

6a.6 NCB and VANCB reporting guidelines

This section of this document is designed to assist in classification of needle NCB and VANCB samples.

6a.6.1 Specimen information & handling

Proper interpretation of core biopsies requires knowledge of details of both clinical and imaging findings (mammography/US) and this information should be provided on the request form. The completed request form should include clinical details, specifying the imaging findings, the site of biopsies and the number of cores.

The tissue fixation protocol should be based on a standard procedure agreed between the departments involved.

Biopsies performed from microcalcifications should be x-rayed to determine the presence of calcification. A comment regarding the presence of **representative** microcalcification of the mammographic lesion in the sample should be provided along with the specimen x-ray.

Biopsies should be placed in fixative solution immediately and sent promptly to the laboratory.

The specimen x-ray procedure should not interfere with prompt fixation. Before fixation, the specimen should be arranged straight, if multiple in parallel. It is recommended that no more than four NCB specimens should be placed into a single cassette.

In cases of mammographic microcalcification, examination of further levels should be performed if calcification is not immediately apparent on histological examination. In problem cases further levels and/or radiography of paraffin blocks may be helpful.

Figure 1a: Specimen NCB/VANCB reporting form

Breast screening NCB/VANCB Form					
Surname _____		Forenames _____			
		Date of Birth _____			
Screening No _____		Hospital No _____			
Centre _____		Report No _____			
Side	Right	Left	Quadrant : _____		
Clinical details: _____					

Radiological category	R1	R2	R3	R4	R5
Radiological Appearance					
Spiculate mass	Stellate lesion	Microcalcification,	coarse	branching	
Well defined mass	Architectural distortion		fine	clustered	
Localisation technique	Palpation	Stereotactic	Ultrasound guided		
Specimen type	NCB	VANCB	Number of cores _____		
Calcification present on specimen X-ray		Yes	No	Radiograph not seen	
Histological calcification	Absent	In benign changes	In malignancy	In both	
Opinion of Pathologist	B1. Uninterpretable/Normal tissue only B2. Benign B3. Lesion of uncertain malignant potential B4. Suspicious of malignancy B5. Malignant				
	a. In situ carcinoma b. Invasive carcinoma c. Invasive status not assessable d. Other malignancy				
Pathologist _____		Operator taking biopsy _____			
		Date _____			
Comment _____					

Figure 1b: NCB/VANCB reporting form

Optional further information:				
Benign lesion				
Fibroadenoma	Solitary papilloma	Multiple papilloma		
Fibrocystic change	Sclerosing adenosis	Complex sclerosing lesion/radial scar		
Periductal mastitis/duct ectasia			Columnar cell change	
Other (please specify) _____				
Epithelial proliferation				
Not present	Present without atypia	Present with atypia (ductal)		
Present with lobular intraepithelial neoplasia				
Columnar cell change with atypia				
Malignant lesion				
In situ carcinoma	Not present	Ductal	Lobular with necrosis	
DCIS grade	High	Intermediate	Low	Not assessable
Invasive Carcinoma	Not present	Present		
Oestrogen receptor status:	Positive	Negative	____ Quick Score (Allred)	
Not performed				
Comment _____				

6a.6.2 Recording basic information

Information on the nature of the mammographic abnormality and clinical characteristics should be provided by the breast screening radiologist requesting the pathology examination.

Centre/Location

Give the name of the assessment centre, clinic, department etc., where the specimen was obtained.

Side

Indicate right or left.

For specimens from both sides use a separate form for each side.

Quadrant

The quadrant of the index lesion can be entered here.

Radiological category

The radiological assessment of the lesion should be entered here.

Radiological appearance

This section of the form is to be filled in by the clinician to indicate the radiological abnormality.

- Spiculate mass
- Stellate lesion
- Well defined mass
- Microcalcification. This should be classified by the requesting radiologist into coarse or fine and branching or clustered
- Architectural distortion

Localisation technique

Please choose one of the following terms:

- Palpation NCB guided by palpation
- Stereotactic NCB/VANCB guided by stereotaxis
- Ultrasound guided CB/VANCB guided by ultrasound.

Number of cores

If known indicate the number of NCB samples taken.

It is recommended that any cores containing calcification are so identified and are sent separately to the pathologist.

Calcification present on specimen X-ray

Indicate whether there is calcification visible on the specimen radiograph if available.

Histological calcification

Indicate whether calcification has been identified in the sample and if present whether it is associated within a benign or malignant lesion.

Pathologist

The name of the pathologist giving the histological opinion. The pathologist should be registered at the screening office.

Operator

Enter the name of the operator performing the biopsy.

Date

Enter the date of reporting the slides.

Comment field

This free text field is included for extra information to be recorded.

Optional further information

In some cases it may be helpful to record further information. This will be particularly so where

neo-adjuvant or primary chemotherapy is contemplated or for VANCB specimens where the whole lesion may be resected. The fields chosen are the relevant fields from the main histology form (See section B).

6a.6.3 Reporting categories

It is important to remember that histological examination of both NCB and VANCB samples is performed to fulfil the assessment process role by giving a pathology category classification (B1-5) and not designed to give a definitive diagnosis, although this is possible in the majority of cases. Thus whilst most samples can be readily categorised as normal, benign or malignant, it must be recognised that a small proportion (probably less than 10%) of samples cannot. The following reporting guidelines have been devised in recognition of this and should be used for all screen-detected lesions (microcalcification, architectural deformities and mass lesions). It is also important to remember that, although there are five reporting categories similar to those used in fine needle aspiration cytology (FNAC), these are not equivalent.

B1. Normal tissue/uninterpretable

This indicates a core of normal tissue whether or not breast parenchymal structures are present; thus this category is equally appropriate for a core including normal breast ducts and lobules or mature adipose/fibrous tissue only. A B1 report should include a description of the components present and comment should be made regarding the presence of breast epithelial structures. Cores with B1 diagnoses may contain microcalcification, for example within involutinal lobules. It is important in these cases that discussion between pathology and radiology colleagues is undertaken to confirm the appropriateness of the microcalcification in the histological specimen. Small foci of calcification within involuted lobules are common and frequently too small to be visible mammographically, thus a report that merely records the presence of this calcification without additional comment on its nature, size and site may be misleading and lead to false reassurance. Mammograms do not demonstrate microcalcification, either singly or in clusters, less than 100 microns in diameter.

Exceptionally some specimens may be classified as uninterpretable, for example due to excessive crush artefact or composed of blood clot only. Such samples should also be classified as B1 although some experts would prefer these to be classified as B0.

B2. Benign lesion

A core is classified as B2 Benign when it contains a benign abnormality. This category is appropriate for a range of benign lesions including fibroadenomas, fibrocystic changes, sclerosing adenosis and duct ectasia and extends to include other non-parenchymal lesions such as abscesses and fat necrosis.

In some cases it may be difficult to determine whether a specific lesion is present, for example if minor fibrocystic changes are seen. The multi-disciplinary approach is once again vital in these cases to determine whether the histopathological features are in keeping with the radiological and clinical findings. It may be appropriate and prudent to classify the lesion as B1, rather than B2 if only very minor changes are present; such histopathological features would clearly be insufficient to explain a well-defined mass lesion and classification as B2 would be inappropriate.

B3. Lesion of uncertain malignant potential

This category mainly consists of lesions which may provide benign histology on NCB, but either are known to show heterogeneity or to have an increased risk (albeit low) of associated malignancy.

The B3 category has a lower rate of malignancy on further surgical biopsy (25%) when compared with B4 (66%). The majority of B3 lesions require surgical excision, but all these cases should be discussed at a preoperative multidisciplinary meeting.

I. Papillary lesions

Papillary lesions may show intralesional heterogeneity and the limited sampling achieved with NCB may miss areas of in situ cancer. The majority of these lesions should, therefore, also be designated B3 of uncertain malignant potential. On rare occasions when a small lesion has been very widely sampled and submitted for pathological examination, a benign B2 classification may be considered. Conversely, when a sample of a papillary lesion in a NCB shows atypia, for example strongly suspicious of papillary carcinoma in situ, a B4 designation may occasionally be more appropriate.

II. Radial scar/complex sclerosing lesion

Biopsies which show features of a radial scar/complex sclerosing lesion such as areas of hyalinisation, elastosis, or tubular entrapment with epithelial proliferation should be categorised as B3 if they represent the cause of the radiological abnormality^{30a}. Actually, these lesions are heterogeneous and a proportion of them are associated with atypia or malignancy (in general LIN or low grade DCIS).

III. Lobular intraepithelial neoplasia (LIN)

A small cell regular epithelial proliferation within moderately distended lobules which is considered by the pathologist to represent lobular intraepithelial neoplasia or LIN (regrouping ALH and LCIS) should be classified as B3: this process does not necessarily have the same management implications as a diagnosis of DCIS but surgical diagnostic excision might be considered. Lobular intraepithelial neoplasia is frequently a co-incidental finding in a core biopsy from a screen-detected lesion however and multidisciplinary discussion is essential as the abnormality identified radiologically may not be represented. Furthermore, it may be that LIN encountered serendipitously in a breast surgical excision does not carry the same risk and prognosis as LIN diagnosed via a targeted NCB/VANCB of a mammographic abnormality³¹. These cases must be managed cautiously.

On occasions it may be impossible to classify a small cell epithelial proliferation in lobules and/or ducts as either lobular neoplasia or low grade DCIS and in these circumstances a numerically higher category (B4 or B5) is prudent and should be considered. In these cases, E-cadherin may help in the differential diagnosis³². Pleomorphic LIN may also be classified as B5. There is at present, however, no definite follow-up information on these lesions and management should be discussed in a multidisciplinary forum.

IV. Atypical epithelial proliferation of ductal type

The definition of atypical ductal hyperplasia (ADH) is derived from surgical resection specimens and relies on a combination of histological, morphological and size extent criteria. There is a range of severity from those which are insufficient for a definite diagnosis of DCIS but highly suspicious to those which only show a minor degree of atypia, normally architectural which requires further assessment. In some cases, the appropriate categorisation may be B4 in lesions highly suspicious of DCIS. These proliferations must be clearly separated from usual epithelial hyperplasia (see pitfalls).

A definitive diagnosis of ADH is not possible on NCB. It has been shown that core biopsy samples which include atypical epithelial proliferative foci of ductal type, of insufficient extent for classification as DCIS, on subsequent surgical resection may form part of an established in situ neoplastic lesion with or without associated invasion. This view is based on several studies which describe the subsequent surgical diagnosis in cases described as ADH in NCB. In over 50% of cores surgical excision biopsy has shown either in situ or invasive carcinoma³³. The limited tissue sampling which can be undertaken by NCB guns (often by stereotactic methods for foci of microcalcification) may thus provide insufficient material for definitive diagnosis of low grade DCIS if only a few involved duct spaces are obtained. In all cases open biopsy is indicated to evaluate the lesion, define its extent, and to exclude invasive growth. It should be clear that ADH cannot be diagnosed on NCB, and that it is incorrect to use the term ADH for cases where core biopsies include atypical intraductal epithelial proliferative foci, or an area of well differentiated DCIS insufficient in extent for classification as DCIS. These should be diagnosed as atypical epithelial proliferation of ductal type.

V. Phyllodes tumour

Fibroepithelial lesions suggesting phyllodes tumour (cellular stroma, stromal overgrowth and possibly some mitotic activity) should also be designated B3. Thus the presence of a cellular stroma within a fibroepithelial lesion should prompt a search for other features that may aid in discrimination from a fibroadenoma. In practice, however, this distinction is often impossible and careful appraisal of the entire clinical picture will usually allow appropriate management to be undertaken. Obviously malignant cases should be classified as B5.

B4. Suspicious of malignancy

Technical problems such as crushed or poorly fixed cores which contain probable carcinoma but cannot provide the definitive diagnosis are best included as B4. Similarly, apparently neoplastic cells contained within blood clot or adherent to the outer aspect of the sample should be classified as B4 suspicious.

A complete single duct space bearing an unequivocal high-grade epithelial proliferative process can be classified as B5 malignant. However care must be taken if one or only part of a duct space is seen containing a highly atypical epithelial process particularly if no necrosis is present; this may be regarded as suspicious rather than definitively malignant. In particular great care should be taken if the epithelial cells show any features of an apocrine phenotype, which may represent an atypical apocrine proliferation rather than DCIS.

The management of cases classified as B4 will usually be either diagnostic excision biopsy of the area or repeat NCB sampling to obtain a definitive diagnosis. **Definitive therapeutic surgery should not be undertaken as a result of a B3 or B4 NCB diagnosis.**

B5. Malignant

This category is appropriate for cases of unequivocal malignancy on NCB. Further categorisation into in situ and invasive malignancy should be undertaken whenever possible. Other forms of malignancy such as malignant lymphoma may also be classified as B5.

I. Lobular intraepithelial neoplasia (see above B3)

Lobular intraepithelial neoplasia is included in the B3 category, as it does not have the same management implications as a diagnosis of DCIS or invasive malignancy. Nevertheless the pleomorphic variant or LIN with comedo-necrosis may be classified as B5.

II. Ductal carcinoma in situ

One of the benefits of NCB is that it can allow distinction between in situ and invasive carcinoma. It should however be borne in mind that, due to sampling error, exclusive presence of DCIS in the core does not exclude the possibility of an invasive focus being present. In approximately 20% of cases sampled by standard methods co-existing invasive carcinoma will be identified in the subsequent surgical excision specimen²¹. The nuclear grade, architecture and the presence of necrosis within the DCIS can be indicated on the NCB report. In particular, the presence of associated calcification should be recorded. Biopsies of skin for Paget's disease may also be recorded as non-operative diagnostic procedures and can be classified accordingly.

III. Invasive carcinoma

An advantage of NCB over FNAC is the ability to diagnose invasion positively. Invasive mammary carcinoma can be unequivocally identified in NCB with a positive predictive value of 98%³⁴. As noted above, however, the negative predictive value for invasion is only 80% when only DCIS is identified. Assessment of grade and type of carcinoma may be achieved (although concordance with final grade and type are not absolute and, if performed, should be interpreted with caution)^{35,36}.