










# Imaging and Clinical Findings of Xanthogranulomatous Inflammatory Disease of Various Abdominal and Pelvic Organs: A Pictorial Essay

복부와 골반의 다양한 장기에서 발생한 황색육아종성  
염증 질환의 영상 및 임상 소견: 임상화보

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Xanthogranulomatous (XG) inflammatory disease is a rare benign disease involving various organs, including the gallbladder, bile duct, pancreas, spleen, stomach, small bowel, colon, appendix, kidney, adrenal gland, urachus, urinary bladder, retroperitoneum, and female genital organs. The imaging features of XG inflammatory disease are nonspecific, usually presenting as a heterogeneous solid or cystic mass. The disease may also extend to adjacent structures. Due to its aggressive nature, it is occasionally misdiagnosed as a malignant neoplasm. Herein, we review the radiological features and clinical manifestations of XG inflammatory diseases in various organs of the abdomen and pelvis.

**Index terms** Abdomen; Pelvis; Xanthogranulomatous Cholecystitis;  
Xanthogranulomatous Pyelonephritis; Ultrasonography;  
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## INTRODUCTION

Xanthogranulomatous (XG) inflammatory disease is a rare and aggressive form of chronic inflammation. It is pathologically characterized by soft yellow nodules composed of lipid-laden foamy macrophages, histiocytes, and plasma cells, with a variable number of foreign body giant cells (1). In XG inflammation, the organ tissue is eventually replaced by abundant xanthoma cells, granuloma formation, and dense proliferation of fibrin and collagen deposition, resulting in tissue destruction that extends into the adjacent structures (2).

Although various mechanisms have been proposed, including defective lipid transport, immunological disorders, infection caused by low-virulence organisms, reactions to specific infectious agents, and lymphatic obstruction (3), the precise pathogenesis of XG inflammation has not been elucidated.

XG inflammation is commonly seen in cholecystitis and pyelonephritis; however, it can also occur in other organs, including the bile duct, pancreas, spleen, stomach, small bowel, colon, appendix, adrenal gland, urachus, urinary bladder, retroperitoneum, and female genital organs (4). Owing to their aggressive nature, the clinical and radiologic findings of XG inflammatory diseases are often misdiagnosed as malignant neoplasms. Therefore, accurate diagnosis is essential for proper patient management and surgical planning. Imaging plays a substantial role in the diagnosis and assessment of disease progression and its associated complications (4).

Herein, we review the radiological features and clinical manifestations of XG inflammatory disease in various abdominal and pelvic organs.

## XANTHOGRANULOMATOUS CHOLECYSTITIS

XG cholecystitis is a rare chronic inflammatory disease characterized by grayish-yellow nodules or streaks in the gallbladder wall (5). It commonly occurs in women aged 60–70 years. Patients with XG cholecystitis present with various signs and symptoms of cholecystitis, including right upper quadrant pain, vomiting, jaundice, leukocytosis, and a positive Murphy's sign (5). The pathogenesis of XG cholecystitis is unclear; however, one theory suggests that it may occur due to extravasation of bile into the gallbladder wall (with the involvement of the Rokitansky–Aschoff sinuses), or through a small ulceration in the mucosa (5, 6).

Sonographic features of XG cholecystitis includes diffuse wall thickening with hypoechoic intramural nodules or a hypoechoic band surrounding the gallbladder (Fig. 1) (5, 7). Hypoechoic nodules indicate abscesses and xanthogranulomas, alone or in combination (8). Similarly, CT and MRI can reveal intramural nodules, thickened gallbladder walls, gallstones, and ductal dilatation (Fig. 2) (5-7). Intramural nodules are hypoattenuated on CT and show T2 hyperintensity and signal drops on opposed phase MRI (5-7). Complications of XG cholecystitis include gallbladder perforation, abscess formation, fistulous tract formation to the duodenum or skin, liver or colon invasion, and Mirizzi syndrome (4).

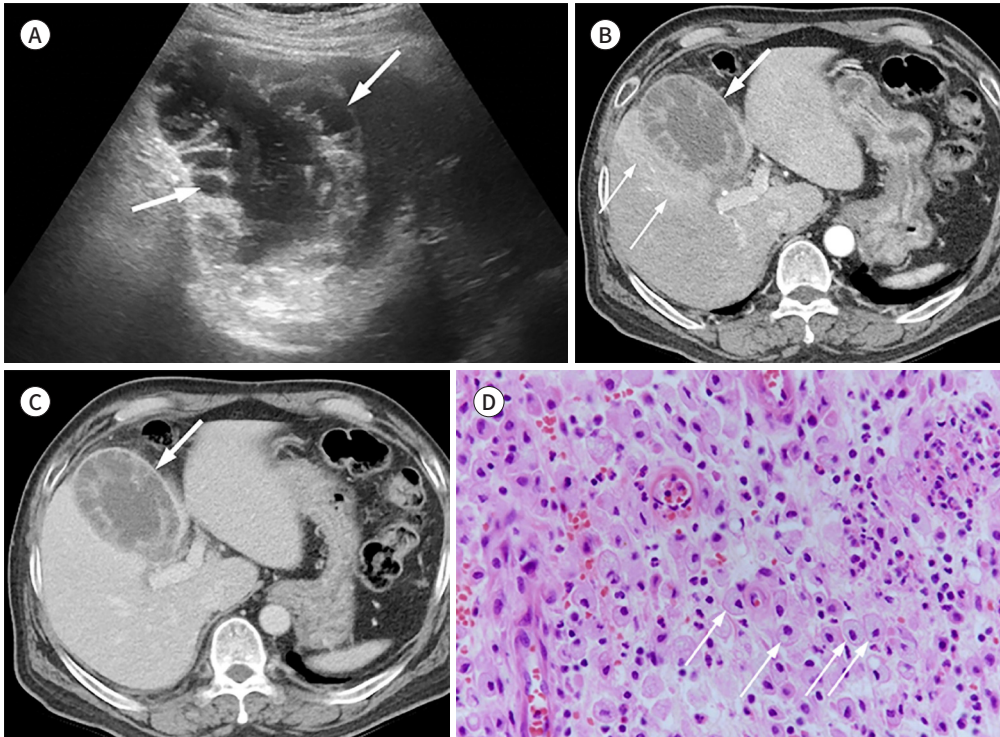
Clinical and radiological findings of XG cholecystitis often resemble those of gallbladder cancer. Several studies have attempted to determine the characteristic imaging features that distinguish XG cholecystitis from gallbladder cancer. The following features significantly in-

**Fig. 1.** A 64-year-old male with xanthogranulomatous cholecystitis.

**A.** Transverse sonography shows diffuse wall thickening of gallbladder with hypoechoic intramural nodules (arrows) and sludges in gallbladder lumen.

**B, C.** Arterial (**B**) and portal (**C**) phases CT reveal diffuse wall thickening of gallbladder with multiple intramural hypodense nodules (arrows) along with transient arterial enhancement of liver parenchyma adjacent to the gallbladder (thin arrows in **B**).

**D.** Photomicrography shows diffuse sheets of foamy lipid-laden macrophages (arrows) and lymphocytes (hematoxylin and eosin,  $\times 400$ ).



dicating XG cholecystitis over malignancy: diffuse gallbladder wall thickening, continuous mucosal line, intramural nodules, gallstones, pericholecystic fat stranding, transient hepatic attenuation differences, absence of intrahepatic bile duct dilatation, and absence of hepatic invasion (6, 7). Intramural nodules corresponding to xanthoma deposition are key features for differentiating XG cholecystitis from malignancies (5).

## XANTHOGRANULOMATOUS CHOLANGITIS

XG cholangitis is a rare disease, with only 14 reported cases (9). Although it usually develops because of the extension of inflammatory lesions from XG cholecystitis into the bile ducts (10), cases of isolated XG cholangitis have been reported (9). The most common symptoms of XG cholangitis include abdominal pain and jaundice. However, its pathogenesis remains unclear. It occurs secondary to recurrent cholangitis and leads to chronic bile duct inflammation, scarring, and XG reaction (9).

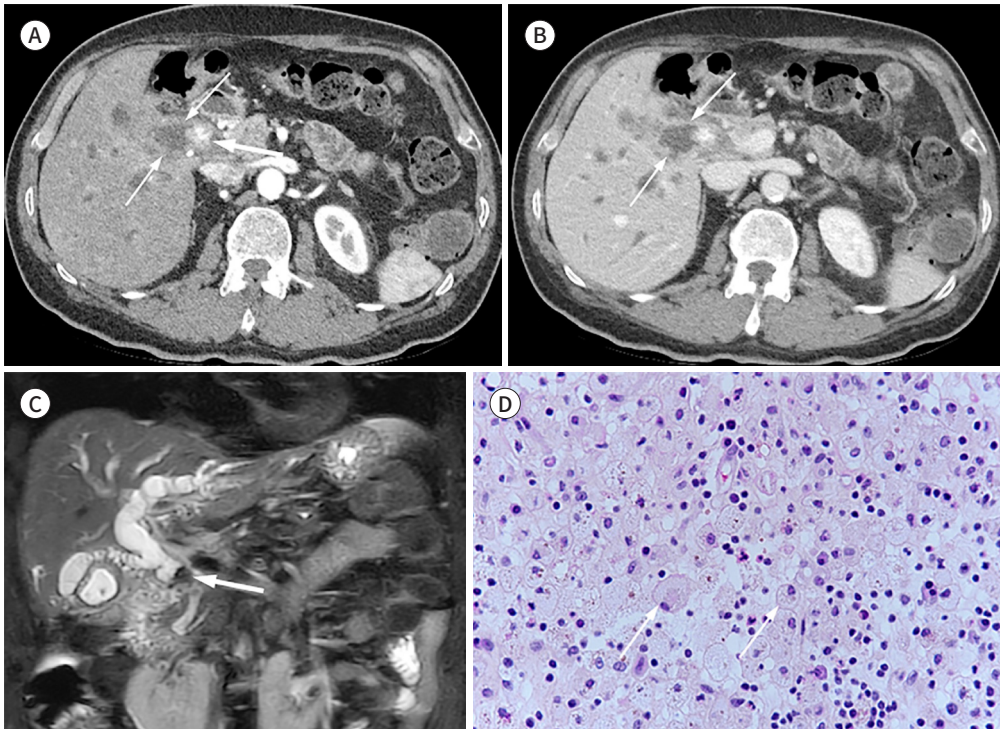
Although there are few reports of XG cholangitis imaging features, bile duct stricture with associated wall thickening is a known finding (11, 12). In our case, XG cholecystitis was diagnosed concurrent with XG cholangitis. CT and MRI revealed a stone in the dilated extrahe-

Fig. 2. A 70-year-old male with xanthogranulomatous cholangitis.

A, B. Arterial (A) and portal (B) phases CT show wall thickening of dilated extrahepatic bile duct (thin arrows) with a stone (arrow). Intrahepatic bile ducts are dilated.

C. Coronal T2-weighted image shows a hypointense stone in dilated extrahepatic bile duct (arrow). Diffuse wall thickening of gallbladder with tiny stones is seen. Intrahepatic bile ducts are dilated.

D. Photomicrography shows numerous foamy lipid-laden macrophages (arrows), lymphocytes, and plasma cells (hematoxylin and eosin,  $\times 400$ ).



patric bile duct, bile duct wall thickening, and intrahepatic bile duct dilatation (Fig. 2). XG cholangitis can mimic cholangiocarcinoma, primary sclerosing cholangitis, recurrent pyogenic cholangitis, inflammatory pseudotumors, and Mirizzi syndrome (12).

## XANTHOGANULOMATOUS PNCREATITIS

XG pancreatitis is a rare form of chronic pancreatitis. The exact pathophysiology of XG pancreatitis remains unclear; however, it typically results from a combination of duct obstruction, infection, and repeated hemorrhages. Moreover, it is occasionally associated with pancreatic neoplasms (13, 14). XG pancreatitis commonly occurs in middle-aged men and presents with abdominal pain (15). Laboratory findings, including WBC count, serum C-reactive protein, amylase, lipase, and CA19-9 are mostly within the normal range or clinically insignificant (14).

There are various CT and MRI indications for XG pancreatitis, ranging from a cystic mass to a solid mass which can mimic pancreatic neoplasms (14). The solid type of XG pancreatitis presents a lobulated heterogeneous mass with hypovascular and progressive enhancement (Fig. 3) (14). Differentiating XG pancreatitis from a malignant tumor is difficult; however, in patients with XG pancreatitis, upstream pancreatic duct dilatation is generally absent, and

**Fig. 3.** A 68-year-old female with xanthogranulomatous pancreatitis.

**A, B.** Axial enhanced CT reveal a heterogeneously hypodense lesion with irregular contour (arrows), which is protruded from the pancreatic tail.

**C.** Gross specimen of distal pancreas shows a capsular lesion filled with mud-like contents (arrows).



CA19-9 levels are within the normal range (14). The cystic type of XG pancreatitis presents as a lobulated cystic mass with an irregular thick wall, similar to intraductal papillary mucinous neoplasms or other pancreatic cystic tumors (13, 14). MRI findings can help distinguish the cystic type from other cystic tumors. In XG pancreatitis, T2-weighted imaging usually demonstrates a low signal intensity relative to the fluid as well as obvious diffusion restriction (13, 14).

## XANTHOGRANULOMATOUS INFLAMMATION OF THE GASTROINTESTINAL TRACT

The involvement of the gastrointestinal tract in XG inflammation is rare, with only 14 cases reportedly affecting the stomach (16, 17), 2 affecting the terminal ileum (18, 19), and 5 affecting the colon (20, 21). Of the five cases with colonic involvement, two involved the sigmoid colon, two involved the ascending colon, and one involved the cecum (20, 21).

Most gastric lesions present as mass-forming lesions, with submucosal involvement or wall thickening (16). Cases of XG gastritis may be misdiagnosed preoperatively as submucosal tumors or advanced gastric cancer (16). XG inflammation of the terminal ileum presents with mucosal ulceration, wall thickening (18), or mass formation (19), whereas XG colitis presents as a submucosal (21) or ulceroinfiltrative (20) mass. Symptoms of XG colitis are abdominal pain, constipation, and diarrhea (20, 21). XG inflammation of the gastrointestinal tract is difficult to distinguish from other malignancies. Therefore, intraoperative pathological diagnosis using frozen sections may be required to plan the surgical extent (20).

## XANTHOGRANULOMATOUS APPENDICITIS

XG appendicitis is a rare chronic inflammatory disease of the appendix and is associated with long-standing or recurrent inflammation (22). XG appendicitis occurs in patients who undergo interval appendectomy after perforation of acute appendicitis (23). Although several studies have been published in pathology journals, imaging findings have rarely been described (24-26). All imaging findings reported were obtained using CT (24-26). Previously reported CT findings of XG appendicitis have only been described as secondary findings of ap-

pendicitis, such as diffuse bowel wall thickening of the terminal ileum and cecum (24), pericecal abscesses (25), or large ileocecal phlegmon (26). In addition, these reports did not provide CT findings of XG appendicitis in the appendix itself.

In our case, the CT findings of XG appendicitis demonstrated severe asymmetric appendiceal wall thickening and scant intraluminal fluid (Fig. 4). This result correlates with the pathology of mural fibrosis, chronic transmural inflammation with lymphoid aggregates, and subserosal spread.

## XANTHOGRANULOMATOUS INFLAMMATION OF THE PERITONEUM

To the best of our knowledge, XG inflammation of the peritoneum is rare, and only one case has been reported (27), with CT findings of an approximately 1.7 cm-sized omental nodule and multiple peritoneal nodules. Bile leakage after laparoscopic cholecystectomy is considered a cause of XG inflammation of the peritoneum (27).

In our case, a 1.2 cm-sized heterogeneous enhancing soft tissue mass in the peritoneal cavity of the epigastric portion was observed on CT of a 77-year-old man with a history of cholecystectomy for gallbladder cancer (Fig. 5). The mass was located near the surgical incision

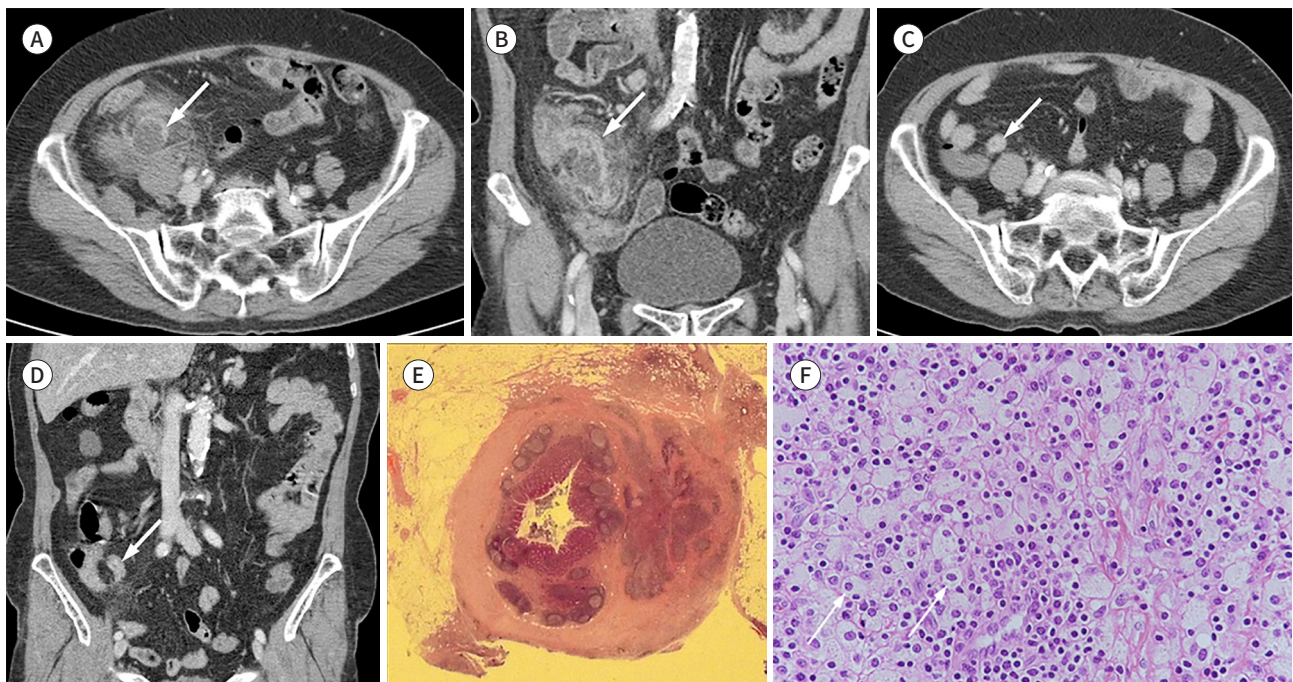
**Fig. 4.** A 69-year-old male with xanthogranulomatous appendicitis.

**A, B.** Axial (**A**) and coronal (**B**) enhanced CT show perforated appendicitis with periappendiceal abscess formation (arrows). Periappendiceal fat infiltration is noted.

**C, D.** Follow-up enhanced CT after 40 days shows diffuse wall thickening of appendix (arrows). Periappendiceal abscess and periappendiceal fat infiltration are disappeared.

**E.** Low-power photography reveals diffuse mural fibrosis and thickening and transmural chronic inflammation with lymphoid aggregates of appendix.

**F.** Photomicrography shows numerous foamy lipid-laden macrophages (arrows), lymphocytes, and plasma cells (hematoxylin and eosin,  $\times 400$ ).



site on the abdominal wall. We believe that the XG inflammation of the peritoneum in our case may have been related to surgery. XG inflammation in the peritoneum was confirmed by surgical resection.

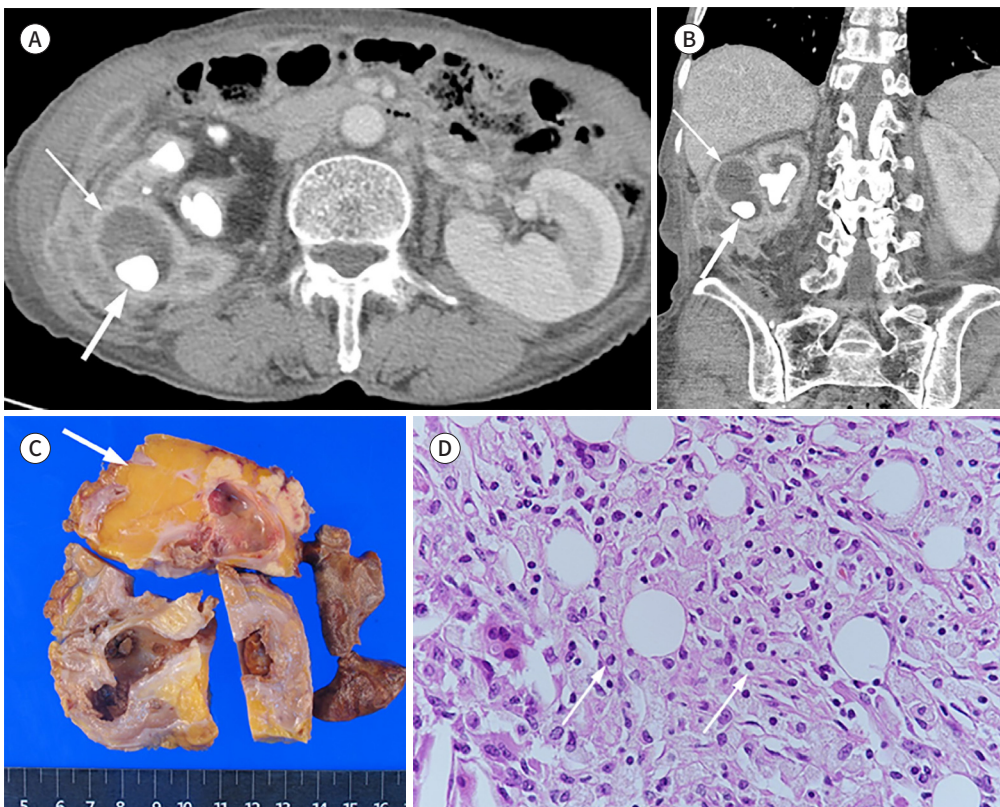
**Fig. 5.** A 77-year-old female with xanthogranulomatous inflammation of the peritoneum and a history of cholecystectomy for gallbladder carcinoma 4 months ago.

- A.** Axial enhanced CT shows a small hypodense nodule at peritoneum of the epigastric portion (arrow) and perilesional fat infiltration.
- B.** After 6 months, the nodule became smaller and more contrast enhanced on enhanced CT (arrow).
- C.** PET CT shows a focal hypermetabolic lesion in anterior abdominal cavity (arrow).



**Fig. 6.** A 70-year-old female with xanthogranulomatous pyelonephritis.

- A, B.** Axial (A) and coronal (B) enhanced CT show calyceal dilatation, staghorn calculi (thick arrows), and parenchymal thinning (thin arrows).
- C.** Excised specimen shows staghorn calculi and xanthogranulomatous inflammation represented by irregular yellow tissue in collecting system (arrow).
- D.** Photomicrography shows diffuse sheets of foamy lipid-laden macrophages (arrows) and inflammatory cells (hematoxylin and eosin,  $\times 400$ ).



## XANTHOGRANULOMATOUS PYELONEPHRITIS

XG pyelonephritis (XGP) is an uncommon but severe form of chronic renal infection. It commonly occurs in middle-aged patients with a history of recurrent urinary tract infections, diabetes, or kidney stones (28). The clinical features are nonspecific, with flank pain and fever being the most common symptoms (29). XGP is a chronic granulomatous disease in which the renal parenchyma is destroyed and ultimately replaced with lipid-laden (foamy) macrophages, often extending beyond the kidneys (2).

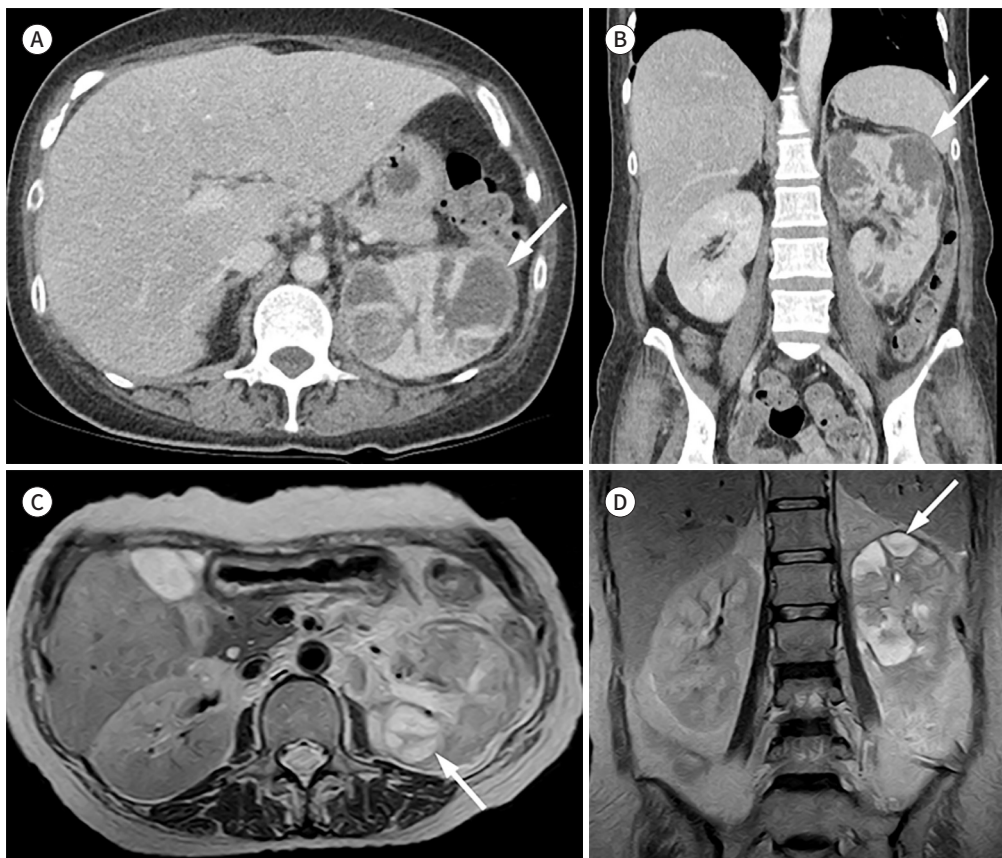
Sonography typically shows diffuse enlargement of the kidney, with renal parenchymal thinning or loss of normal corticomedullary differentiation (29). Multiple anechoic or hypoechoic masses similar to the “bear paw sign” on CT can be observed (30). Central echogenicity with posterior acoustic shadowing indicates pelvic staghorn calculus (30). XGP is focal in 10% of cases and can mimic renal tumors (2).

CT demonstrates renal enlargement and replacement of the renal parenchyma by multiple low-attenuation blown-out calyces with cortical thinning, representing a “bear paw sign” (Fig. 6) (30, 31). Low-attenuation calyces show either dilated calyces or renal parenchyma filled with

**Fig. 7.** A 48-year-old female with xanthogranulomatous pyelonephritis.

**A, B.** Axial (**A**) and coronal (**B**) enhanced CT reveal multiple ill-defined hypodense lesions in left kidney (arrows).

**C, D.** Axial (**C**) and coronal (**D**) T2-weighted images show multiple hyperintense masses with cortical thinning of left kidney (arrows).





pus, debris, or hemorrhage (31). A large staghorn calculus with contraction of the renal pelvis can also be seen (29).

The MRI findings are nonspecific and overlap with those of tuberculosis and malignancy. Lipid-laden foamy macrophages are hyperintense on T1-weighted images (28). Cavities and abscesses are hypointense on T1-weighted images and hyperintense on T2-weighted images (Fig. 7) (30).

Complications associated with XGP include perforation with an abscess and/or fistula formation involving the spleen, paraspinal muscles, or psoas muscle (2, 28).

## XANTHOGRANULOMATOUS INFLAMMATION OF THE URACHUS AND URINARY BLADDER

The urachus is an embryonic fibrous remnant of the allantois that extends from the apex of the bladder to the umbilicus. Involution of the urachus is usually complete at birth; however, incomplete involution results in urachal abnormalities such as patent urachus, umbilical-urachal sinus, vesicourachal diverticulum, and urachal cysts (32). Few cases of XG inflammation in the urachus have been reported (33-35). The main symptoms include lower abdominal mass, abdominal pain, and dysuria (35).

Sonography and CT reveal a midline supravescical soft tissue mass with adjacent organ involvement. On sonography, it usually appears as a mixed echogenic mass in the anterosuperior portion of the bladder (Fig. 8) (33). CT demonstrates heterogeneous attenuation with variable contrast enhancement (33) or peripheral rim enhancement (Figs. 8, 9) (34). The differential diagnosis of urachal carcinoma based on imaging is difficult; however, a supravescical mass with calcification would significantly favor urachal carcinoma over inflammation (32).

XG cystitis is a rare, benign, chronic inflammatory disease; to date, 30 cases have been reported in the literature (in English) (36). However, a majority of the reported cases were associated with urachal cysts or remnants, and the location of XG cystitis in these cases was at the dome of the bladder (Fig. 9) (37).

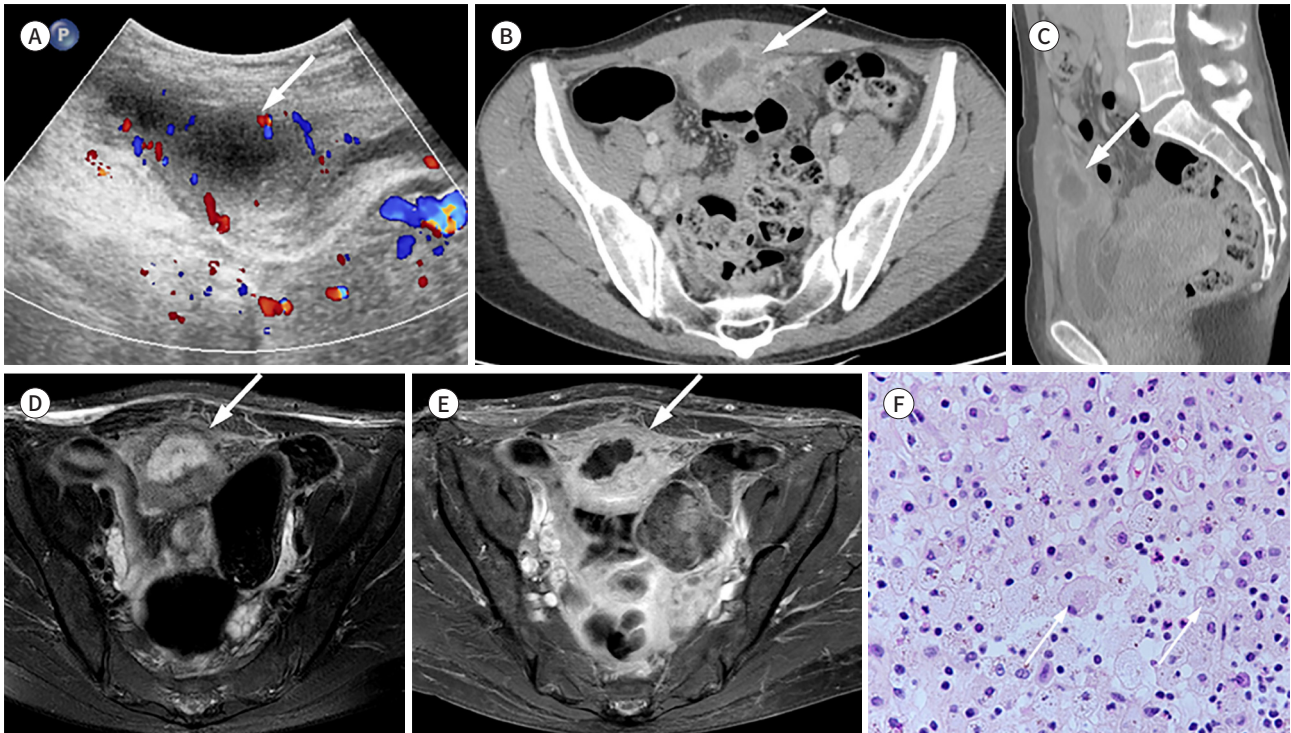
XG cystitis without a relationship with the urachus is very rare, with only four cases demonstrating this condition (36, 38). Common symptoms of XG cystitis without a relationship with the urachus include lower abdominal pain and dysuria; it may also be asymptomatic (36, 38). Imaging findings of XG cystitis without a relationship to the urachus include a polypoid mass (36) and wall thickening of the urinary bladder (38). XG cystitis is difficult to distinguish from actinomycosis, which is another chronic inflammation. However, actinomycosis usually occurs in women and is associated with intrauterine device insertion, which can be helpful in the differential diagnosis (39).

## XANTHOGRANULOMATOUS PROSTATITIS

XG prostatitis is a rare form of granulomatous prostatitis that can mimic prostate cancer clinically and radiologically (40-42). Patients with clinical presentation of symptoms are typically in their 60s. It presents with lower urinary tract infections, such as frequency and dysuria (40-42). Prostate-specific antigen levels are typically elevated (41, 42).

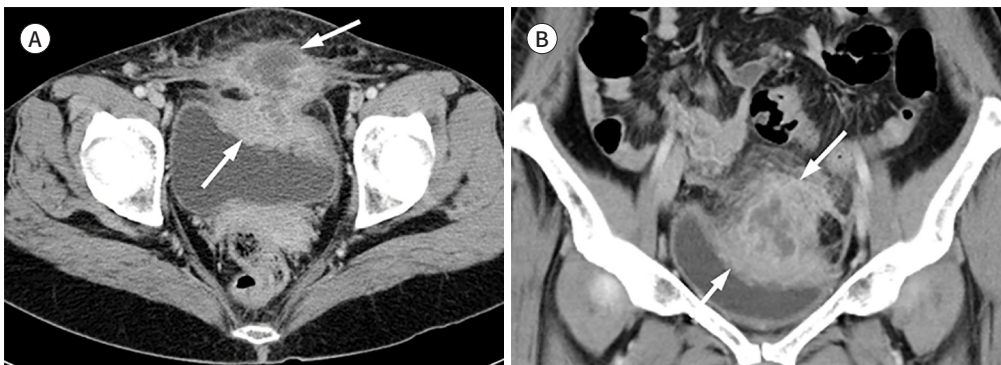
**Fig. 8.** A 31-year-old female with xanthogranulomatous inflammation of urachus.

- A.** Sagittal color Doppler sonography shows a tear-drop shaped hypoechoic mass with blood flow within the peripheral portion in suprav vesical space (arrow).  
**B, C.** On axial (**B**) and sagittal (**C**) enhanced CT show an enhancing soft tissue mass with central cystic focus at suprav vesical space (arrows).  
**D.** On T2-weighted MR image, the mass shows intermediate signal intensity with central cystic focus (arrow).  
**E.** On contrast enhanced T1-weighted image, the mass shows homogeneous enhancement with central cystic lesion (arrow).  
**F.** Photomicrography shows diffuse sheets of foamy lipid laden macrophages (arrows) and lymphocytes (hematoxylin and eosin,  $\times 400$ ).



**Fig. 9.** A 42-year-old female with xanthogranulomatous inflammation of the urinary bladder and urachus.

- A, B.** Axial (**A**) and coronal (**B**) enhanced CT reveal enhancing soft tissue mass with central cystic portion at the urinary bladder dome and suprav vesical space (arrows).



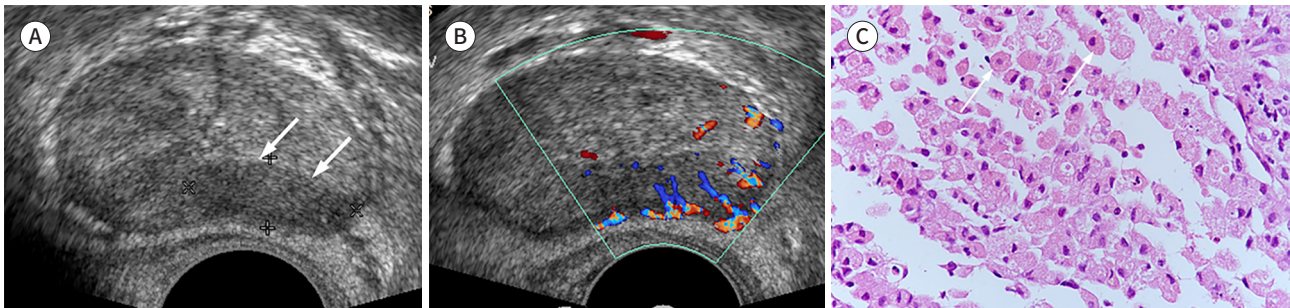
Endorectal sonography typically shows uniform hypoechoic (43) or multiple hypoechoic lesions (Fig. 10) (40). Few cases concerning MRI features have been reported (41, 44). MRI shows T1 hypointensity and T2 hyperintensity in the bilateral peripheral zones, with peripheral enhancement and diffusion restriction (41). Prostatic abscesses are rare in patients with XG prostatitis (45). On sonography and MRI, differentiation between XG prostatitis and pros-

**Fig. 10.** A 69-year-old male with xanthogranulomatous prostatitis.

**A.** Axial sonography shows a hypoechoic lesion in left peripheral zone of the prostate (arrows).

**B.** The mass shows hypervascularity on color Doppler sonography.

**C.** Photomicrography shows diffuse sheets of foamy lipid-laden macrophages (arrows) and lymphocytes (hematoxylin and eosin,  $\times 400$ ).



tate cancer may be difficult (40-42). Hence, a transrectal sonography-guided prostate biopsy should be performed for an accurate diagnosis (40). Although XG prostatitis does not require additional treatment, follow-up is necessary (41).

## XANTHOGANULOMATOUS INFLAMMATION OF THE FEMALE GENITAL TRACT

XG inflammation of the female genital tract is an uncommon chronic inflammation that causes destructive masses in the pelvic cavity. It affects the endometrium, fallopian tubes, or ovaries focally or entirely and is most commonly seen in the endometrium (46).

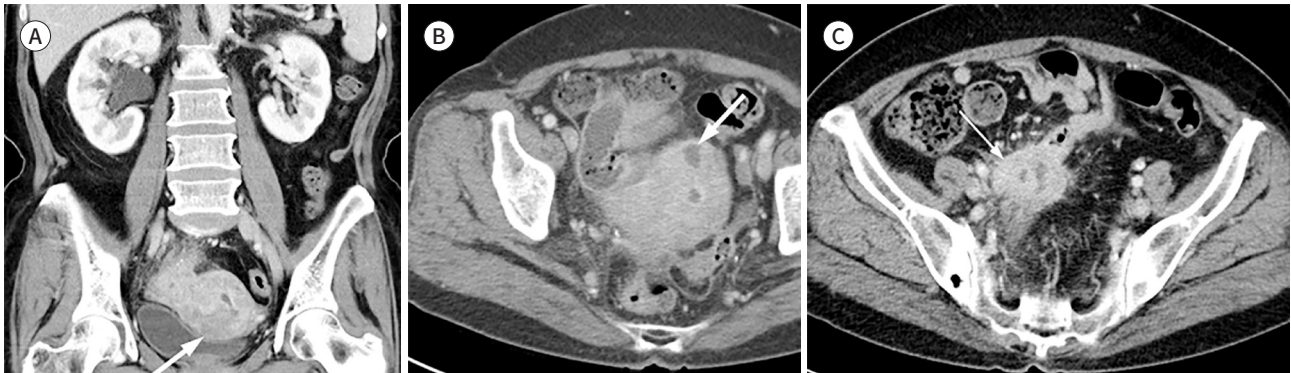
XG endometritis occurs in women aged 59–88 years (average: 72 years) (46). Postmenopausal women with endometrial hyperplasia, endometrial carcinoma, and/or cervical stenosis can develop this condition (47). Clinical manifestations include lower abdominal pain and vaginal discharge. Radiological findings usually include a heterogeneous cystic uterine mass (48) or chronic complex fluid collection in the endometrial canal (47). XG inflammation rarely involves the myometrium and extends to the pelvis with uterine perforation (Fig. 11) (49). Radiological features can easily lead to misdiagnosis of malignant tumors, and XG endometritis may be associated with endometrial cancer. Surgical excision is the standard treatment for XG endometritis (48).

XG salpingitis and oophoritis typically occur in women aged 23–72 years (average: 38.5 years) and mainly affect one side (46). Patients with XG inflammation of the fallopian tube and ovary typically present with a history of pelvic inflammatory diseases that last for months or years. Other clinical presentations include anemia, anorexia, fever, irregular colporrhagia, abdominal pain, and tenderness of the adnexal mass (46).

Sonography, CT, and MRI revealed a thick-walled complex solid cystic tubo-ovarian mass with variably enhanced solid portions (Fig. 12) (50). It usually appears as an invasion of the surrounding tissue and may be difficult to differentiate from actinomycosis (51) or ovarian malignancies (50). On MRI, the presence of non-enhancing nodules in the thickened walls of ovarian cystic masses may be a unique finding to confirm the diagnosis of XG oophoritis (52).

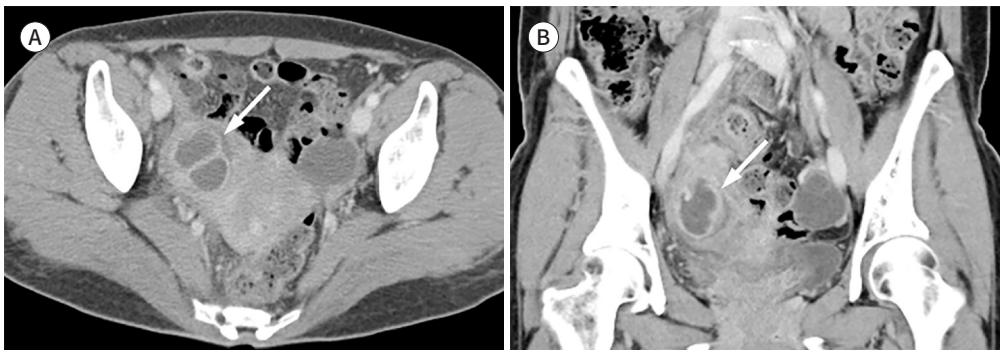
**Fig. 11.** A 59-year-old female with xanthogranulomatous inflammation of the uterus, right salpinx, and right ovary.

**A-C.** Axial (**A, B**) and coronal (**C**) enhanced CT show irregular hypodense abscesses in the uterine body (thick arrows), right salpinx, and right ovary (thin arrow). Multiple fat infiltrations are seen adjacent to the inflamed right ovary.



**Fig. 12.** A 44-year-old female with xanthogranulomatous salpingitis.

**A, B.** Axial (**A**) and coronal (**B**) enhanced CT show a cystic mass with a thick wall and a septum in the right adnexa (arrows). Right salpingectomy, omentectomy, and adhesiolysis were performed. Laparoscopic right salpingectomy was performed. Pathologic diagnosis of xanthogranulomatous salpingitis was made.



## CONCLUSION

XG is a rare and benign inflammatory disease involving various organs. The imaging features of XG inflammatory disease are nonspecific and usually present as heterogeneous solid or cystic masses with destruction of the affected organs. Due to its aggressive nature, it is occasionally misdiagnosed as a malignancy. Hence, it is important for radiologists to familiarize themselves with the relevant imaging findings and consider XG inflammatory disease in the differential diagnosis to reduce the rate of unnecessary surgeries.

### Author Contributions

Conceptualization, Y.D.M.; data curation, Y.D.M., K.H.C., K.S.W., W.K.Y., P.S.H., J.W.K.; formal analysis, Y.D.M., K.H.C., K.S.W., W.K.Y., P.S.H., J.W.K.; supervision, Y.D.M.; writing—original draft, L.S.J., Y.D.M.; and writing—review & editing, all authors.

### Conflicts of Interest

Dal Mo Yang has been a Section Editor of the Journal of the Korean Society of Radiology since 2014; however, he was not involved in the peer reviewer selection, evaluation, or decision process of this article. Otherwise, no other potential conflicts of interest relevant to this article were reported.

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## 복부와 골반의 다양한 장기에서 발생한 황색육아종성 염증 질환의 영상 및 임상 소견: 임상화보

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황색육아종성 염증 질환은 담낭, 담관, 췌장, 비장, 위, 소장, 결장, 충수, 신장, 부신, 요막관, 방광, 후복막, 여성 생식기 등 다양한 장기를 침범하는 드문 양성 질환이다. 황색육아종성 염증 질환의 영상학적 소견은 비특이적이고 대개 비균질한 고형 또는 낭성 종괴로 나타나며 인접한 장기를 침범할 수 있다. 황색육아종성 염증 질환은 공격적인 양상으로 인해 때때로 악성 종양으로 오인될 수 있다. 본 임상화보에서는 복부와 골반의 다양한 장기에서 발생한 황색육아종성 염증 질환의 영상 소견 및 임상양상을 고찰하고자 한다.

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