A guide to

Invasive Breast Cancer Histopathology Reporting

	al details	
<u>\$1.02</u>	Clinical information provided on request form (complete as narrative or use the structured format below)	Text
	Specimen type	See p3
	SENTINEL NODES (if submitted)	
	Location (eg Axillary, Internal mammary)	Text
	Sentinel nodes number	
	Sentinel nodes colour	Text
	Radioactive count	
	Tumour site and laterality (use clock-face analogy)	Text
	Method of localisation	Carbon track Hook wire
	New primary cancer or recurrence	New primary Local recurr. Distant met.
<u>G1.01</u>	Any other relevant information (eg history, clinical dx, neoadjuvant therapy, lab and imaging results)	Text
<u>S1.03</u>	Pathology accession number	Text
<u>\$1.04</u>	Principal clinician	Text
Macro	oscopic findings	
S2 01	No. of specimens submitted	
52.02	Specimen laterality	
52.02	Specifien laterality	Right
<u>\$2.03</u>	Specimen type/Lymph tissue	See p3
	Intraoperative consultation	Not perfomed Performed
	If perfomed record type	See p3
<u>S2.04</u>	Specimen orientation	Not oriented Oriented
	If oriented, record markers and locations	Text
S2.05	Method of localisation	carbon
		hook wire N/A
<u>S2.06</u>	Specimen size	xx mm
	If oriented use the following 3 measures:	
	Medial-lateral length	mm
	Superficial-deep length	 mm
	Superior-inferior length	mm
S2.07	Specimen weight	g
<u>\$2.08</u>	Macroscopically visible tumours?	Absent Present
	If present, record the no. of foci	

Macroscopic findings (cont.) Gross descript. of tumour/s S2.09 (for each tumour record) Nature of tumour Text Tumour size __x_x_mm mm Distance to nearest separate tumour foci _mm from Min. macro. margin clearance (specify margin) from any tumour deposit S2.10 Skin Absent Present If present, record ... ___x__mm Skin dimensions Skin abnormalities Absent OR Ulceration Paget disease Satellite nodules Other (specify) S2.11 Muscle Absent Present S2.12 SENTINEL LYMPH NODES (for each node received record...) Site Axilla Int. mamm chain **Radioactive count** Uptake of dye No Yes - Blue Size __x__x__mm NON SENTINEL LYMPH **NODES/TISSUE** Total number of nodes __to__mm Size range Description Text S2.13 Block identification key Text G2.01 Other macroscopic findings Text **Microscopic findings** S3.01 Multiple tumours? Absent Present If present, record: **Quadrants involved** Text Total no. of tumour deposits (if >2 record) Max span of multifocal ____X___mm tumour bed involved For EACH tumour identified above complete S3.02-S3.04 and consider recording G3.01 S3.02 MAX. INVASIVE TUMOUR

SIZE Whole tumour size Max. size of invasive tumour

___mm

mm



Microscopic findings (cont.)				Microscopic findings (cont.)				
<u>G3.01</u>	Other invasive tumour dimensions	xmm	<u>\$3.16</u>	Paget disease	Absent Present			
<u>\$3.03</u>	Histological grade (record for each tumour) (Refer to p3)	3–5 (Grade 1) 6–7 (Grade 2) 8–9 (Grade 3)	<u>S3.17</u>	Margin involvement by invasive carcinoma or DCIS	Not involved Involved			
<u>S3.04</u>	Invasive carcinoma subtype	See p4		If involved, specify				
<u>\$3.05</u>	Peritumoural lymphovascular invasion	Not identified Present Suspicious		for <i>each</i> involved margin:				
	If suspicious, record the block	Text		Type of involvement	DCIS Invasive ca DCIS & invasive ca			
S3.06	Skin	See p4			Tout			
<u>\$3.07</u>	Muscle	Not involved Involved		Extent of involvement	mm OR			
<u>\$3.08</u>	Treatment effect (after neoadjuvant hormonal or	No definite resp. Partial response		If not involved,	Focal			
	chemotherapy) (Refer to p4)	Complete resp. Not applicable		record for <i>each</i> close margin				
	If no definite or partial response record the estimate	%		Distance of invasive carcinoma to margin	mm OR >10mm			
	of overall level of cellularity for invasive cancer			If DCIS is closer to the margin the invasive ca additionally record:	&mm from DCIS			
	Specify neoadjuvant response classification	Text	<u>\$3.18</u>	Lobular neoplasia	Absent Present			
	system used Result of treatment	Text		If present, record type	Classical Variant			
S3.09	DCIS	See p4		Extent	Focal			
S3.10	Max. extent of breast	mm	C2 02	LCIS at the margin	Extensive			
	involved by DCIS		<u>G3.03</u> S3 10		See p4			
<u>S3.11</u>	Max. dimension pure DCIS	mm	62.20	SENTINEL NODES (SN)	5ee p-			
<u>55.12</u>	DCIS	Intermediate	33.20	Total number of SN				
C2 02	Nuclear and between situat	High		Number of SN with				
<u>G3.02</u>	DCIS	Present		macrometastases				
	If present, record, next most prevalent grade	Low Intermediate High		micrometastases				
S 3.13	Necrosis in DCIS	Absent		tumour cells				
		Present	<u>S3.21</u>	NONSENTINEL NODES (NSN)				
<u>\$3.14</u>	Architecture of DCIS (Select all that apply)	comedo solid		Total number of NSN				
		cribriform micropapillary apocrine papillary other (specify)		Number of NSN with metastases	—			
			<u>\$3.22</u>	Extranodal spread	Absent Present			
S3.15	Microcalcification	Absent	<u>S3.23</u>	Treatment effect in LN	See p4			
		Present	<u>G3.04</u>	Other microscopic comments	lext			
	If present record	in DCIS	Ancil	lary test findings				
	in which tissue(s):	in benign tissue in invasive cancer	<u>\$4.01</u>	Oestrogen receptors	Not performed Performed Pending			
	Lesion(s) with microcalcification	iext		If performed, record				
	Associated with	No	%age nuclei staine		to %			
	necrosis? Size and extent of	Yes Text		Predominant staining intensity	1+ Low 2+ Intermed.			
	microcalcification (if required, per lesion with microcalcification)			ER result	Negative Positive			

Ancillary test findings							
Progest	trone receptors	Not performed Performed Pending					
	If performed, record						
	%age nuclei stained	to %					
	Predominant staining intensity	1+ Low 2+ Intermed. 3+ High					
	PR result	Negative Positive					
<u>S4.03</u> HER2 (1	ISH)	Not performed Performed Pending					
	If performed, record:						
	Number of copies of HER2						
	Number of copies of CEP17 (if assessed)						
	HER2 result	Amplified Non-amp. diploid Non-amp polysomic Indeterminate					
HER2 I	HC (if performed)	0 1+ 2+ 3+ Not performed					

Synthesis and overview						
<u>\$5.01</u>	Tumour stage and group	See p4 and 5				
<u>\$5.02</u>	Year & edition of staging system	Text				
<u>G5.01</u>	Diagnostic summary (include: Specimen type and laterality; Histological grade; Max. tumour size; Margin status; Lymph node status; Lymphovascular invasion)	Text				
<u>\$5.03</u>	Overarching comment	Text				

S1.02 /S2.03 Specimen type

Single select from the following:

- diagnostic open biopsy
- wide local excision (partial mastectomy, quadrantectomy or segmentectomy)
- re-excision
- mastectomy
- mastectomy post neoadjuvant therapy
- other (specify)

Lymph tissue - choose all that apply:

- not submitted
- lymph node biopsy sentinel
- Iymph node biopsy non-sentinel
- axillary sample
- axillary clearance
- other (specify)

S2.03 Intraoperative consult

Choose all that apply:

- frozen section
- imprint cytology
- gross examination for margin assessment
- other (specify)

S3.03 Histological grade

Nuclear grade

Score 1:	Size equivalent to normal breast epithelial
	cells, regular outlines, uniform chromatin;
	inconspicuous nucleoli, little size variation.
Score 2:	Larger nuclei, open vesicular chromatin: visib

- Score 2: Larger nuclei, open vesicular chromatin; visible nucleoli, moderate variability in size and shape
 Score 3: Vesicular nuclei; often with prominent nucleoli;
- exhibiting marked variation in size and shape, occasionally very large and bizarre forms.

Tubular differentiation

- Score 1: >75% of invasive carcinoma forming tubular or glandular structures
- Score 2: 10–75% of invasive carcinoma forming tubular or glandular structures
- Score 3: <10% of invasive carcinoma forming tubular or glandular structures.
- Not assessable*
- * microinvasion only (each focus \leq 1mm)

Mitotic counts

Number of mitoses per 10 high-power fields use the tables below. OR Not assessable (ie microinvasion only (each focus \leq 1mm)

Field diameter	Mitotic frequency score			Field diameter	Mitotic frequency score			Field diameter	Mitotic frequency score		
(mm)	1	2	3	(mm)	1	2	3	(mm)	1	2	3
0.40	≤4	5-9	≥10	0.50	≤7	8-14	≥15	0.60	≤10	11-20	≥21
0.41	≤4	5-9	≥10	0.51	≤7	8-14	≥15	0.61	≤10	11-21	≥22
0.42	≤ 5	6-10	≥ 11	0.52	≤7	8-15	≥16	0.62	≤ 11	12 - 22	≥23
0.43	≤5	6-10	≥ 11	0.53	≤8	9-16	≥17	0.63	≤ 11	12-22	≥23
0.44	≤5	6-11	≥12	0.54	≤8	9-16	≥17	0.64	≤ 11	12-23	≥24
0.45	≤5	6-11	≥12	0.55	≤8	9-17	≥18	0.65	≤12	13-24	≥25
0.46	≤6	7-12	≥13	0.56	≤8	9-17	≥18	0.66	≤ 12	13-24	≥25
0.47	≤6	7-12	≥13	0.57	≤9	10-18	≥19	0.67	≤ 12	13-25	≥26
0.48	≤6	7-13	≥14	0.58	≤9	10-19	≥20	0.68	≤13	14-26	≥27
0.49	≤6	7-13	≥ 14	0.59	≤9	10-19	≥20	0.69	≤13	14-27	≥28

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S3.04 Invasive carcinoma subtype

- Invasive carcinoma of No Special Type (Ductal)
 - Pleomorphic carcinoma
 - Carcinoma with osteoclast like stromal giant cells
 - Carcinoma with choriocarcinomatous features
 - Carcinoma with melanotic features Invasive lobular carcinoma
- Classical
- Tubulolobular
- Alveolar
- Solid
- Pleomorphic
- Mixed
- Others signet ring, histiocytoid, etc
- Tubular carcinoma
- Cribriform carcinoma
- Mucinous carcinoma
- Carcinoma with medullary features
 - Medullarv

•

- Atypical medullary
- Invasive carcinoma NST (ductal) with medullary features
- Carcinoma with apocrine differentiation
- Carcinoma with signet ring cell differentiation
- Invasive micropapillary carcinoma .
- Metaplastic carcinoma
 - Low grade adenosquamous carcinoma
 - Fibromatosis-like metaplastic carcinoma
 - Squamous cell carcinoma
 - Spindle cell carcinoma
 - Metaplastic carcinoma with mesenchymal differentiation
 - Chondroid differentiation •
 - Osseous differentiation
 - Other types of mesenchymal differentiation
- Mixed metaplastic carcinoma
- Myoepithelial carcinoma

Rare Types of Invasive Cancer:

- Carcinomas with Neuroendocrine features
 - Neuroendocrine tumour, well differentiated Neuroendocrine tumour, poorly differentiated (small cell carcinoma)
- Carcinoma with neuroendocrine differentiation
- Secretory carcinoma
- Invasive papillary carcinoma
- Acinic cell carcinoma
- Mucoepidermoid carcinoma •
- Polymorphous carcinoma •
- Oncocytic carcinoma
- Lipid rich carcinoma •
- Glycogen rich/Clear cell carcinoma •
- Sebaceous carcinoma
- Salivary gland/skin adnexal type tumours .
- Adenoid cystic carcinoma
- Adenomyoepithelioma with carcinoma

S3.06 Skin

- Not involved
- Paget disease of the nipple (DCIS extending to skin contiguous with lactiferous sinuses)
- Invasive carcinoma involving dermis or epidermis without ulceration
- Invasive carcinoma involving dermis or epidermis with ulceration
- Ipsilateral satellite skin nodules, ie dermal deposits of invasive carcinoma, separate from the main tumour

S3.23 Treatment effect in LN

V2.3 Guide derived from Invasive Breast Cancer Structured Reporting Protocol 2nd Edition

- nodes negative, no treatment effect
- nodes negative, with treatment effect .
- nodes positive, with treatment effect
- nodes positive, no treatment effect
- Not applicable

S3.08 Treatment effect

- No definite response to pre-surgical therapy in the invasive carcinoma
- Partial response to pre-surgical therapy in the invasive carcinoma, residual carcinoma identified.
- Complete pathologic response in breast and lymph nodes: No residual invasive carcinoma is present in the breast or lymph nodes after pre-surgical therapy
- Not applicable

S3.09 DCIS

- Absent
- Present only in conjunction with invasive carcinoma
- Present only as pure DCIS
- Present as both pure DCIS and in conjunction with invasive carcinoma

G3.03 LCIS at the margin

- LCIS with comedo necrosis present
- Pleomorphic LCIS present

S3.19 Assoc. breast changes

- atypical ductal hyperplasia
- flat epithelial atypia
- lobular neoplasia (ALH/ LCIS)
- radial scars
- sclerosing adenosis
- fibrocystic change
- other breast changes (eq calcification) (specify)

S5.01 Tumour stage and group#

TNM descriptors (Only if applicable; select all that apply)

- m- multiple foci of invasive carcinoma
- r recurrent y - post treatment

Τ4

T4a

T4b

T4c

T4d

as pT4

Both T4a and T4b

Inflammatory carcinoma

Primary Tumour (Invasive Ca) (pT)

- ΤХ Primary tumour cannot be assessed
- No evidence of primary tumour т0
- Tis (DCIS) Ductal carcinoma in situ
- Tis (LCIS) Lobular carcinoma in situ
- Tis (Paget's) Paget disease of the nipple NOT associated with invasive carcinoma and/or carcinoma in situ (DCIS and/or LCIS) in the underlying breast parenchyma. Carcinomas in the breast parenchyma associated with Paget's disease are categorized based on size and characteristics of the parenchymal disease, although the presence of Paget's disease should still be noted.
- T1 Tumour \leq 20 mm in greatest dimension
- T1mi Tumour ≤ 1 mm in greatest dimension
- T1a Tumour >1 mm but \leq 5 mm in greatest dimension
- T1b Tumour >5 mm but ≤10 mm in greatest dimension
- T1c Tumour >10 mm but \leq 20 mm in greatest dimension

Tumour of any size with direct extension to the chest

wall and/or to the skin (ulceration or skin nodules) Note: Invasion of the dermis alone does not qualify

Ulceration and/or ipsilateral satellite nodules and/or oedema (including peau d'orange) of the skin, which

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do not meet criteria for inflammatory carcinoma

Extension to the chest wall, not including only

pectoralis muscle adherence/invasion

- Т2 Tumour >20 mm but \leq 50 mm in greatest dimension
- Т3 Tumour >50 mm in greatest dimension

Regional Lymph Nodes (pN)*

*Note: Classification is based on axillary lymph node dissection with or without sentinel lymph node biopsy. Classification based solely on sentinel lymph node biopsy without subsequent axillary lymph node dissection is designated (sn) for "sentinel node" for example, pN0(sn)

- pNX Regional lymph nodes cannot be assessed (eg previously removed, or not removed for pathologic study)
- pN0 No regional lymph node metastasis identified histologically.

Note: isolated tumour cell clusters (ITC) are defined as small clusters of cells not greater than 0.2mm, or single tumour cells, or a cluster of fewer than 200 cells in a single histologic cross-section. ITCs may be detected by routine histology or by immunohistochemical (IHC) methods. Nodes containing only ITCs are excluded from the total positive node count for purposes of N classification but should be included in the total number of nodes evaluated.

- pN0(i-) No regional lymph node metastases histologically, negative IHC
- pN0(i+)Malignant cells in regional lymph node(s) no greater than 0.2 mm (detected by H&E or IHC including ITC)
- pN0(mol-) No regional lymph node metastases histologically, negative molecular findings (RT-PCR)
- pN0(mol+) Positive molecular findings (RT-PCR)**, but no regional lymph node metastases detected by histology or IHC
- pN1 Micrometastases; or metastases in 1-3 axillary lymph nodes; and/or in internal mammary nodes with metastases detected by sentinel lymph node biopsy but not clinically detected.***
- pN1mi Micrometastases (greater than 0.2 mm and/or more than 200 cells, but none greater than 2.0 mm)
- pN1a Metastases in 1-3 axillary lymph nodes, at least 1 metastasis greater than 2.0 mm
- pN1b Metastases in internal mammary nodes with micrometastases or macrometastases detected by sentinel lymph node biopsy but not clinically detected.***
- pN1c Metastases in 1-3 axillary lymph nodes and in internal mammary lymph nodes with micrometastases or macrometastases detected by sentinel lymph node biopsy but not clinically detected.
- pN2 Metastases in 4-9 axillary lymph nodes; or clinically detected**** internal mammary lymph nodes in the absence of axillary lymph node metastases
- pN2a Metastases in 4-9 axillary lymph nodes (at least one tumour deposit greater than 2.0 mm)
- pN2b Metastases in clinically detected**** internal mammary lymph nodes in the absence of axillary lymph node metastases
- pN3 Metastases in ten or more axillary lymph nodes; or in infraclavicular (level III axillary) lymph nodes; or in clinically detected**** ipsilateral internal mammary lymph nodes in the presence of one or more positive level I, II axillary lymph nodes; or in more than three axillary lymph nodes and in internal mammary lymph nodes with micrometastases or macrometastases detected by sentinel lymph node biopsy but not clinically detected***; or in ipsilateral supraclavicular lymph nodes
- pN3a Metastases in 10 or more axillary lymph nodes (at least one tumour deposit greater than 2.0 mm); or metastases to the infraclavicular (level III axillary lymph) nodes
 - lymph) nodes Used with the permission of the American Joint Committee on Cancer (AJCC), Chicago, Illinois. The original source for this material is the AJCC Cancer Staging Manual, Seventh Edition (2010) published by Springer Science and Business Media LLC, www.springerlink.com

Regional Lymph Nodes (pN)* (cont.)

pN3b Metastases in clinically detected**** ipsilateral internal mammary lymph nodes in the presence of one or more positive axillary lymph nodes; or in more than three axillary lymph nodes and in internal mammary lymph nodes with micrometastases or macrometastases detected by sentinel lymph node biopsy but not clinically detected****

pN3c Metastases in ipsilateral supraclavicular lymph nodes Notes:

**RT-PCR: reverse transcriptase/polymerase chain reaction

- ***'Not clinically detected' is defined as not detected by imaging studies (excluding lymphoscintigraphy) or not detected by clinical examination
- ****'Clinically detected' is defined as detected by imaging studies (excluding lymphoscintigraphy) or by clinical examination and having characteristics highly suspicious for malignancy or a presumed pathologic macrometastasis based on fine needle aspiration biopsy with cytologic examination.

Distant Metastasis (M)

- M0 No clinical or radiographic evidence of distant metastases
- cM0(i+) No clinical or radiographic evidence of distant metastases, but deposits of molecularly or microscopically detected tumour cells in circulating blood, bone marrow, or other nonregional nodal tissue that are no larger than 0.2 mm in a patient without symptoms or signs of metastasis
- M1 Distant detectable metastases as determined by classic clinical and radiographic means and/or histologically proven larger than 0.2 mm

Stage Grouping*

Stage	т	Ν	М
0	Tis	N0	M0
IA	T1*	N0	M0
IB	Т0	N1mi	M0
	T1*	N1mi	M0
IIA	Т0	N1†	MO
	T1*	N1 ⁺	M0
	T2	N0	M0
IIB	T2	N1	M0
	Т3	N0	M0
IIIA	Т0	N2	M0
	T1*	N2	M0
	T2	N2	M0
	Т3	N1	M0
	Т3	N2	M0
IIIB	T4	N0, N1, N2	M0
IIIC	Any T	N3	M0
IV	Any T	Any N	M1

^{*}T1 includes T1mic

 $^{\rm +}$ T0 and T1 tumours with nodal micrometastases only are excluded from Stage IIA and are classified Stage IB.

Notes:

- M0 includes M0(i+)
- The designation pM0 is not valid; any M0 should be clinical.
- If a patient presents with M1 prior to neoadjuvant systemic therapy, the stage is considered stage IV and remains stage IV regardless of response to neoadjuvant therapy.
- Stage designation may be changed if postsurgical imaging studies reveal the presence of distant metastases, provided that the studies are carried out within 4 months of diagnosis in the absence of disease progression and provided that the patient has not received neoadjuvant therapy.
- Post-neoadjuvant therapy is designated with "yc" or "yp" prefi x. No stage group is assigned if there is a complete pathologic response (CR) to neoadjuvant therapy, for example, ypT0ypN0cM0.