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Review paper

Computed tomography imaging of ovarian peritoneal carcinomatosis: a pictorial review

Binit Sureka^{A,B,E,F}, Virendra Meena^{B,E,F}, Pawan Garg^{B,F}, Taruna Yadav^{E,F}, Pushpinder Singh Khera^{A,F}

All India Institute of Medical Sciences, Jodhpur, India

Abstract

Gynaecological malignancies are the most common malignancies in women and also an important public health issue. In developing countries, there is a paucity of screening facilities and cancer awareness, so patients present at an advanced stage of disease, which severely limits the prognosis and clinical outcome. Among the gynaecological malignancies, ovarian malignancy has the second highest incidence in women according to the incidences. Ovarian malignancy is usually diagnosed at the advanced stages, and to improve the patient's survival, debulking surgery is very important. Advanced-stage disease is treated with either debulking surgery followed by adjuvant chemotherapy or initial neoadjuvant chemotherapy followed by debulking surgery. Imaging is very important in patient selection, in determining who will benefit from neoadjuvant chemotherapy just before debulking surgery. This article highlights the role of computed tomography (CT) in the detection of patterns of spread of ovarian malignancy, important for staging and management.

Key words: ovary, cancer, carcinomatosis, pathways, spread.

Introduction

Ovarian cancer is the fifth most common malignancy among women worldwide and the second most common gynaecological malignancy [1]. Ovarian malignancy is usually diagnosed at advanced stage of the disease with five-year survival rate in stage III disease being 37% and stage IV disease being 24% [2]. The International Federation of Gynaecology and Obstetrics (FIGO) ovarian cancer staging system was first published in 1973 and was revised in 1988 and 2014. Table 1 and Figure 1 summarise the new FIGO 2014 staging of ovarian cancer [3,4].

The treatment planning of advanced ovarian malignancies includes both primary debulking surgery followed by adjuvant chemotherapy, and if non-resectable disease is found on preoperative imaging, neoadjuvant chemotherapy followed by debulking surgery. Several recent randomised control trials clearly show similar disease-free survival rates between stage III or stage IV disease in patients who initially underwent chemotherapy followed by debulking surgery [5].

Detection of peritoneal disease on preoperative imaging is very difficult, and interpretation of imaging findings requires in-depth knowledge of complex peritoneal anatomy, peritoneal fluid flow direction, and specific anatomic locations where debulking surgery is technically difficult. The presence of disease at these sites is very helpful for surgeons for planning of surgery.

This article describes the role of radiologists in planning the management of advanced ovarian malignancies. Radiologists provide essential information about routes of disease dissemination and common sites for intra- and extraperitoneal disease. Preoperative imaging is very helpful in detection of sites of potentially non-resectable disease or resectable disease, which is very helpful in the planning of debulking surgery. Early signs of peritoneal spread of

Correspondence address:

Prof. Binit Sureka, Diagnostic and Interventional Radiology, All India Institute of Medical Sciences, Basni Industrial Area, 342005, Jodhpur, India, e-mail: binitsurekapgi@gmail.com

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A Study design · B Data collection · C Statistical analysis · D Data interpretation · E Manuscript preparation · F Literature search · G Funds collection

ovarian malignancy may be missed if subtle deposits are not detected. Precise interpretation of ovarian malignancy spread on computed tomography is necessary for good quality reporting of ovarian malignancies for optimal management of disease.

Stage I	IA: Tumour limited to one ovary or fallopian tube, capsule intact, no tumour on surface, no tumour			
Tumour confined to ovaries	cells in ascites or washings			
	IB: Tumour involves both ovaries or fallopian tubes, otherwise like stage IA			
	IC: Tumour involves one or both ovaries with any of the following: capsule rupture, tumour on surface			
	of the ovary, positive peritoneal washings or ascites			
	IC1: Intraoperative spill			
	IC2: Capsule rupture before surgery or tumour on ovarian or fallopian tube surface			
	IC3: Malignant cells in the ascites or peritoneal washings			
Stage II	IIA: Extension and/or implant on uterus and/or fallopian tubes			
Tumour involves one or both ovaries with	IIB: Extension to other pelvic intraperitoneal tissues			
pelvic extension (below the pelvic brim)				
or primary peritoneal cancer				
Stage III	IIIA: Positive retroperitoneal lymph nodes and /or microscopic metastasis beyond the pelvis			
Tumour involves one or both ovaries with	IIIA1: Positive retroperitoneal lymph nodes only			
cytological or histologically confirmed spread	IIIA1(i) Metastasis \leq 10 mm			
to the peritoneum outside the pelvis and/or	IIIA1(ii) Metastasis > 10 mm			
metastasis to the retroperitoneal lymph nodes	IIIA2: Microscopic, extrapelvic (above the brim) peritoneal involvement \pm positive retroperitoneal lymph nodes			
	IIIB: Macroscopic, extrapelvic, peritoneal metastasis $\leq 2 \text{ cm} \pm \text{positive retroperitoneal lymph nodes}$.			
	Includes extension to capsule of liver/spleen			
	IIIC: Macroscopic, extrapelvic, peritoneal metastasis > 2 cm \pm positive retroperitoneal lymph nodes.			
	Includes extension to capsule of liver/spleen			
Stage IV	IVA: Pleural effusion with positive cytology			
Distant metastasis excluding peritoneal	IVB: Hepatic and/or splenic parenchymal metastasis, metastasis to extra-abdominal organs			
metastasis	(including inguinal lymph nodes and lymph nodes outside of the abdominal cavity)			

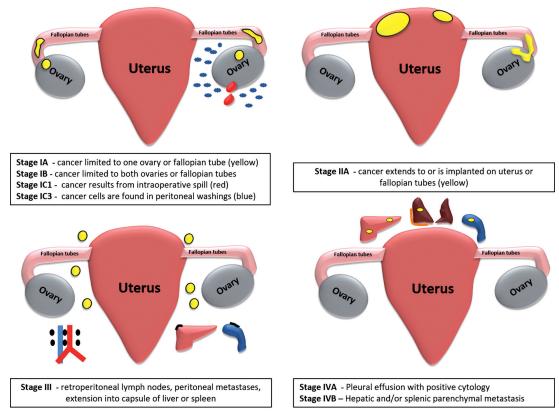


Figure 1. Drawing showing FIGO staging of ovarian cancer

Clinical and laboratory findings in ovarian cancer

Early stage ovarian cancers are usually asymptomatic. Late-stage ovarian cancers, which account for approximately 60% of cases, often have nonspecific symptoms, not recognised as symptoms of cancer [6]. Nonspecific symptoms of ovarian cancer are back pain, fatigue, abdominal pain/bloating, constipation, or urinary symptoms.

In addition to these nonspecific symptoms, ovarian cancer may present with paraneoplastic syndromes such as subacute cerebellar degeneration; sudden onset of seborrheic keratoses (Leser-Trélat sign); or unexplained spontaneous, recurrent, or migratory venous thrombotic events (Trousseau syndrome). Advanced disease may present with symptoms of regional spread or metastasis, such as bowel or ureteral obstruction, or shortness of breath [7,8]. Sex cord-stromal tumours can present early at stage I because of hormonal manifestations such as precocious puberty, abnormal uterine bleeding, and virilisation [9].

Cancer antigen (CA) 125 is the biomarker commonly tested for ovarian cancer. CA125 is elevated in about 80% of epithelial ovarian cancers overall, but in only 50% of early-stage epithelial ovarian cancers [8]. Biomarkers for nonepithelial ovarian cancers include inhibin A/B for sex cord-stromal tumours, and serum a-fetoprotein and quantitative beta human chorionic gonadotropin for germ cell tumours [10]. Human epididymal protein 4 (HE4) is a relatively new tumour marker. Elevated serum HE4 levels have been shown to have higher sensitivity (89%) and specificity (92%) than CA125 for distinguishing ovarian cancer from benign ovarian disease in premenopausal women, particularly in early-stage disease. As well as these biomarkers, complete blood count, blood chemistry including liver function tests and calcium (to assess for paraneoplastic syndromes), and serum biomarkers should be obtained if ovarian cancer is suspected.

Computed tomography technique

The optimal abdominal and pelvic computed tomography (CT) technique for staging ovarian cancer includes water as an oral contrast media, and in the absence of contraindications to intravenous contrast media, the use of intravenous contrast media is mandatory. Opacification of the gastrointestinal tract with oral contrast media helps to distinguish between luminal lesions and serous or mesenteric deposits; however, small, calcified deposits may be masked by oral contrast [11]. At our institution, image acquisition is done at 70 seconds after injection of contrast media. Acquisition for an abdominal and pelvic CT extends from the lower thorax to the inguinal regions with reconstruction of images with an axial slice thickness of 1-2 mm [10]. To include chest or not is an institutional policy. At our institute, we routinely scan the whole chest to look for lung metastases.

Routes of dissemination

Ovarian cancer usually spreads by direct infiltration into surrounding pelvic organs. Ovarian malignancies most commonly involve the adjacent adnexal structures like the uterus, fallopian tube, and contralateral ovary. Pelvic organs like the rectum, sigmoid colon, urinary bladder, and pelvic side wall may also be invaded. Spread of ovarian malignancy outside the pelvis occurs by intraperitoneal seeding, lymphatic transmission, or haematogenous spread.

Intraperitoneal seeding

Intraperitoneal seeding is most common mode of spread of ovarian malignancy, and during laparotomy 70% patients have peritoneal metastasis. The ovary is covered by single layer of epithelium and in epithelial ovarian malignancy, tumour cells are detached from the epithelium and spread in peritoneal fluid. Tumour cells initially shed into the dependent aspect of the rectouterine pouch and then move into lateral paravesical region because of gravity, and then move in a cranial direction with circulating peritoneal fluid through the right paracolic gutter to the hepatorenal pouch, right subphrenic space and then inferiorly into the inframesocolic compartment (Figure 2). The rectouterine pouch, right lower quadrant, sigmoid colon, and right paracolic gutter are the most common sites for peritoneal deposits due to gravity-dependent areas in the peritoneal cavity [12]. Peritoneal deposits are most commonly seen in the right paracolic gutter rather than the left due to the presence of the phrenicocolic ligament on the left side.

Lymphatic dissemination

Ovaries have three main drainage pathways – paraaortic and paracaval nodes are most common sites of lymph node metastasis, and they are drained by lymphatic vessels that follow the ovarian veins. The second most common site of lymph node metastasis is along the external iliac and obturator lymph nodes along the broad ligament (Figure 3) and inguinal lymph nodes along the round ligament [13].

Haematogeneous spread

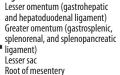
Haematogeneous spread is the least common mode of spread of ovarian malignancy. It often involves colon, liver, small intestine, and lung in decreasing order. Ovarian malignancy can rarely involve the spleen, kidney, and brain (Figure 4) [14].

Sites of tumour dissemination

Peritoneal involvement

Visible metastatic implants outside the pelvis and retroperitoneal lymphadenopathy are seen in 60% of epithelial

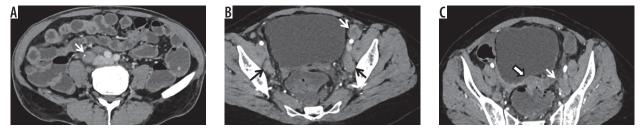
Ligaments



Lymph nodes Suprarenal (celiac axis, porta hepatis) Cardiophrenic

Pelvis •• Bony and pelvic side wall invasion Pelvic vessels encasement Bladder trigone involvement

Figure 2. A) Drawing showing the peritoneal space anatomy and pathways of spread of tumour cells into the peritoneal cavity. B) Coronal contrast-enhanced CT showing tumoural deposits into the peritoneal cavity along the serous surface, perihepatic space, paracolic spaces, ligaments, lymph nodes, and pelvis



B

surface

Paracolic gutters

Serous surfaces

cavity, bowel

Perihepatic space Gallbladder fossa

Falciform ligament

Porta hepatis

deposits

Liver, spleen, subphrenic

space, diaphragm, pleural

Bilateral paracolic spaces

Peritoneal and serous

Figure 3. Lymphatic dissemination in a case of ovarian cancer: A) Contrast-enhanced CT images showing enlarged lymph node (white arrow) in right common iliac region with calcific foci; B) bilateral internal iliac (black arrow) and left external iliac group (white arrow); C) obturator group (white arrow) with a deposit in rectouterine pouch (arrowhead)

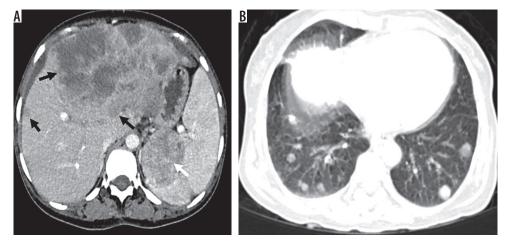


Figure 4. Haematogeneous spread from ovarian cancer: A) Contrast-enhanced CT images showing large metastatic deposit in liver (black arrow) and spleen (white arrow); B) and multiple metastatic nodules in bilateral lungs

ovarian malignancy [15]. Subtle thickening, nodularity, soft tissue in omental fat, peritoneal enhancement, and loculated ascites are subtle signs of peritoneal spread (Figure 5). Negative oral contrast is recommended for the detection of subtle serous deposits on the bowel surface.

Morphologically, there are three types of implants: solid, cystic, and mixed. In all three categories, calcifications may or may not be present. Different patterns of implants have been described:

- micronodular pattern characterised by the presence of tiny 1-5-mm nodules involving the serosa and subserosal fat;
- nodular pattern more than 5-mm-sized nodules, which are oval or round in shape with spiculated margins;

A

Diaphragmatic surface

TC

Rectouterine pouch

Greater omentum

Lesser sac

Liver surface

•••••

\.**.***

Paracolic gutters

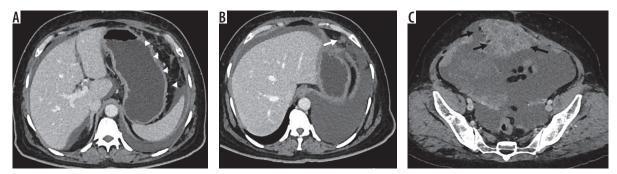


Figure 5. Contrast-enhanced computed tomography showing various types of peritoneal involvement in peritoneal carcinomatosis: A) subtle thickening and fat stranding; B) nodular pattern C); mass-like pattern with neoplastic ascites



Figure 6. Contrast-enhanced computed tomography image showing nodular deposits (arrowheads) in mesentery

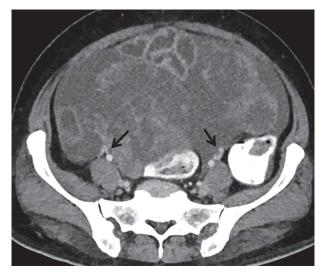


Figure 7. Pelvic involvement in ovarian cancer – contrast-enhanced computed tomography image showing tense ascites with plaque-like thickening of the peritoneum encasing the bilateral iliac vessels (arrows)

 plaque-like pattern – confluence of nodular implants giving a sheet-like pattern typically found in subdiaphragmatic spaces;

- mass-like pattern similar to plaque-like pattern, commonly found in the pelvis;
- omental cake diffuse involvement of greater omentum with fibrotic reaction leading to consolidation of omental fat;
- teca aspect or ileal freezing sleeve-like covering of small bowel loops by thickened a layer of visceral peritoneum with or without bowel obstruction;
- calcified implants nodular calcific deposits in peritoneum along with calcified lymph nodes, especially seen in serous ovarian adenocarcinoma;
- neoplastic ascites free fluid in the peritoneal cavity may be due to increased capillary permeability and fluid production or blockage of lymphatics leading to decreased absorption.

Mesentery

Implantation of tumour cells in small bowel mesentery is seen usually surrounding the terminal ileum. Morphological appearances of mesenteric involvement can be in the form of plaque-like thickening, reticulonodular lesions, subtle soft tissue haziness, or a mass-like deposit (Figure 6).

Lymph nodes

The presence of enlarged suprarenal lymph nodes at the level of the celiac axis, porta hepatis, and cardiophrenic lymph nodes makes the disease prone to suboptimal resection, and patients should go for neoadjuvant chemotherapy before surgery [16].

Pelvis

Pelvic side wall invasion should be suspected when the mass lesion is within 3 mm of the pelvic side wall, or surrounds or distorts more than 90% of the circumference of iliac vessels (Figure 7) [17].

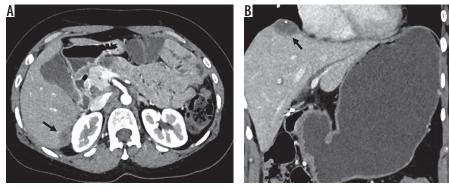


Figure 8. Subserosal deposits in ovarian cancer. Contrast-enhanced computed tomography images in two different patients showing subserosal implants in liver surface (arrow)

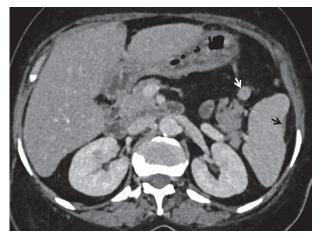


Figure 9. Subcapsular splenic deposit. Contrast-enhanced computed tomography image showing subcapsular splenic deposit (black arrow) and another nodular deposit in the gastrosplenic ligament (white arrow)

Visceral surfaces

Liver

The location, site, and size of liver implants should be documented in the report. Deposits in the region of Morrison's pouch, inferior vena cava, and around the right hepatic vein must be documented, because deposits in this region can cause excessive haemorrhage at the time of surgery. Secondary invasion of surface or subserosal implant (Figure 8) into the liver parenchyma should be well documented in the report [16].

Spleen

Subcapsular splenic deposits and parenchymal metastases have to be differentiated because this allows the surgeon to decide between splenectomy and spleen-sparing surgery (Figure 9) [17].

Subphrenic space

Right subphrenic involvement is very common because of preferential flow of peritoneal fluid from the right para-

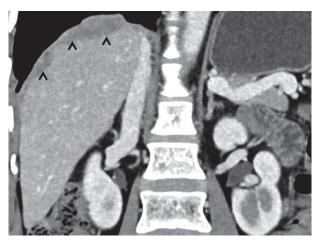


Figure 10. Subphrenic deposit in ovarian cancer. Coronal contrast-enhanced computed tomography image showing nodular thickening in right sub-phrenic region (arrowheads) indenting the liver surface

colic gutter. Involvement of the subphrenic region is best visualised in coronal and sagittal sections of post-contrast CT images. Imaging features of sub phrenic involvement include the presence of nodules, masses, thickening, and enhancement of the hemi diaphragm. Finding of subphrenic involvement must be written in radiologists' reports because sub phrenic involvements usually require upper-quadrant peritonectomy (Figure 10) [18,19].

Bowel

Magnetic resonance imaging (MRI) is superior to CT for assessment of bowel involvement in ovarian malignancies. Imaging findings of bowel involvement in ovarian malignancy include segmental or focal bowel wall thickening, focal nodules, serous infiltrations, or any definitive mass. Extensive bowel resections are not recommended due to functional complications (Figure 11) [20-22].

Pleural cavity

The presence of pleural effusion must be documented, although this finding is not specific unless tumour cells are detected in the fluid cytology.



Figure 11. Bowel involvement in ovarian cancer. Contrast-enhanced computed tomography image showing symmetrical nodular thickening (arrowheads) of the ileal loops in a case of ovarian cancer

Ligaments

Greater omentum

Imaging features of involvement of greater omentum range from infiltrative fat stranding to discrete and confluent masses forming omental cake. Involvement of transverse mesocolon also has to be commented upon because a modified surgical approach is required for the same.

Lesser omentum

The lesser omentum is a part of the peritoneum, which suspends the lesser curvature of the stomach, and it is composed of gastro-hepatic ligament and hepatoduodenal ligament. The left gastric artery, vein, and gastro-hepatic lymph nodes are contents of the gastro hepatic ligament, and it is recognised by identification of the left gastric artery on post-contrast CT images. The portal vein, common bile duct, hepatic artery, and portocaval lymph nodes are content of the hepatoduodenal ligament. Subtle soft tissues in the lesser omentum, and nodules and lymphadenopathy in the gastro hepatic and hepatoduodenal ligaments must be written in radiology reports. Involvement of the lesser omentum precludes surgery because of the complicated anatomy of the lesser omentum, which poses significant surgical challenges (Figure 12) [18,19].

Lesser sac

The lesser sac is a potential space between the stomach and pancreas. The lesser sac on the left side is outlined by the splenorenal ligament, gastrosplenic ligament, and splenopancreatic ligament. The lesser sac, on the right side, connects with the greater sac through the foramen of Winslow. Involvement of the lesser sac must be included in the radiological report because its involvement makes ovarian malignancy unresectable. Thickening, nodule, mass, fat stranding, and the presence of fluid in the lesser sac are considered to be signs of disease involvement (Figure 13).

Perihepatic space

The gallbladder fossa and fissure for the falciform ligament are the main components of the perihepatic space. Involvement of the falciform ligament should be looked for carefully in multiple planes to differentiate from liver parenchymal lesions. The falciform ligament, gallbladder fossa and periportal space should be carefully evaluated, because deposits more than 2 cm in size are predictors of non-optimal debulking (Figure 14) [23].

Differential diagnosis and computed tomography mimics of peritoneal carcinomatosis

Various pathologies involving the peritoneum can mimic peritoneal carcinomatosis. Table 2 summarises the mimics of peritoneal carcinomatosis and their imaging features [24-33].

Comparison between computed tomography and magnetic resonance imaging in the detection of peritoneal implants

CT is the imaging modality of choice for staging ovarian cancer. However, CT is limited in detecting small perito-

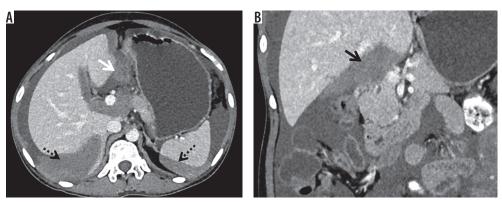


Figure 12. Lesser omentum involvement in ovarian cancer. Axial and coronal CT images showing soft tissue deposits in gastrohepatic ligament (white arrow) and in hepatoduodenal ligament (black arrow) of lesser omentum region. Perisplenic and perihepatic enhancing deposits (dashed arrows) with neoplastic ascites also evident

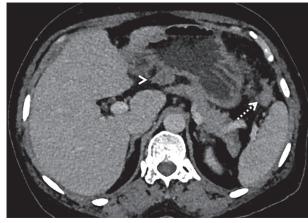


Figure 13. Lesser sac deposit in ovarian cancer. Contrast-enhanced computed tomography image showing metastatic deposits in between pancreas and stomach (arrowhead) and in gastrosplenic ligament (dashed arrow)

neal metastases. The sensitivity and specificity of CT in detecting tumour implants larger than 1 cm range from 85-93% and 91-96%, respectively, but the sensitivity decreases to 25-50% in detecting implants that are 1 cm or smaller. MRI combining delayed gadolinium-enhanced imaging and diffusion-weighted MR imaging (DWM-RI) is more accurate than CT for estimating peritoneal tumour burden [34]. Low *et al.* [35] have shown that MRI is superior to CT in estimating peritoneal cancer index, with sensitivity 95%, specificity 70%, and accuracy 88%,

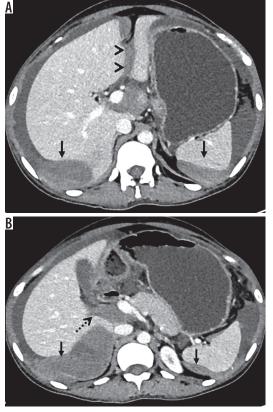


Figure 14. Perihepatic deposits in ovarian cancer. Axial contrast-enhanced computed tomography images showing: A) deposits in the falciform ligament (arrowheads), periportal region; B) around gallbladder fossa (dashed arrow) with perihepatic and perisplenic enhancing deposits (arrows)

Table D Differential	استنبع مشمع منتجا		A			
Table 2. Differential	diadnosis and	computed	tomodraphy	mimics of	peritoneai	carcinomatosis
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Carcinomatosis mimics	Imaging features
Tuberculous peritonitis	Three types — wet, fixed fibrotic, and dry plastic types. Young age, presence of free or loculated ascites, necrotic lymph nodes, matted bowel loops, and smooth peritoneal thickening confirms the diagnosis [24]
Peritoneal Iymphomatosis	Associated findings of splenomegaly, aneurysmal dilatation of bowel loop segment, confluent homogeneously enhancing non-necrotic lymph nodes encasing the vessels producing a 'sandwich sign' [25]
Primary peritoneal mesothelioma	Older age of presentation, strong association of asbestos exposure, two imaging patterns – nodular mass like large confluent masses and diffuse nodular thickening of peritoneum, no calcification, ascites (loculated or diffuse) with omental caking [26]
Pseudomyxoma peritonei	Gelatinous ascites causing classical scalloping of the surface of abdominal organs [27]
Peritoneal sarcomatosis	Associated with gastrointestinal stromal tumour, liposarcoma, and leiomyosarcoma. Distinguishing imaging features include heterogeneous bulky masses, hypervascularity with or without hemoperitoneum and variable presence of ascites [28]
Desmoplastic small round cell tumour	Rare aggressive malignancy that occurs in adolescents and young adults, mostly male, and carries a poor prognosis. Imaging features are characterised by single or multiple peritoneal masses in the omentum or paravesical region, which spread later to the entire abdominal cavity [29]
Splenosis	Spontaneous auto-transplantation of splenic tissue after traumatic injury or splenectomy, ^{99m} Tcechnetium-labelled RBC scintigraphy with SPECT/CT imaging is the modality of choice for diagnosis [30]
Melanosis	Unusual condition characterised by either focal or diffuse brown or black pigmentation of the peritoneum, usually surgico-pathological diagnosis; associated with various causes like metastatic melanoma, mature ovarian cystic teratoma, ovarian serous cystadenoma, enteric duplication cyst, peritoneal cyst [31]
Sclerosing peritonitis	Chronic inflammatory condition of the peritoneum with an unknown aetiology, eventually progresses to sclerosis and membrane formation with subsequent cocoon formation. Imaging features are central accumulation of the small intestine encased by a dense membrane, intestinal obstruction, ascites, calcification bowel serous surface [32]
Omental infarct	Fatty, large (> 5 cm) encapsulated mass, with soft-tissue stranding adjacent to the ascending colon [33]
Fat saponification	Sequelae of pancreatitis resulting in mesenteric and retroperitoneal fat necrosis. Imaging shows scattered tiny soft tissue density nodules and fluid in vicinity of inflammation [33]

Potentially nonresectable disease Extensive involvement of small bowel or mesenteric root Lymph node above celiac axis Pleural involvement Pelvic sidewall invasion Bladder trigone involvement Intrahepatic metastases Metastatic implants near right hepatic vein Implants > 2 cm in diameter in diaphragm, lesser sac, porta hepatis, gallbladder fossa, falciform ligament, gastrosplenic or gastrohepatic ligament



Resectable disease that requires surgical subspecialty expertise Subcapsular implants with hepatic parenchymal invasion Limited hepatic parenchymal metastases Bladder or ureteral invasion Bowel serous deposits

Figure 15. Diagram showing resectable and non-resectable parameters of ovarian cancer

compared to CT, which had sensitivity 55%, specificity 86%, and accuracy 63%. DWMRI with a high b-value (e.g. 800 s/mm²) increases the specificity in peritoneal carcinomatosis detection because of better suppression of the bowel contents, and the combination of DW-MRI and contrast-enhanced MRI further improve the detection accuracy [36]. For treatment response evaluation, functional imaging with DWMRI or PET-CT is recommended. Postoperative inflammation and oedema would result in high ADC, in contrast to tumours that have low ADC, allowing for differentiation of disease recurrence from postoperative fibrosis and inflammation. DWMRI is superior to PET-CT in detecting perihepatic and serous deposits because of physiological uptake in the liver and bowel and respiratory motion interference [37]. PET-CT is a helpful imaging adjunct in the presence of rising tumour markers but equivocal CT or MRI, in detecting disease recurrence in ovarian cancer [38].

Conclusions

Accurate peritoneal anatomy and knowledge of imaging features of peritoneal carcinomatosis is essential for appropriate management and care of patents with ovarian cancer. It is important to be aware of the various resectable and nonresectable parameters of ovarian cancer (Figure 15). CT is the most widely used imaging modality for assessment; however, MR with advanced functional tools can be helpful in assessing the tumour response to treatment.

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Conflict of interest

The authors report no conflicts of interest.

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