

Original article

Morphological and Crystal Chemical Characteristics of Gallbladder Biomineralization

Roman Moskalenko¹, Sergiy Danilchenko², Artem Piddubnyi¹, Oleksandr Kravets³,
Inna Chorna⁴, Olena Kolomiets¹, Anatolii Romaniuk¹

¹*Department of Pathology, Sumy State University, Sumy, Ukraine*

²*Institute of Applied Physics of National Academy of Science, Sumy, Ukraine*

³*Department of General Surgery, Sumy State University, Sumy, Ukraine*

⁴*Department of Biophysics, Biochemistry, Pharmacology and Biomolecular Engineering,
Sumy State University, Sumy, Ukraine*

SUMMARY

Pathological biomineralization can be found in some gallbladder (GB) diseases such as chronic calculous cholecystitis (CCCh), gallbladder cancer (GBC) and porcelain gallbladder (PGB).

The aim of the work was to analyze the morphology of pathological biomineralization in GB tissue in CCCh, GBC and PGB.

Five cases of PGB, 10 samples of CCCh and 5 cases of GBC with biomineralization were selected for this study. All samples were examined by histology, histochemistry and scanning electron microscopy with X-ray diffraction.

The X-ray diffraction of mineral deposits of PGB wall and GB concretions revealed their different mineral composition. All PGB and GBC samples had the presence of hydroxyapatite. Calcium-containing GB concretions were composed of calcium carbonate with the presence of trace amounts of other calcium phosphate phases (vaterite, dolomite).

We did not find cancer in all PGB cases. The different crystal phases of biominerals were found in the wall (PGB and GBC) and in the GB cavity (CCCh) during pathology development. The difference between mineral content of biominerals can be caused by various conditions and mechanisms of their formation.

Key words: cholecystitis, gallbladder cancer, porcelain gallbladder, hydroxyapatite, calcium carbonate

Corresponding author:

Roman Moskalenko

E-mail: r.moskalenko@med.sumdu.edu.ua

INTRODUCTION

Gallbladder (GB) diseases are often accompanied by pathological biomineralization (PBM) development. PBM occurs in chronic calculous cholecystitis (CCCh) (cholelithiasis), gallbladder cancer (GC) and porcelain gallbladder (PGB) (1).

Chronic calculous cholecystitis is extremely common pathology. Thereby, the causes, formation mechanisms and chemical composition of GB stones have been well studied (2, 3). Other pathologies, such as gallbladder cancer (GBC) and PGB, were studied to a lesser degree according to their low incidence. Moreover, the PBM in GBC and PGB are actually "terra incognita" (1).

GBC is more common in women with a ratio 5:1. The age of patients ranges from 50 to 70 years (1, 2). GBC prevalence depends on the geographical and ethnic features and varies from 1.59 cases per 100.000 (Maori women, New Zealand) to 22 cases per 100.000 (women, North India) (4). Calcification of the GB wall is associated with GBC in 12-62% of cases (5-7).

PGB is a rare manifestation of chronic GB disease. It has dense calcification deposits in the GB wall and can be found in 0.06-0.8% of cholecystectomies (8). The causes of PGB are exactly unknown. PGB was first described in 1929 as a GB with fragile consistency, uncolored wall and its widespread calcification (6).

The aim of the work was to analyze the morphology of pathological biomineralization in GB tissue in CCCh, GBC and PGB.

MATERIALS AND METHODS

Ethics statement

A written informed consent was obtained from all patients. This research was approved by the Medical Ethics Committee of The Sumy Regional Clinical Hospital and Medical Institute of Sumy State University (Protocol No.3/6, June 07 2016).

Sample collection

All patients were selected during the period 2012-2014 at the surgical department of the Sumy Regional Clinical Hospital. Patients were hospital-

ized routinely with the diagnosis: "Chronic calculous cholecystitis." Laparoscopic cholecystectomy and abdominal drainage were performed in patients under endotracheal anesthesia.

All five cases of PGB were evaluated as clinical findings in female patients at the age of 61.4 ± 1.54 years.

PBM in CCCh was represented as an intraluminal GB concretions. The stones with a high inorganic component were used for this study. The organic calculi were destroyed during the sample processing (burning in a muffle furnace). Therefore, all CCCh samples are presented by GB calcium stones. Ten female patients with age range 54.4 ± 1.77 years were examined.

The study also involved five cases of GBC with PBM (1 male and 4 female patients, mean age 67.6 ± 4.32 years) (Table 1).

Table 1. The list of samples

Case	Age, years	Sex	Diagnosis	Mineral
1	58	f	PGB	Hydroxyapatite
2	59	f	PGB	Hydroxyapatite
3	66	f	PGB	Hydroxyapatite
4	64	f	PGB	Hydroxyapatite
5	66	f	PGB	Hydroxyapatite
6	48	f	CCCh	Calcite, Tricalcium magnesium phosphate
7	50	f	CCCh	Calcite, Tricalcium magnesium phosphate
8	52	f	CCCh	Calcite, vaterite
9	55	f	CCCh	Calcite, vaterite
10	55	f	CCCh	Calcite, dolomite
11	60	f	CCCh	Calcite, dolomite
12	62	f	CCCh	Calcite
13	63	f	CCCh	Calcite, dolomite
14	63	f	CCCh	Calcite, dolomite
15	51	f	CCCh	Calcite
16	57	m	GBC	Hydroxyapatite
17	58	f	GBC	Hydroxyapatite
18	70	f	GBC	Hydroxyapatite
19	75	f	GBC	Hydroxyapatite
20	78	f	GBC	Hydroxyapatite

Histology and histochemistry

The presence of significant mineral deposits in some cases required an intensive decalcification with EDTA.

All tissue samples were fixed in 10% formaldehyde in PBS, dehydrated in ethanol and xylene, embedded in paraffin. 4- μm -thick sections were used for the hematoxylin-eosin, Alizarin red and Von Kossa staining.

X-ray diffraction

Mineral component was isolated by burning at 200 $^{\circ}$ C for 1 hour. X-ray diffraction studies were performed on diffractometer DRON4-07 ("Burevestnik", Russia). Radiation $\text{CuK}\alpha$ (wavelength 0.154 nm) was used under conditions of focusing due to Bragg-Brentano (θ -2 θ) (2 θ - Bragg's angle) (9). The current and the voltage at the X-ray tube were 20 mA and 30 kV, respectively. The samples were scanned at the continuous recording mode (speed 2 $^{\circ}$ /min) in a range of angles 2 θ between 10 and 60 $^{\circ}$. All procedures of the experimental data processing were performed by the usage of the licensed software support package and results processing (DIFWIN-1, TOO "Etalon TCP"). Identification of crystalline phases was carried out by automatic comparison of the obtained results with the database cards Powder Diffraction File 2 without imposing restrictions on the elemental composition of the samples; the software package

Crystallographica Search-Match (Cryosystems, Oxford) was used in the work.

Scanning electron microscopy

Scanning electron microscopy (SEM) with X-ray microanalysis was performed on tissue sections of GB by the REMMA 100U (SEMI, Ukraine). Briefly, 10 μm thick tissue sections were placed on the graphite plates. Paraffin sections were submerged to the three sets of xylene for 5 minute each, followed by three sets of 96 $^{\circ}$ C ethanol for 5 min and finally rinsed with distilled water. Slides were placed into thermostat at 56 $^{\circ}$ C for 10 minutes for drying.

RESULTS

Morphological study

PGBs maintained their shape after their surgical removal. GBs were whitish-gray and dense during palpation. The GB wall was thickened to 1.0 - 1.2 cm, dense, the tissue was crumbled. The mucosa exfoliated easily as thin plates with gray-yellowish color. Cracks were found on a mucosal surface of PGB, which was similar to the surface of old porcelain tableware (Figure 1A).

Macropreparations of PGB had a different spread of PBM and ranged from the total calcification of the organ to calcification of large parts of the GB wall (more than 50% of the GB wall).

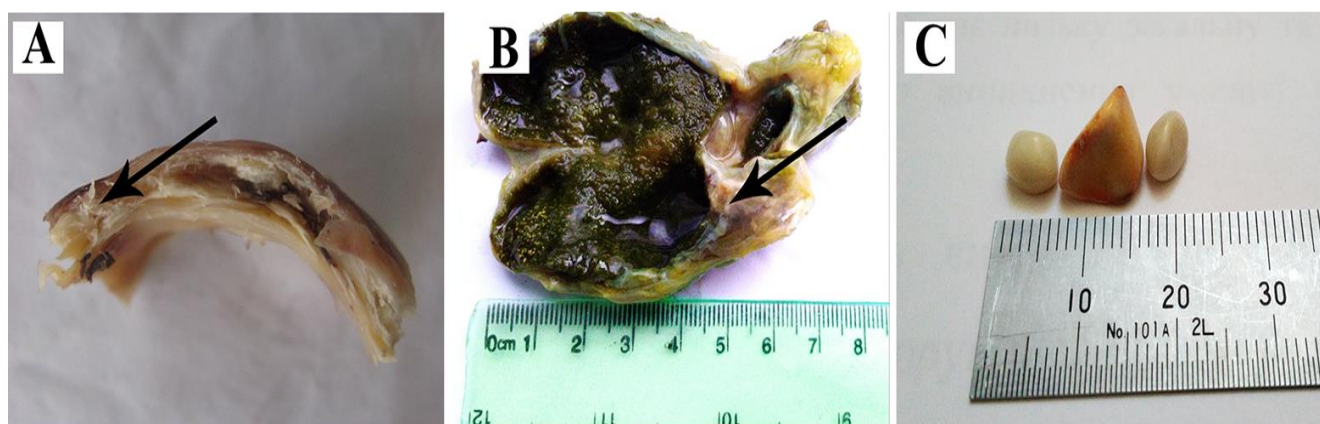


Figure 1. GB with PBM. A. The PGB wall with a massive deposition of biominerals (indicated by arrow), the preparation is fixed in formalin; B – macropreparation of GBC (tumor thickening of the walls is showed by the arrow); C – calculus from the patients with CCCh.

GBC specimens were collapsed, had gray-pink color of serous membranes. GB tissues were soft; the walls were thickened to 0.6 - 0.7 cm (Figure 1B). Visible small gray nodules with a diameter of 0.1 - 0.2 cm, hemorrhages, fibrous tissue overgrowth were found in GBC. One GBC patient had polyps with the diameter of 0.1-0.3 cm at the GB mucosa.

GBs with CCCh were of pink-grayish color and they were collapsed. The slight wall thickening to 0.4 - 0.5 cm was defined (normal 0.2 - 0.3 cm). The GB mucosa had velvety surface with pinpoint hemorrhages. The GB wall had fibrous tissue overgrowth

and hemorrhages. White-yellowish round shaped calculi with a diameter of 0.3-2.5 cm (Figure 1C) were found inside the GB.

Histological examination

Histological examination of the gallbladders shows the typical pathological changes of organ tissue. A moderate mixed cell inflammatory infiltration was revealed in the wall tissues of all PGB cases. Inflammation was accompanied by fibrosis, hemorrhages, muscular hypertrophy, hyalinosis and stagnation of secretion (Figure 2A, C).

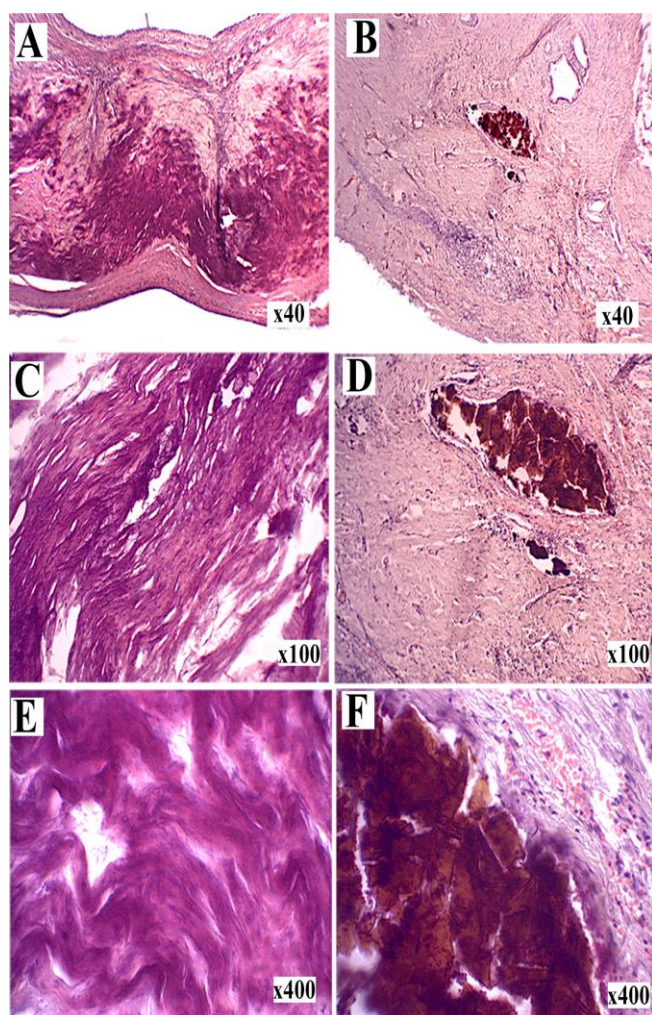


Figure 2. Histological examination of PGB wall and GBC. A, C, E –PGB; B, D, F –GBC. Calcified material deposits mainly in the muscular layer of the PGB wall in the form of roughly dispersed sediments (A) and mineralized fibers in the type of wire (C, E). Biom mineral deposits of GBC are localized within the tumor glands with desquamated epithelium (B, D), which are surrounded by blood vessels and inflammation (F). Hematoxylin-eosin staining. Magnification is indicated in the lower right corner of micrographs.

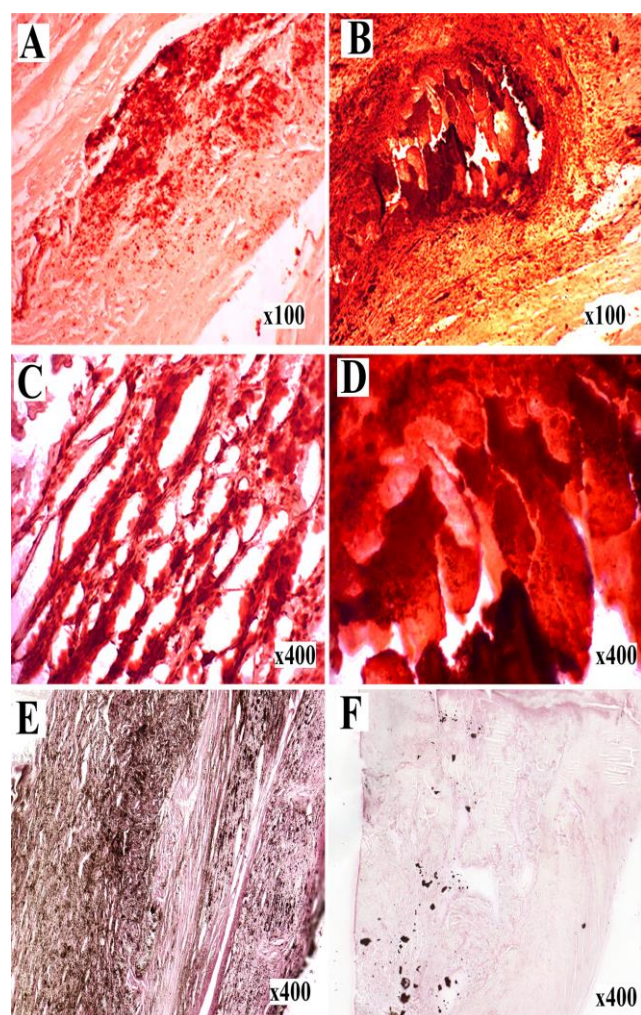


Figure 3. Alizarin red and von Kossa staining of PGB and GBC tissues. A, C, E –PGB. Calcium deposits as sand (A) and fibers (C). B, D, F - GBC. Solitary (single) mineral deposits (B) in the GB wall which are fragmented during histological processing (D). von Kossa - positive deposits are colocalized with Alizarin red - positive structures. A-D – Alizarin red staining, E - F – von Kossa staining. Magnification is indicated in the lower right corner of micrographs.

Calcium deposits were predominantly located in the muscular layer along the muscle fibers and connective tissue. The size of these deposits varied from 0.1 to 5.0 cm. Muscle fibers were similar to "metallic net" in the places of evident calcification (Figure 2E, 3C). Histological examination of CCCh tissue revealed a diffuse inflammatory infiltration with lymphocytes and histiocytes, connective tissue excrescence, small hemorrhages and edema. Interstitial deposits of calcium compounds were not found.

GBC was found as a spread of atypical glandular tissue in the mucosa and muscle layer of GB. Cancerous tissue formed single atypical glands and tenia of tumor cells. All GBC cases were represented as adenocarcinoma. Focal inflammatory infiltration, connective tissue excrescence, and blood vessels formation were observed around the tumor tissue

and calcifications in the GB wall (Figure 2F). Calcifications appeared as small single formations, which were found often in malignant glands or they were associated with the tumor. Desquamation of typical epithelial was found in glands with calcium deposits (Figure 2B, D, F; 3B, D).

Scanning electron microscopy

SEM revealed mineralized elements of GBs as white and gray colored objects with the signs of degradation as fragmentation and cracks (artifact material damage during the samples' processing). Electronic scanograms of PGB wall showed the presence of massive biomineral deposits. Calcium compounds were colocalized with fibrous tissue elements (muscles, connective tissue) (Figure 4A). SEM of mineralized GBC tissues revealed single foci

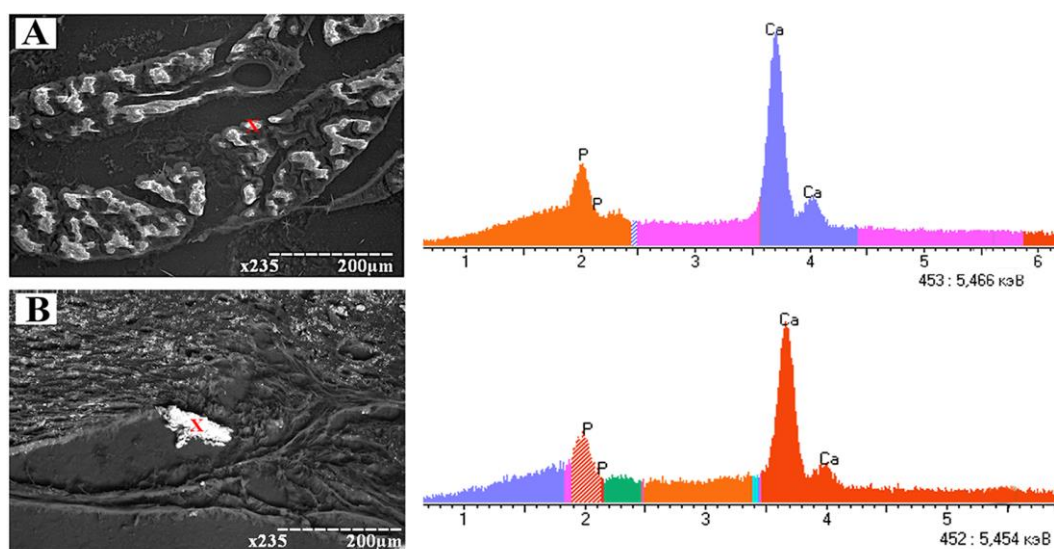


Figure 4. SEM with x-ray microanalysis. A – PGB wall and energy-dispersive X-ray spectroscopy of mineralized tissue; B – GBC tissue and energy-dispersive X-ray spectroscopy of calcified area. Red cross indicates the point of microanalysis. Magnifications are indicated in the lower right corner of micrographs.

of calcification with jagged borders. They were round-shaped and corresponded to the neoplastic glands shape (Figure 4B). X-ray diffraction of PGB and GBC showed a similar chemical composition and ratio of calcium and phosphorus, which corresponds to hydroxyapatite (Figure 4). X-ray diffraction patterns of PGB (Figure 5) corresponded to the structural data of hydroxyapatite ($\text{Ca}_{10}(\text{PO}_4)_6(\text{OH})_2$, JCPDS № 9-0432).

A significant broadening and overlap of diffraction peaks indicated a low level of the material crystallinity, a small size of the region with a regular periodic structure (crystallites or mosaic blocks) and a big amount of defects, which led to the lattice microdeformations. The semiquantitative evaluation of the crystallite size using Scherrer's formula (9) in a perpendicular direction to the crystallographic plane (002) gave the values which were close

to the typical size of bone crystallites. It should be noted that the line, corresponding to the longitudinal size of the apatite crystallites (e.g. 002), was broadened considerably less than the line corresponding to the transverse size (e.g., 310). This suggests a certain crystals' elongation along the hexagonal axis at a relatively small size of the transverse direction. Such crystals' morphology of biological apatite (plate or

rod like) is typical for bone tissue and similar to some synthetic biomaterials.

Structural studies of pathological deposits, which were located in the gallbladder, showed that the major phase of the formed deposit was calcium carbonate (CaCO_3 , JCPDS № 83-577 and/or JCPDS № 88-1810) that is characterized by a high degree of crystallinity (Figure 6).

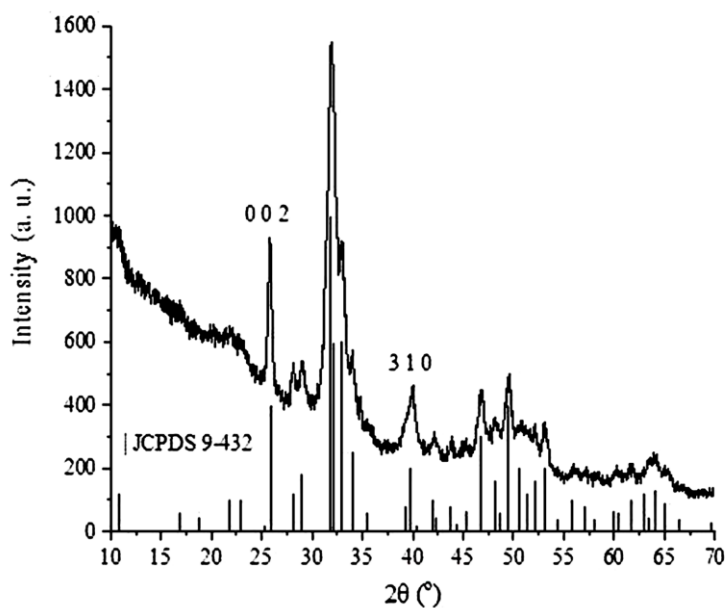


Figure 5. Patterns of X-ray diffraction of the GB wall mineral formations; vertical lines correspond to the angular positions and relative intensities of the lines of the standard JCPDS № 9-0432; line with the indices 002 and 310 were used to estimate the size of the crystallites.

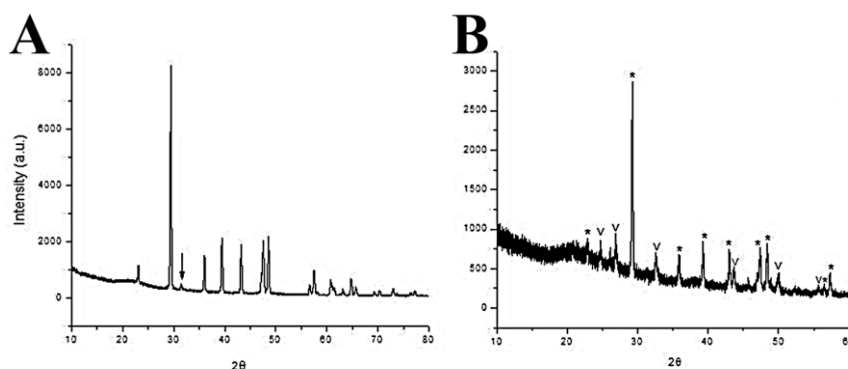


Figure 6. X-ray diffraction patterns of PBM formations in the GB; A – the strongest line of additional trace calcium phosphate phase ($\beta\text{-(Ca,Mg)}_3(\text{PO}_4)_2$ and/or apatite) is marked by the arrow, B – * - lines of calcite, JCPDS № 88-1810, v - vaterite lines, JCPDS № 74-1867.

The trace amounts (a few percent) of another crystal phase or phases, presumably, β -tricalcium magnesium phosphate (β -(Ca,Mg)₃(PO₄)₂) and/or apatite, were detected. In the one sample, a mixture of two different structural (polymorphic) forms of calcium carbonate - vaterite (JCPDS № 74-1867) and calcite (JCPDS № 88-1810), with a predominant concentration of the last one, was found (Figure 6B).

DISCUSSION

There are two types of PGB, according to the calcification level: complete (covers the entire organ, penetrates the muscle layer) and incomplete (multifocal, point deposits) (5, 10). The combination of GBC and PGB with incomplete calcification type, according to various data, ranges between 0% and 5% (11). There was no information about the combination of complete type of PGB and malignant tumors. This can indicate that two types of calcification cause different risk of GBC development. Despite this, a preventive simple cholecystectomy is the variant of PGB treatment (12).

Consequently, we did not detect cancer in all examined PGB samples. The results of histological and histochemical examinations show similarity of pathological changes (presence of chronic inflammation). Obviously, the idea that PGB is a type of chronic cholecystitis has a morphological basis (8). It is also known that PGB is associated with cholelithiasis in 90% of cases (13).

Differential diagnosis between PGB and calcified carcinoma of GB requires more precise criteria, as the correct diagnosis has a decisive influence on further treatment of the patient and on the perspectives for recovery.

Historically, the idea about the link between the PGB and the risk of GBC has changed. First works from the 60-70's in the 20th century described a rather high incidence of malignancy (malignant transformation) of GB - up to 61% (6, 14). Recent studies claim that the incidence of carcinomas in PGB ranges from 0% to 5% (5, 10). Also, it was found that the complete type of PGB is not associated with GBC (5, 8, 11).

The results of our study and literature review

Table 2. Histopathological features of different forms of gallbladder calcification

	<i>CCC</i>	<i>GBC</i>	<i>PGB</i>
Size	Enlarged	Enlarged	Reduced in size
Wall thickness	Slightly thickened	Thickened	Thickened
Morphology	GB wall with normal structure, fibrosis, hyalinosis, inflammation	Fibrosis, atypical glands with invasive grow	The number of cells is reduced, massive deposits of small and big biominerals
Localization of biominerals	In the lumen (stones)	In the wall (in tumor tissue) as single biominerals	Walls are totally calcificated, solid layer, sometimes it spreads to the whole organ
Mineral compound	Calcium carbonate, dolomite	Hydroxyapatite	Hydroxyapatite
Fibrosis	Moderate	Moderate	Severe
Hyalinosis	Moderate	Moderate	Severe

are presented in Table 2.

PGB and GBC have common features of PBM. The result of PBM is the formation of hydroxyapatite. The common element is the development of dystrophy and necrosis in the GB walls. They are accompanied by cell death and by the occurrence of the building material for mineralization (calcium and phosphates). The reason of cell death in the case of PGB is inflammation, and in the case of GBC the damaging factor is infiltrative growth and disinte-

gration of tumor cells, which leads also to secondary inflammation.

The prevalence of hydroxyapatite was found in our previous studies of biomineral formations of aorta and heart valves, prostate, and eye (15-17). We can assume that the similar feature for all these cases is a local tissue damage with the collagen fibers denudation, that is a matrix for bone mineralization. Therefore, we suggest that bioapatite crystals are similar to bone and synthetic biomaterials (15).

Unlike PGB and GBC, mineralization develops in organ cavity in conditions of CCCh. Calcium-containing concretions are more resistant to mechanical destruction in comparison to stones, which are organic or have a significant organic component. Calcium stones can have a different shape, their color ranges from snow-white (calcium palmitate) to dark brown (calcium carbonate - laterite), even black (calcium bilirubinate) (3). Due to the results of our research, calcium carbonate with the signs of small amounts of calcium phosphate phases (vaterite, dolomite) forms mainly in CCCh.

We have also studied the pathological biominerals of non-apatite nature in the pancreas (18). Biominerals with calcium carbonate develops in the pancreatic tissue and ducts as well as in the lumen of GB. Apparently, this is due to the functions of bicarbonate buffer systems.

The difference of mineral content of biominerals can be caused by various conditions and formation mechanisms. It is obvious that different pH and the environment are in GB wall and cavity. It is caused by the influence of various pathological conditions. One of the possible reasons could be an insufficient amount of phosphorus and lack of collagen matrix in the GB cavity.

CONCLUSIONS

Different crystal phases of biominerals were found in the wall (PGB and GBC) and in the GB cavity (CCCh) during pathology development. Intraperietal biominerals were represented by hydroxyapatite, stones from the GB cavity consisted predominantly of calcium carbonate with phosphate additives. These results indicate different conditions, causes and mechanisms of their formation

Acknowledgements

This research has been performed with the financial support of grants of the Ministry of Education and Science of Ukraine No. 0117U003937 "The development of tumor diagnosis method of reproductive system organs using cellular adhesion molecules of cancer-embryonic antigen" and No. 0118U003570 "The efficiency of "liquid biopsy" and tissue biopsy in the diagnosis and treatment of malignant tumors", Erasmus+ Project 2017-1-SE01-KA107-034386 between Sumy State University (Sumy, Ukraine) and Umeå University (Umea, Sweden).

Conflicts of interest

Authors have no conflict of interest to declare.

References

1. Cunningham SC, Alexander HR. Porcelain gallbladder and cancer: ethnicity explains a discrepant literature? *Am J Med* 2007; 120:17-8. <https://doi.org/10.1016/j.amjmed.2006.05.028>
2. Sun H, Tang H, Jiang S, et al. Gender and metabolic differences of gallstone diseases. *World J Gastroenterol* 2009; 15 (15):1886-91. <https://doi.org/10.3748/wjg.15.1886>
3. Qiao T, Ma RH, Luo XB, et al. The Systematic Classification of Gallbladder Stones. *PLoS ONE* 2013; 8(10). <https://doi.org/10.1371/journal.pone.0074887>
4. Hundal R, Shafer EA. Gallbladder cancer: epidemiology and outcome. *Clin Epidemiol* 2014; 6: 99-109. <https://doi.org/10.2147/CLEP.S37357>
5. Stephen AE, Berger DL. Carcinoma in the porcelain gallbladder: a relationship revisited. *Surgery* 2001; 129 (6):699-703. <https://doi.org/10.1067/msy.2001.113888>
6. Ashur H, Siegal B, Oland Y, et al.. Calcified gallbladder (porcelain gallbladder). *Arch Surg* 1978; 113:594-6. <https://doi.org/10.1001/archsurg.1978.01370170056010>
7. Patel S, Roa JC, Tapia O, et al. Hyalinizing cholecystitis and associated carcinomas: clinicopathologic analysis of a distinctive variant of cholecystitis with porcelain-like features and accompanying diagnostically challenging carcinomas. *Am J Surg Pathol* 2011; 35(8):1104-13. <https://doi.org/10.1097/PAS.0b013e31822179cc>
8. Palermo M, Nunez M, Duza GE, et al. Porcelain gallbladder: a clinical case and a review of the literature. *Cir Esp* 2011; 89 (4):213-7. <https://doi.org/10.1016/j.ciresp.2010.09.012>
9. Elliott JC. Structure and chemistry of the apatites and other calcium orthophosphates, Elsevier, Amsterdam, 1994: 404.
10. Towfish S, McFadden DW, Cortina GR, et al. Porcelain gallbladder is not associated with gallbladder carcinoma. *Am Surg* 2001; 67:7-10.
11. Yun EJ, Yoon DY, Choi CS, et al. Calcified carcinoma of the gallbladder with calcified nodal metastasis presenting as a porcelain gallbladder: a case report. *Cancer Res Treat* 2011; 43 (1): 71-4. <https://doi.org/10.4143/crt.2011.43.1.71>
12. Kwon AH, Inui H, Matsui Y, et al. Laparoscopic cholecystectomy in patients with porcelain gallbladder base dont he preoperative ultrasound findings. *Hepatogastroenterol* 2004; 51:950-3.
13. Puttasubbappa PS, Pallavi P. Porcelain gallbladder mimicing carcinoma gallbladder - a case report and review of literature. *Indian J Surg* 2013; 75 (Suppl 1): S208-209. <https://doi.org/10.1007/s12262-012-0545-1>
14. Polk HC. Carcinoma and calcified gallbladder. *Gastroenterol* 1966; 50:582-5. [https://doi.org/10.1016/S0016-5085\(66\)80037-9](https://doi.org/10.1016/S0016-5085(66)80037-9)
15. Danilchenko SN, Kuznetsov VN, Stanislavov AS, et al. The mineral component of human cardiovascular deposits: morphological, structural and crystal-chemical characterization. *Crystal Res Technol* 2013, 48 (3): 153-62. <https://doi.org/10.1002/crat.201200443>
16. Moskalenko R, Romanyuk A, Danilchenko S, et al. Morphogenetic aspects of biomineralization on the background of benign prostatic hyperplasia. *Georgian Medical News* 2013; 214 (1): 54-61.
17. Moskalenko R, Romanyuk A, Danilchenko S, et al. Rare case of pathological biomineralization of eye tissue. *Čes a slov Oftal* 2014; 70 (4):160-3.
18. Kravets OV, Danilenko IA, Smorodska OM, et al. Morphological and crystal chemical characteristic of pancreatic lithiasis. *Wiad Lek* 2018; 71, 1, cz. II : 237-41.

Morfološke i hemijske kristalne karakteristike biomineralizacije žučne kese

Roman Moskalenko¹, Sergiy Danilchenko², Artem Piddubnyi¹, Oleksandr Kravets³,
Inna Chorna⁴, Olena Kolomiets¹, Anatolii Romaniuk¹

¹Državni univerzitet u Sumiju, Departman za patologiju, Sumi, Ukrajina

²Institut za primenjenu fiziku nacionalne akademije za nauku, Sumi, Ukrajina

³Državni univerzitet u Sumiju, Departman za opštu hirurgiju, Sumi, Ukrajina

⁴Državni univerzitet u Sumiju, Departman za biofiziku, biohemiju, farmakologiju i biomolekularni
inženjering, Sumi, Ukrajina

SAŽETAK

Patološka biomineralizacija viđa se kod nekih bolesti žučne kese, poput hroničnog kalkuloznog holecistitisa, karcinoma žučne kese i porcelanske žučne kese.

Cilj rada bila je analiza morfologije patološke biomineralizacije u tkivu žučne kese kod hroničnog kalkuloznog holecistitisa, karcinoma žučne kese i porcelanske žučne kese.

Pet slučajeva porcelanske žučne kese, deset uzoraka hroničnog kalkuloznog holecistitisa i pet slučajeva porcelanske žučne kese sa biomineralizacijom, uključeno je u ovu studiju. Svi uzorci su pregledani histološki, histohemijski i pomoću skenirajuće elektronske mikroskopije sa difrakcijom x-zraka.

Difrakcija x-zraka mineralnih depozita zida porcelanske žučne kese ukazala je na različit sastav minerala. Kod svih uzoraka porcelanske žučne kese i karcinoma žučne kese, uočeno je prisustvo hidroksiapatita. Kalcifikati žučne kese sastojali su se od kalcijum-karbonata, sa prisustvom ostalih faza kalcijum-fosfata u tragovima (veterit, dolomit).

Karcinom nije utvrđen u svim slučajevima porcelanske žučne kese. Različite kristalne faze biominerala uočene su u zidu žučne kese (porcelanska žučna kesa i karcinom žučne kese) i u žučnoj kesi (hronični kalkulozni holecistitis) za vreme patološkog razvoja. Razlika između mineralnog sastava biominerala može biti uzrokovana različitim stanjima i mehanizmima njihovog formiranja.

Ključne reči: holecistitis, karcinom žučne kese, porcelanska žučna kesa, hidroksiapatit, kalcijum-karbonat