclude malignancy.

The average age of the patients was 53,5 years (12-86). There were 61 males and 33 females. All biopsies were performed under image guidance (64 with the use of CT-scan and 30 with radioscopy). Thirty lesions were localized to the lower limb, 15 to the upper limb, 23 to the spine, 22 to the pelvis and 4 to the trunk.

Patients were initially submitted to imageguided FNA. The most suitable route was chosen in order to avoid noble structures such as neurovascular bundles and organs. After the selection of the area, the whole path from the skin to the periosteum, was anesthetized with 3-5 ml of 2% Lidocaine. A 22-gauge needle was introduced and a cytoaspiration was performed (Fig. 1). Samples were kept in CytoRich®Red Preservative Fluid allowing further cytobiological studies. Immediately after the aspiration, within the same cutaneous region, an 8-gauge needle biopsy (ZamarCatchsystem®) was introduced 3-4 times in order to get a macroscopically sufficient sample which was immediately placed in a sterile flask containing formaldehyde.

In the majority of these cases a diagnosis of bone tumour was necessary to start treatment but in a few the exclusion of malignancy was also mandatory. All procedures were performed by the same orthopaedic surgeon and all samples were analysed by the same pathologist.

Results of both biopsy techniques were compared with final diagnoses which were established by surgical specimen (41 patients) or ulterior clinical and imaging evaluation (53 patients) since in some benign tumours, metastases and haematopoietic lesions no surgery is needed. With the exception

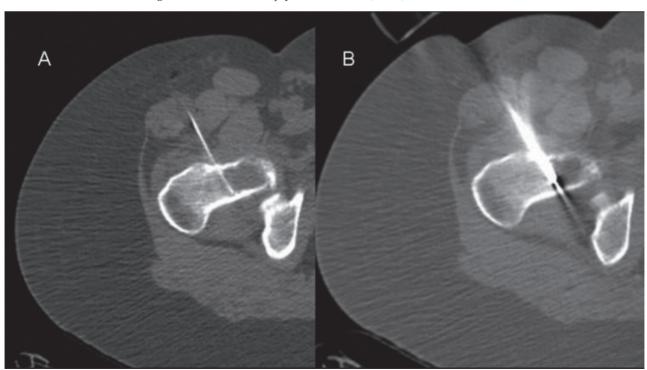


Fig. 1. - Percutaneous biopsy of a metastasis (breast). A - FNA; B- CNB

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of metastases, the minimum follow up to confirm these diagnoses was 2,5 years. Exclusion of malignancy or infection, when clinically suspected, was included in the group of diagnosis.

The diagnostic yield (ratio between the number of diagnosis obtained and the number of all procedures) and accuracy (ratio between the confirmed diagnosis and the number of established diagnosis) were determined for both procedures and compared using the t test for proportion, set to a 95% confidence interval. The proportion of patients in which FNA and CNB were able to exclude malignancy, establish diagnosis and initiate treatment were calculated for both techniques and compared using the Chi-Square test. Specificity, sensibility, positive predictive value (PPV) and negative predictive value (NPV) of both techniques were analysed using MedCalc version® 15.11.4. The MedCalc software was used for statistical analysis and a p value < 0.05 was considered to represent a significant difference between both techniques.

All procedures followed were in accordance with the ethical standards of the responsible committee on human experimentation and with the Helsinki Declaration of 1975, as revised in 2000.

RESULTS

There were no complications associated with the procedures and all patients were discharged on the same day, with the prescription of a simple analgesic.

Final diagnoses were: 29 metastases, 28 primitive malignant tumours, 13 benign tumours, 12 haematological diseases and 5 infections. In 7 cases pathology could be excluded. Results for all patients are summarised in table 1.

With FNA 70 diagnoses were possible (74,5%). Two of them were wrong - a spinal discitis was initially taken as a giant cell tumour and a low-grade chondrosarcoma of the scapula was assumed as an enchondroma (Table 1 – cases 9 and 32). Accurate diagnoses were then 97,1%. With this technique, 15 results (16%) were completely inconclusive but in 9 cases, although a diagnosis was not obtained, the pathologist could differentiate a benign lesion (n = 5) from a malignant one (n = 4) and this differentiation was correct in all cases. Excluding the inconclusive cases, and regarding determining malignancy, FNA had 98,3% sensibility, 100% specificity, 100% positive predictive value and 95,2% negative predictive value (Table 2).

In 92 patients (97,9%) a diagnosis was obtained with CNB. Of these, 91 (98,9%) were accurate. Only 1 benign lesion was misdiagnosed - a lowgrade chondrosarcoma of the proximal femur was assumed as an osteochondroma (Table 1 – case 19). Regarding determining malignancy CNB had 98,5% sensibility, 100% specificity, 100% positive predictive value and 96,2% negative predictive value (Table 2).

The diagnostic yield was significantly lower (p < .0001) with FNA than with CNB. There was no statistical difference (p=0.4046) between the diagnostic accuracy when using both techniques nor there were differences in the sensitivity, sensibility, PPV and NPV (Table 2). Comparing the possibility of exclude malignancy FNA and CNB were statistically similar (Table 3). However, FNA was inferior to CNB establishing an accurate diagnosis and initiating a treatment (Table 3).

DISCUSSION

All cytological and histological results should always be interpreted integrating clinical and imaging information. Percutaneous biopsy also depends on the operator technique and on the experience of the pathologist (15) and this is especially important in FNA.

The percutaneous biopsy's first challenge is obtaining an appropriate sample, which means sufficient in quantity and representative of the lesion. This point is measured by yield, and values can vary between 69 and 97% for FNA (5,7) and up to 97% for CNB (5,7).

There are two main reasons that help explaining the wide variation of rates in FNA: the type of lesion selected and the accomplishment of preliminary evaluation. Lesions with lower diagnostic yield by percutaneous biopsy are cysts, lesions with and surrounding cortex and lesions with a dense calcified matrix (5,10). In this study, from the

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IB and final diagnosis. FNA - Fne Nedle Apiration. CNB - Core Needle Biopsy.	
Table 1 – Demographic and pathologic information for 94 patientes with the FNA, CN	B – henion M – malionant Y – match N – nonmatch ND – non diaonostic

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COMPARISON OF FINE NEEDLE ASPIRATION AND CORE NEEDLE BIOPSY

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Ŋ	ND	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Υ	Y	Υ	Υ	Υ	Υ	ND	Υ	Υ	Υ	ND	Υ	Υ	Υ	Υ	Υ	ND	Υ	Υ	ND	Ν	Υ	ND	Z	ND
В	Μ	М	Μ	Σ	Μ	Μ	Σ	Σ	В	Σ	М	М	Σ	М	М	в	Σ	М	Μ	М	В	В	В	М	М	М	М	М	ND	М	М	QN	М	X
ND	ND	Μ	Μ	Μ	Μ	Μ	Μ	Μ	В	Μ	Μ	Μ	Μ	Μ	Μ	QN	Μ	Μ	Μ	ND	В	В	В	Μ	Μ	ND	Μ	Μ	ND	Μ	Μ	ND	Μ	ND
Infection	Lymphoma	Metastasis	Metastasis	Metastasis	Myeloma		Metastasis	Metastasis	Exclusion tumour	Metastasis	Myeloma	Metastasis	Metastasis	Metastasis	Metastasis	Enchondroma	Lymphoma	Metastasis	Myeloma	Metastasis	Giant Cell Tumor	Exclusion tumour	Exclusion tumour	Myeloma	Ewing sarcoma	Chondrosarcoma	Metastasis	Chondrosarcoma	Osteosarcoma	Osteosarcoma	Metastasis	Ewing sarcoma	Ewing Sarcoma	Metastasis
Infection	Lymphoma	Metastasis	Metastasis	Metastasis	Myeloma	Chondrosarcoma	Metastasis	Metastasis	Exclusion tumour	Metastasis	Myeloma	Metastasis	Metastasis	Metastasis	Metastasis	Enchondroma	Lymphoma	Metastasis	Myeloma	Metastasis	Giant Cell Tumor	Exclusion tumour	Exclusion tumour	Myeloma	Ewing sarcoma	chondrosarcoma	Metastasis	Chondrosarcoma	Inconclusive	Osteosarcoma	Metastasis	Inconclusive	Ewing Sarcoma	Metastasis
Inconclusive	Inconclusive	Metastasis	Metastasis	Metastasis	Myeloma	Chondrosarcoma	Metastasis	Metastasis	Exclusion tumour	Metastasis	Myeloma	Metastasis	Metastasis	Metastasis	Metastasis	Inconclusive	Lymphoma	Metastasis	Myeloma	Inconclusive	Giant Cell Tumor	Exclusion tumour	Exclusion tumour	Myeloma	Ewing sarcoma	Inconclusive	Metastasis	Chondrosarcoma	Inconclusive	Malignant lesion	Metastasis	Inconclusive	Malignant lesion	Inconclusive
Spine	Femur	Humerus	Femur	Acetabulum	Spine	Femur	Humerus	Iliac	Iliac	Iliac	Iliac	Iliac	Tibia	Humerus	Metacarpal	Sacrum	Sacrum	Rib	Iliac	Spine	Tibia	Femur	Femur	Sternum	Scapula	Femur	Humerus	Humerus	Iliac	Iliac	Iliac	Femur	Femur	Iliac
Male	Female	Female	Male	Male	Male	Female	Female	Female	Female	Male	Male	Female	Female	Female	Male	Male	Male	Female	Male	Female	Male	Male	Male	Male	Male	Female	Male	Male	Female	Female	Male	Male	Male	Male
76	73	77	61	60	69	68	74	70	70	85	64	59	78	76	60	53	53	59	83	53	38	56	56	81	22	80	76	24	67	67	78	20	20	66
60	61	62	63	64	65	99	67	68	69	70	71	72	73	74	75	76	77	78	62	80	81	82	83	84	85	86	87	88	89	06	16	92	93	94

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COMPARISON OF FINE NEEDLE ASPIRATION AND CORE NEEDLE BIOPSY

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	Fine Needle Aspiration	Core Needle Biopsy	p value
Diagnostic yield	70/94 (74.5%)	92/94 (97.9%)	< 0.0001
Diagnostic accuracy	68/70 (97.1%)	91/92 (98.9%)	0.4046
Specificity	100.0%	100.0%	1.0
Sensibility	98.31%	98.51%	0.9288
Positive predictive value	100.0%	100.0%	1.0
Negative predictive value	95.24%	96.15%	0.8792

Table II – Accuracy of Fine Needle Aspiration (FNA) and Core Needle Biopsy (CNB) regarding determining malignancy in comparison to the final diagnosis. P values indicate de differences of both biopsy techniques.

24 non-diagnostic FNA it was possible to find at least 18 lesions with these characteristics. The preliminary evaluation comes from the observation of the sample by the pathologist during the procedure, allowing its repetition if necessary, with substantially improved results when compared to studies where this evaluation is not performed (5,10). In this study, the quantity and quality of the sample was decided by the executant alone without the presence of the pathologist. It is possible that this is one of the reasons for the poor diagnostic yield (74,5%) of FNA.

All the cases of non-diagnostic results were due to technical issues with samples such as tissue scarcity, acellularity or an artifactually distorted specimen. than 95% (1,7,10,12,13). Here, the accuracy of FNA was equivalent to that of CNB on all accounts and close the highest published rates (3,6,11,12), showing the reliability of this technique in diagnosing benign tumours, sarcomas, metastases, infections, haematologic disease lesions and in excluding pathology.

In many cases of musculoskeletal tumours, the specific diagnosis has a minor role in the initiation of treatment. The histological grade, staging and anatomical location are the most important factors for therapeutic decisions and it may even be said that the existing protocols are less based on the histological subtype. Some authors go further, referring to the minor importance of histological subtype and highlighting the relevance

Table III – Comparison of FNA and CNB excluding malignancy, establishing diagnosis and initiating treatment.

	Fine Needle Aspiration	Core Needle Biopsy	p value
Excluding malignancy	78/79 (98.7%)	91/92 (98.9%)	0.9047
Establishing diagnosis	68/79 (86.1%)	91/92 (98.9%)	0.0011
Initiating treatment	73/94 (77.7%)	91/94 (96.8%)	0.0001

The accuracy of a diagnostic technique is the most important parameter in its assessment, and obtaining an exact result is its main objective. In different studies, the diagnostic accuracy of FNA varies between 67% and 99%, where the lowest values are obtained in smaller samples (4,6,11). If it were only considered studies with high samples (n > 300) this value would be greater

of the distinction between sarcoma and metastasis, since the treatment of most sarcomas in adults is primarily based on its size, location and proximity to vital structures (2). Kilpatrick and colleagues (9) considered FNA sufficient to initiate treatment in 87% of bone tumours. In a study conducted in 2010, concerning soft tissue masses of extremities, definitive treatment could be initiated based solely

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on FNA in 81.3% of benign, in 78% of malignant and in 43% of indeterminate tumours (13). Assuming the same criteria, the technique in the present study would therefore allow for the initiation of treatment in all 68 patients with a diagnosis proven correct and in the other 5 in which malignancy had been excluded. This would represent 73 of the 94 (77,7%). In other words, although statistically inferior to CNB due to the inconclusive results, if these are results were excluded, FNA would be a reliable technique and would enable the treatment.

Finally, caution should be taken in malignancies since the initial treatment is different according to each diagnosis. The utility of cytogenetics in the routine work-up of sarcomas collected by FNA has been reinforced (8). Nevertheless this was not done in this study.

In conclusion, FNA is reliable and enables the initiation of treatment every time it establishes a diagnosis or excludes malignancy. The number of inconclusive cases, the real problem with this technique, can potentially be decreased by a better selection of the lesions to be analysed by this technique and by the preliminary evaluation by a pathologist. Until then, CNB remains the preferable method for bone tumours diagnosis.

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