# **Predictors for adverse outcomes and recurrence of pyometra: a 10-year retrospective study**

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**Objective:** To identify predictors for adverse outcomes and recurrence in patients with pyometra.

**Methods:** Medical records of patients with a diagnosis of pyometra admitted to Tuen Mun Hospital between 1 January 2014 and 31 December 2023 were retrospectively reviewed. Patients' clinical characteristics, laboratory findings, management options, and clinical outcomes (including treatment- and disease-related complications and recurrence) were collected. Patients with or without composite adverse outcomes were compared, as were patients with or without recurrence. Multivariate logistic regression was used to determine independent predictors for composite adverse outcomes and recurrence.

**Results:** In total, 152 patients (median age, 79 years) were included in the analysis; the incidence of pyometra was 0.003%. Composite adverse outcome was more likely to occur in those with diabetes mellitus (adjusted odds ratio [aOR]=6.76, p=0.001) or congestive heart failure or a history of acute myocardial infarction (aOR=4.40, p=0.028), those with the longest diameter of the intrauterine pus collection (aOR=1.26, p=0.009), and those with extended-spectrum  $\beta$ -lactamase–positive bacteria (aOR=6.07, p=0.013), whereas composite adverse outcome was less likely to occur in those with vaginal bleeding (aOR=0.14, p=0.002). Of the patients, 24.0% had recurrence. The risk of recurrence increased with the presence of enterococci (aOR=3.31, p=0.022) and those with the longest diameter of the intrauterine pus collection (aOR=1.16, p=0.033).

**Conclusion:** Pyometra is rare and often associated with malignancy and severe complications. Patients at risk of developing adverse outcomes include those with diabetes mellitus, congestive heart failure or a history of acute myocardial infarction, large pus collection, and infection with extended-spectrum  $\beta$ -lactamase–producing organisms. Patients at risk of recurrence include those with a large pus collection or the presence of enterococci. At-risk patients should be monitored vigilantly. Early diagnosis and intervention are crucial to improving clinical outcomes.

Keywords: Bacteria; Pyometra; Recurrence

## Introduction

Pyometra is the accumulation of pus in the uterine cavity caused by interference with its natural drainage<sup>1,2</sup>. The obstruction in the cervical canal can stem from a malignant or benign tumour, surgery, radiotherapy, or senile cervicitis<sup>3</sup>. Its incidence ranges from 0.01% to  $0.5\%^{1.4.5}$ ); it mainly occurs in postmenopausal women, with an incidence of  $13.6\%^5$ . Pyometra has a possible association with malignancy; untreated cases may result in spontaneous uterine rupture, bacteraemia, pelvic abscess formation, generalised peritonitis, and septic shock<sup>6-9</sup>.

Studies of pyometra primarily consist of case series and case reports. This study aimed to identify predictors for adverse outcomes and recurrence of pyometra and evaluate the microbial profile and antibiotic resistance of patients.

#### Methods

Medical records of patients with a diagnosis of pyometra admitted to Tuen Mun Hospital between 1

January 2014 and 31 December 2023 were retrospectively reviewed. Pyometra was defined as a uterine pus collection confirmed by both pelvic ultrasound and drainage of purulent discharge from the cervix. Patients with haematometra or hydrometra were excluded.

Patients' clinical characteristics, laboratory findings, management options, and clinical outcomes (including treatment- and disease-related complications and recurrence) were collected. The composite adverse outcome was defined as the presence of any of the diseaserelated complications (uterine rupture, sepsis, septic shock, intensive care unit admission, organ derangement, disseminated intravascular coagulopathy, emergency hysterectomy for pyometra, all-cause mortality at 28 days,

Correspondence to: Dr Sze Tik Eugenia LO Email: eugenialo2013@gmail.com and prolonged hospitalisation of  $\geq 21$  days). Treatmentrelated complications entailed drainage-related perforation and failure to drain. Recurrence was defined as any readmission for pyometra within 1 year. Sepsis was defined as an infection leading to organ dysfunction, as reflected by a sequential organ failure assessment (SOFA) score of  $\geq 2^{10}$ . Septic shock was defined as a requirement for vasopressors to maintain a mean arterial pressure (MAP) of  $\geq 65$  mmHg and a serum lactate level of >2 mmol/L (>18 mg/dL), despite adequate fluid resuscitation<sup>11</sup>. Sepsis-induced coagulopathy was defined as a SOFA score of  $\geq 4$ , based on platelet count and prothrombin time (PT)<sup>12</sup>.

Patients with or without composite adverse outcomes were compared using the Student's t test or Mann-Whitney U test for continuous variables and the Chi-squared test or Fisher's exact test for categorical variables. Similarly, patients with or without recurrence were compared. Variables with a p value of <0.05 in the univariate analysis were included in the multivariate logistic regression to determine independent predictors for composite adverse outcomes and recurrence. Statistical analyses were performed using SPSS (Windows version 26.0; IBM Corp, Armonk [NY], United States). A p value of <0.05 was considered statistically significant.

#### Results

In total, 152 patients (median age, 79 years) were included in the analysis (Table 1). There were 43 959 gynaecological admissions during the study period, giving an incidence of 0.003%. Among the patients, 59.9% had impaired ambulation, 48.0% used incontinence pads regularly, and most had medical comorbidities.

Of 152 cultures, 55.9% grew ≥2 microorganisms (Table 2); 16.4% were extended-spectrum  $\beta$ -lactamase (ESBL)-producing bacteria, of which 96% were Escherichia coli and 4.0% were Klebsiella spp. Their isolate detection rates were 33.8% and 6.7%, respectively. Both ESBL-positive and -negative E coli strains had similar resistance rates to amoxicillin/clavulanic acid (coamoxiclav) [7.1% and 6.4%, respectively, Table 3]. One ESBL-positive E coli isolate was resistant to second- and third-generation cephalosporins. Compared with ESBLnegative E coli isolates, ESBL-positive E coli isolates displayed higher antibiotic resistance against levofloxacin (56.5% vs 46.8%) and trimethoprim-sulfamethoxazole (41.7% vs 27.7%) and lower antibiotic resistance against gentamicin (16.7% vs 36.2%). ESBL-negative Klebsiella spp had a <20% resistance rate to most antibiotics, except for trimethoprim-sulfamethoxazole, whereas one ESBL-

Table	1.	Clinical	characteristics	of	patients	with
pyome	etra	a (n=152).				

Characteristic	Value*
Age, y	79 (69-89)
Ethnicity	(0, 0,)
Chinese	151 (99.3)
Pakistani	1 (0.7)
Menopause	148 (97.4)
Impaired ambulation	91 (59.9)
Incontinence pad use	73 (48.0)
Living in nursing home	63 (41.5)
Comorbidity	05 (41.5)
Hypertension	98 (64.5)
Stroke	61 (40.1)
Diabetes mellitus	56 (36.8)
Dementia	44 (29.0)
	44 (29.0) 29 (19.1)
Congestive heart failure / history of acute myocardial infarction	29 (19.1)
Chronic kidney disease	21 (13.8)
Osteoarthritis	10 (6.6)
Long-term medication	10 (0.0)
Steroid	9 (5.9)
5101010	9 (3.9) 4 (2.6)
Other immunosuppressants	, ,
Previous pyometra	10 (6.6)
Intrauterine contraceptive device	6 (4.0)
Endometrial polyp	1 (0.7)
Submucosal fibroid	18 (11.8)
Neighbouring infections: pyosalpinx or tubo-ovarian abscess	6 (4.0)
Enteric or rectal fistula	3 (2.0)
Malignancy	37 (24.3)
Corpus	12 (7.9)
Cervix	21 (13.8)
Vagina	1(0.7)
Double primary (ovary and corpus)	1(0.7) 1(0.7)
Colorectal	
	2 (1.3) 3 (2.0)
Instrumentation (hysteroscopy, dilatation, or curettage of uterus) within 1 year	5 (2.0)
History of loop electrosurgical excision	3 (2.0)
procedure	5 (2.0)
History of pelvic radiotherapy	8 (5.3)
Any of the above	61 (40.1)
Clinical presentation	01 (10.1)
Asymptomatic	9 (5.9)
Vaginal discharge	58 (38.2)
Abdominal pain	25 (16.5)
Fever	25 (16.5)
Maximum temperature, °C (in 25	38.3 (38-39)
women with fever)	50.5 (50-57)
Vaginal bleeding	97 (63.8)
Duration, d (in 117 symptomatic	10 (3-60)
women)	10 (5-00)
Uterine size, wk	6 (6-8)

Data are expressed as median (interquartile range) or No. (%) of patients

Table 2. Laboratory findings of patients withpyometra (n=152).

Laboratory finding	Value*
Pyometra size on pelvic ultrasound	
Longest diameter, cm	2.8 (2.1-4.6)
Shortest diameter, cm	1.2 (1.0-2.3)
Laboratory parameters	
White cell count, $\times 10^{9}/L$ (n=129)	8.1 (6.5-12.3)
Platelet, $\times 10^{9}/L$ (n=129)	245 (187-323)
Urea, mmol/L (n=124)	5.2 (3.9-7.3)
Creatinine, µmol/L (n=124)	66 (57-83.8)
Alanine transaminase, U/L (n=119)	12 (8-16)
Alkaline phosphatase, U/L (n=119)	82 (70-101)
Bilirubin, μmol/L (n=119)	8 (6-10)
Albumin, g/L (n=119)	34 (31-38)
International normalised ratio (n=99)	1.1 (1.0-1.2)
Prothrombin time, s (n=99)	12.6 (11.8-13.7)
Activated partial thromboplastin time,	25.5 (24.1-27.9)
s (n=99)	
C-reactive protein, mg/L (n=75)	33 (3.7-107)
Positive culture	
Genital tract swab	124 (81.6)
Blood	4 (2.6)
Genital tract microorganisms	
Gram-positive bacteria	
Streptococci	35 (23.0)
Enterococci	19 (12.5)
Peptostreptococci	14 (9.2)
Peptoniphilus spp	14 (9.2)
Actinomyces spp	12 (7.9)
Staphylococci	8 (5.3)
Gram-negative bacteria	
Escherichia coli	71 (46.7)
Bacteroides spp	25 (16.5)
Klebsiella pneumoniae	15 (9.9)
Extended-spectrum $\beta$ -lactamase	25 (16.5)
positive	
Fungus	
<i>Candida</i> spp	18 (11.8)
Others <sup>†</sup>	63 (41.5)
Polymicrobial (≥2 microorganisms)	85 (55.9)

Data are expressed as median (interquartile range) or No. (%) of participants.

Diphtheroids (3.9%), Proteus mirabilis (3.9%), Prevotella spp (3.9%), Lactobacillus spp (3.9%), Clostridium spp (3.3%), Fusobacterium spp (2.6%), Actinobaculum spp (2.6%), Morganella morganii (2.0%), Citrobacter spp (2.0%), Porphyromonas spp (2.0%), methicillin-resistant Staphylococcus aureus (1.3%), Gardnerella vaginalis (1.3%), Haemophilus parainfluenzae (1.3%), Dialister micraerophilus (1.3%), Pseudomonas aeruginosa (0.7%), Bifidobacterium spp (0.7%), Parvimonas micra (0.7%), Trueperella bernardiae (0.7%), Providencia stuartii (0.7%), anaerococci (0.7%), Burkholderia cepacia complex (0.7%), group A streptococci (0.7%), and commensals (0.7%). positive *Klebsiella* isolate was sensitive to gentamicin, levofloxacin, and trimethoprim-sulfamethoxazole. No streptococcal isolates displayed antibiotic resistance against penicillin. For enterococci, 22.2% showed resistance to ampicillin and none to vancomycin. The most frequently used antibiotic regimen was co-amoxiclav and levofloxacin (57.2%), followed by co-amoxiclav and metronidazole (24.3%), levofloxacin and metronidazole (13.8%), and cefuroxime and metronidazole (12.5%). The median duration of antibiotic use was 14 days; 25% of patients required change of antibiotics upon availability of culture results.

With regard to clinical management, 88.8% of patients were treated with endometrial aspiration, followed by antibiotics alone (5.9%), Foley drainage (2.6%), emergency hysterectomy (2.0%), and laparotomy with drainage (0.7%) [Table 4]. For the three patients with emergency hysterectomy, the first one underwent staging laparotomy and hysterectomy for an infected uterine tumour, pyometra, and pyosalpinx; the histopathological outcome was carcinosarcoma. The second patient presented with septic shock and underwent staging laparotomy and hysterectomy for an infected ovarian tumour and pyometra; the histopathological outcome was endometrioid adenocarcinoma involving the endometrium and ovary. The third patient had uterine rupture and septic shock secondary to pyometra and pyosalpinx. In the patient who underwent laparotomy with drainage, she presented with urosepsis and septic shock secondary to pyometra, uterine rupture, and tubo-ovarian abscess. There were seven (4.6%) treatmentrelated complications including drainage-related uterine perforation (n=2) and drainage failure (n=5).

A total of 33 (21.7%) patients had disease-related complications, particularly organ derangement (n=24), sepsis (n=15), and prolonged hospitalisation (n=13) [Table 4]. The rate of all-cause mortality at 28 days was 2.0% (n=3); all three patients had multiple comorbidities before admission. One patient was admitted for drug-related acute kidney injury. Imaging examination incidentally revealed pyometra, and drainage was performed. The patient had cardiac arrest and died despite resuscitation. The remaining two patients died of sepsis resulting from pyometra and other coexisting infections (urinary tract infection, pneumonia, and right-leg infective collection).

Composite adverse outcome was more likely to occur in those with diabetes mellitus (adjusted odds ratio [aOR]=6.76, p=0.001) or congestive heart failure / a history of acute myocardial infarction (aOR=4.40, p=0.028), those

Antibiotic	Streptococci (n=35)		Enterococci (n=19)		ESBL-negative Escherichia coli (n=47)		ESBL-positive Escherichia coli (n=24)				ESBL-positive Klebsiella Pneumoniae (n=1)	
	No. of patients	Drug resist- ance, %	No. of patients	Drug resist- ance, %	No. of patients	Drug resist- ance, %	No. of patients	Drug resist- ance, %	No. of patients	Drug resist- ance, %	No. of patients	Drug resist- ance, %
Ampicillin	-	-	18	22.2	-	-	-	-	-	-	-	-
Penicillin	35	0	1	0	-	-	-	-	-	-	-	-
Amoxicillin/clavulanic acid	-	-	1	0	47	6.4	14	7.1	14	7.1	-	-
Second-generation cephalosporin	-	-	1	100	47	0	1	100	14	7.1	-	-
Third-generation cephalosporin	-	-	-	-	8	0	1	100	4	0	-	-
Fourth-generation cephalosporin	-	-	-	-	2	0	-	-	1	0	-	-
Gentamicin	-	-	1	100	47	36.2	24	16.7	14	7.1	1	0
Levofloxacin	2	0	1	0	47	46.8	23	56.5	14	14.3	1	0
Erythromycin	32	18.8	1	100	-	-	-	-	-	-	-	-
Clindamycin	14	28.6	-	-	-	-	-	-	-	-	-	-
Ticarcillin/clavulanic acid	-	-	-	-	47	0	-	-	14	7.1	-	-
Trimethoprim- sulfamethoxazole	21	4.8	-	-	47	27.7	24	41.7	14	21.4	1	0
Piperacillin/tazobactam	-	-	1	100	4	0	-	-	2	0	-	-
Amikacin	-	-	-	-	4	0	9	0	2	0	-	-
Ertapenem	-	-	-	-	1	0	4	25	-	-	-	-
Imipenem	-	-	-	-	2	0	5	0	1	0	-	-
Meropenem	-	-	-	-	2	0	4	0	1	0	-	-
Vancomycin	-	-	17	0	1	0	-	-	-	-	-	-
Cefoperazone/sulbactam	-	_	-	-	2	0	-	-	1	0	-	

#### Table 3. Antibiotic resistance of extended-spectrum β-lactamase (ESBL)–positive and –negative bacteria.

with the longest diameter of the intrauterine pus collection (aOR=1.26, p=0.009), and those with ESBL-positive bacteria (aOR=6.07, p=0.013), whereas composite adverse outcome was less likely to occur in those with vaginal bleeding (aOR=0.14, p=0.002) [Table 5].

Of the patients, 24.0% had recurrence, and the median time interval from the index admission to recurrence was 1 (interquartile range, 0.6-2.8) month. The risk of recurrence increased with the presence of enterococci (aOR=3.31, p=0.022) and those with the longest diameter of the intrauterine pus collection (aOR=1.16, p=0.033) [Table 6].

#### Discussion

Pyometra is a rare condition that mainly affects older women; 74.1% of cases are idiopathic<sup>1</sup>. It probably results from genital atrophy, compounded by poor immunity and suboptimal hygiene<sup>5</sup>. In our study, malignancy was the most common predisposing factor, present in 24.3% of patients. Malignancy may distort intrauterine anatomy and obstruct outflow tract drainage; cervical cancer was the most frequently identified cancer.

In our study, 21.7% of patients had disease-related complications; 3.9% had septic shock, which is lower than the 14.6% reported in a study in South Korea, although it

Outcomes	Value*
Management	
Endometrial drainage	135 (88.8)
Foley drainage	4 (2.6)
Antibiotics alone	9 (5.9)
Hysterectomy	3 (2.0)
Laparotomy and drainage	1 (0.7)
Antibiotic regimen	
Co-amoxiclav + metronidazole	37 (24.3)
Levofloxacin + metronidazole	21 (13.8)
Co-amoxiclav + levofloxacin	87 (57.2)
Cefuroxime + metronidazole	19 (12.5)
Ceftriaxone + metronidazole	9 (5.9)
Piperacillin/tazobactam	15 (9.9)
Vancomycin	6 (4.0)
Carbapenem	16 (10.5)
Pre-drainage antibiotics (n=140;	124 (88.6)
excluding those with antibiotics alone and hysterectomy)	
Duration of antibiotics, d	14 (14-14)
Disease-related complications	
Uterine rupture	2 (1.3)
Sepsis	15 (9.9)
Septic shock	6 (4.0)
Intensive care unit admission	2 (1.3)
Organ derangement	24 (15.8)
Disseminated intravascular coagulopathy	7 (4.6)
All-cause mortality at 28 days	3 (2.0)
Emergency hysterectomy	3 (2.0)
Prolonged hospitalisation	13 (8.6)
Treatment-related complications	
Drainage related perforation	2 (1.3)
Failure to drain	5 (3.3)
Recurrence (n=146; excluding those with mortality and hysterectomy)	35 (24.0)
Time interval of recurrence from index admission, m	1.0 (0.6-2.8)

 Table 4. Management and clinical outcomes of patients with pyometra (n=152).

Data are expressed as median (interquartile range) or No. (%) of patients.

lacked a clear definition of septic shock<sup>13</sup>. In our study, four patients underwent emergency hysterectomy or laparotomy with drainage secondary to septic shock and/or uterine rupture. Such an association was also reported in the study in South Korea<sup>13</sup>. The morbidity and mortality rates were

high when pyometra was complicated with uterine rupture and subsequent peritonitis<sup>14,15</sup>. The poor prognosis was partly attributed to the non-specific clinical presentations, leading to misdiagnosis and delayed management. Therefore, pyometra with uterine rupture should be considered in patients with haemodynamic instability and early intervention is imperative.

Cervical stenosis predisposes patients to recurrence. The incidence of recurrence has been reported to be 22% to  $31.4\%^{4,16}$ . It was 24.6% in patients treated in a single hospital in South Korea between 2010 and 2021<sup>17</sup>. Similarly, our patients were predominantly treated with endometrial drainage, which yielded a recurrence rate of 24.0%. However, another South Korean study in 2024 reported a recurrence rate of  $6.3\%^{13}$ . The lower recurrence rate can be attributed to the use of Foley catheter drainage<sup>5,13</sup>. Detection of recurrence can be affected by symptom severity, patient and caregiver awareness, and follow-up availability and duration.

Older women with chronic diseases and restricted mobility had an increased risk of pyometra<sup>18</sup>. Diabetes mellitus predisposes patients to immunosuppression, whereas congestive heart failure and a history of acute myocardial infarction reduce mobility. A Taiwanese study found that infection-related admissions in patients with heart failure were associated with higher mortality, decompensation, myocardial infarction, stroke, and worsened long-term prognosis<sup>19</sup>. Patients presenting with vaginal bleeding were less likely to experience adverse outcomes, because postmenopausal bleeding could prompt early medical attention and timely treatment, preventing complications. The longest diameter of the intrauterine pus collection was a predictor for both adverse outcomes and recurrence. Similarly, the incidence of septic shock increases with the longest diameter of pyometra, albeit not significantly in the multivariate analysis<sup>13</sup>. Pyometra size reflects disease severity, chronicity, and drainage interference; larger collections increase the risk of incomplete drainage and recurrence. We recommend monitoring with interval scans to ensure complete drainage and early detection of recurrence in patients with a large pyometra collection, thus improving outcomes and reducing complications.

Common causative organisms in the pus collection include *E coli*, *Bacteroides* spp, streptococci, peptostreptococci, and *Klebsiella pneumoniae*<sup>5,13,17,18,20</sup>. In a 2015 review in Hong Kong, *E coli*, *Bacteroides fragilis*, streptococci, and peptostreptococci were the most

Table 5. Predictors	for adverse	outcomes in	patients	with pyometra.
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Variable	Without adverse	With adverse	Odds ratio	p Value	Adjusted odds	p Value	
	outcomes (n=119)*	outcomes (n=33)*	(95% confidence interval)		ratio (95% confidence interval)	•	
Age, y	78 (68-88)	84 (71-90.5)	-	0.311	-	-	
Menopause	116 (97.5)	32 (97.0)	0.83 (0.08-8.23)	>0.99	-	-	
Impaired ambulation	66 (55.5)	25 (75.8)	2.51 (1.05-6.02)	0.035	2.24 (0.62-8.04)	0.217	
Incontinence pad use	55 (46.2)	18 (54.6)	1.40 (0.64-3.03)	0.397	-	-	
Living in nursing home	47 (39.5)	16 (48.5)	1.44 (0.66-3.13)	0.354	-	-	
Comorbidity							
Hypertension	75 (63.0)	23 (69.7)	1.35 (0.59-3.10)	0.479	-	-	
Stroke	49 (41.2)	12 (36.4)	0.82 (0.37-1.81)	0.618	-	-	
Diabetes mellitus	34 (28.6)	22 (66.7)	5.00 (2.19-11.42)	<0.001	6.76 (2.13-21.44)	0.001	
Dementia	33 (27.7)	11 (33.3)	1.30 (0.57-2.98)	0.530	-	-	
Congestive heart	18 (15.1)	11 (33.3)	2.81 (1.16-6.77)	0.019	4.40 (1.18-16.44)	0.028	
failure / history of acute myocardial infarction							
Chronic kidney disease	15 (12.6)	6 (18.2)	1.54 (0.55-4.35)	0.403	-	-	
Osteoarthritis	8 (6.7)	2 (6.1)	0.90 (0.18-4.43)	>0.99	-	-	
Long-term medications							
Steroid	6 (5.0)	3 (9.1)	1.88 (0.45-7.97)	0.408	-	-	
Other	4 (3.4)	0	-	0.577	-	-	
immunosuppressants							
Previous pyometra	9 (7.6)	1 (3.0)	0.38 (0.05-3.13)	0.691	-	-	
Presence of any	46 (38.7)	15 (45.5)	1.33 (0.61-2.88)	0.481	-	-	
predisposing factor							
Clinical presentation							
Asymptomatic	9 (7.6)	0	-	0.207	-	-	
Vaginal discharge	40 (33.6)	18 (54.5)	2.37 (1.08-5.19)	0.029	3.23 (1.03-10.14)	0.440	
Abdominal pain	19 (16.0)	6 (18.2)	1.17 (0.43-3.22)	0.761	_	-	
Fever	12 (10.1)	13 (39.4)	5.80 (2.31-14.52)	<0.001	1.97 (0.53-7.37)	0.314	
Maximum temperature, °C	38.2 (37.7-38.6)	38.5 (38.1-39)	1.95 (0.68-5.64)	0.155	-	-	
(in 25 women with fever)							
Vaginal bleeding	84 (70.6)	13 (39.4)	0.27 (0.12-0.60)	<0.001	0.14 (0.04-0.49)	0.002	
Uterine size, wk	6 (6-8)	6 (6-12)	-	0.239	-	-	
Pyometra size							
Longest diameter, cm	2.8 (2.0-3.8)	3.2 (2.2-8.9)	-	0.043	1.26 (1.06-1.50)	0.009	
Genital tract microorganisms Gram-positive bacteria							
Streptococci	29 (24.4)	6 (18.2)	0.69 (0.26-1.84)	0.455	-	-	
Enterococci	12 (10.1)	7 (21.2)	2.40 (0.86-6.70)	0.132	-	-	
Peptostreptococci	9 (7.6)	5 (15.2)	2.18 (0.68-7.03)	0.186	-	-	
Peptoniphilus spp	12 (10.1)	2 (6.1)	0.58 (0.12-2.71)	0.735	-	-	
Actinomyces spp	8 (6.7)	4 (12.1)	1.91 (0.54-6.80)	0.293	-	-	
Staphylococci	6 (5.0)	2 (6.1)	1.22 (0.23-6.32)	0.685	-	-	
Gram-negative bacteria							
Escherichia coli	52 (43.7)	19 (57.6)	1.75 (0.80-3.81)	0.157	-	-	
Bacteroides spp	18 (15.1)	7 (21.2)	1.51 (0.57-4.00)	0.404	-	-	
Klebsiella pneumoniae	10 (8.4)	5 (15.2)	1.95 (0.62-6.15)	0.319	-	-	
Extended-spectrum	15 (12.6)	10 (30.3)	3.01 (1.20-7.56)	0.015	6.07 (1.45-25.34)	0.013	
β-lactamase positive							
Fungus Candida ann	9 (6 7)	10 (20.2)	602 (215 1604)	-0.001	4.05 (0.00 17.90)	0.075	
Candida spp	8 (6.7) 62 (52 1)	10 (30.3)	6.03 (2.15-16.94)	<0.001 0.072	4.05 (0.92-17.86)	0.065	
Polymicrobial (≥2 microorganisms)	62 (52.1)	23 (69.7)	2.12 (0.93-4.83)	0.072	-	-	

 $^{\ast}$  Data expressed as median (interquartile range) or No. (%) of patients.

Variable	Without recurrence (n=111)*	With recurrence (n=35)*	Odds ratio (95% confidence interval)	p Value	Adjusted odds ratio (95% confidence interval)	p Value
Age, y	79 (69-88)	84 (71-90)	-	0.430	-	-
Menopause	108 (97.3)	35 (100)	-	>0.99	-	-
Impaired ambulation	62 (55.9)	25 (71.4)	1.98 (0.87-4.50)	0.102	-	-
Incontinence pad use	51 (46.0)	19 (54.3)	1.40 (0.65-3.00)	0.389	-	-
Living in nursing home Comorbidity	44 (39.6)	17 (48.6)	1.44 (0.67-3.09)	0.350	-	-
Hypertension	73 (65.8)	23 (65.7)	1.00 (0.45-2.22)	0.996	-	-
Stroke	40 (36.0)	19 (54.3)	2.11 (0.98-4.55)	0.055	-	-
Diabetes mellitus	39 (35.1)	14 (40.0)	1.23 (0.56-2.69)	0.602	-	-
Dementia	30 (27.0)	13 (37.1)	1.60 (0.71-3.56)	0.252	-	-
Congestive heart failure / history of acute myocardial infarction	23 (20.7)	3 (8.6)	0.36 (0.10-1.28)	0.101	-	-
Chronic kidney disease	15 (13.5)	4 (11.4)	0.83 (0.26-2.67)	>0.99	-	-
Osteoarthritis Long-term medications	5 (4.5)	4 (11.4)	2.74 (0.69-10.81)	0.218	-	-
Steroid	7 (6.3)	2 (5.7)	0.90 (0.18-4.55)	>0.99	-	-
Immunosuppressants	4 (3.6)	0	-	0.573	-	-
Previous pyometra	7 (6.3)	3 (8.6)	1.39 (0.34-5.70)	0.703	-	-
Presence of any predisposing factor	44 (39.6)	13 (37.1)	0.90 (0.41-1.97)	0.792	-	-
Uterine size, wk	6 (6-8)	6 (6-10)	-	0.145	-	-
Pyometra size						
Longest diameter, cm	2.8 (2.1-3.8)	3.4 (2.3-7.1)	-	0.045	1.16 (1.01-1.32)	0.033
Genital tract microorganisms Gram-positive bacteria						
Streptococci	24 (21.6)	10 (28.6)	1.45 (0.61-3.43)	0.396	-	-
Enterococci	10 (9.0)	9 (25.7)	3.50 (1.29-9.49)	0.019	3.31 (1.19-9.20)	0.022
Peptostreptococci	10 (9.0)	4 (11.4)	1.30 (0.38-4.45)	0.743	-	-
Peptoniphilus spp	10 (9.0)	4 (11.4)	1.30 (0.38-4.45)	0.743	-	-
Actinomyces spp	7 (6.3)	5 (14.3)	2.48 (0.73-8.37)	0.160	-	-
Staphylococci	6 (5.4)	2 (5.7)	1.06 (0.20-5.51)	>0.99	-	-
Gram-negative bacteria						
Escherichia coli	48 (43.2)	20 (57.1)	1.75 (0.81-3.77)	0.151	-	-
Bacteroides spp	21 (18.9)	4 (11.4)	0.55 (0.18-1.74)	0.305	-	-
Klebsiella pneumoniae	11 (9.9)	4 (11.4)	1.17 (0.35-3.95)	0.757	-	-
Extended spectrum β-lactamase positive	20 (18.0)	4 (11.4)	0.59 (0.19-1.85)	0.359	-	-
Fungus						
<i>Candida</i> spp.	14 (12.6)	2 (5.7)	0.42 (0.09-1.95)	0.359	-	-
Polymicrobial (≥2 microorganisms)	61 (55.0)	23 (65.7)	1.57 (0.71-3.47)	0.262	-	-
Management options			-	0.483	-	-
Endometrial drainage	99 (89.2)	34 (97.1)				
Foley drainage	3 (2.7)	1 (2.9)				
Antibiotics alone	7 (6.3)	0				
Laparotomy and drainage	1 (0.9)	0				
Composite adverse outcomes	22 (19.8)	5 (14.3)	0.67 (0.24-1.94)	0.462	-	-

Table 6. Predictors for recurrence in patients with pyometra.

\* Data expressed as median (interquartile range) or No. (%) of patients.

prevalent pathogens<sup>5</sup>. Our department does not routinely test for sexually transmitted pathogens because most of our patients had impaired ambulation and lived in nursing facilities<sup>13</sup>. The microbial data of our patients suggest that the pyometras originate from gastrointestinal tract and urinary tract pathogens.

Infection with ESBL-producing Enterobacteriaceae is associated with adverse outcomes and thus higher mortality rate, treatment failure, longer hospital stays, and greater hospital expenses<sup>21,22</sup>. Correct empirical antibiotic use is a key contributor; treatment with carbapenem results in significantly lower mortality (3.7%), compared with quinolones (36.3%), cephalosporins (40%), and  $\beta$ -lactam/  $\beta$ -lactamase inhibitors (50%)<sup>21</sup>. The high efficacy of carbapenems against ESBL strains is the result of minimal hydrolysis by ESBL enzymes and reduced susceptibility to the inoculum effect<sup>21,23</sup>. In Hong Kong, the IMPACT guideline supports carbapenem as the preferred agent for serious ESBL-positive infections<sup>24</sup>, as reflected in the low resistance rate in our study.

In our study, enterococci were associated with higher recurrence rates. As gut commensal and opportunistic pathogens, they exhibit resilience, antibiotic resistance, and biofilm formation, complicating treatment and leading to recurrent infections<sup>25,26</sup>. Risk factors for enterococcal overgrowth include prolonged hospitalisation, urinary catheterisation, and repeated antibiotic use<sup>25,26</sup>, which were common in our patients. Efforts to reduce enterococcal colonisation should include judicious antibiotic use, enhanced surveillance, and preventive measures to minimise transmission<sup>26</sup>.

In our study, polymicrobial cultures were present in 55.9% of our patients. Postmenopausal changes, including reduced lactobacilli and increased anaerobes, result in vaginal flora dysbiosis and predisposition to polymicrobial infections<sup>27</sup>. Therefore, broad-spectrum antibiotics should be prescribed to target common pathogens, particularly anaerobes. Co-amoxiclav, levofloxacin, cefuroxime, and metronidazole were the common antibiotics used in our patients. Co-amoxiclav provides good coverage for both Gram-positive and Gram-negative bacteria. Metronidazole primarily targets anaerobes, which are common in female genital tract infections<sup>28</sup>. High quinolone resistance among E coli (50% in our patients and 40% in Hong Kong<sup>24</sup>) raises concerns about levofloxacin as a viable option. Our study supports the suitability of cefuroxime for ESBL-negative infections, but it is ineffective against ESBL-positive strains, consistent with guidelines in Hong Kong<sup>24</sup>.

In Hong Kong, co-amoxiclav and metronidazole would be a reasonable empirical regimen for pyometra, whereas carbapenem should be reserved for severe ESBLpositive infections. Duration of antibiotic use must be customised to individual needs. A microbiologist's input should be considered in patients with recurring pyometra or non-responders to antibiotics.

Our study has several limitations. Only inpatients were included; milder cases of pyometra managed in an outpatient setting were excluded. This selection bias potentially overestimates the severity and frequency of adverse outcomes. Consequently, generalisation of our findings to all pyometra cases is limited. Additionally, subgroup analysis for each adverse outcome was not feasible owing to the small sample size. The retrospective nature of the study is affected by missing data and loss to follow-up. Prospective randomised controlled trials are needed to determine the most optimal management strategies for improvement of clinical outcomes.

#### Conclusion

Pyometra is rare and often associated with malignancy and severe complications. Patients at risk of developing adverse outcomes include those with diabetes mellitus, congestive heart failure or a history of acute myocardial infarction, large pus collection, and infection with ESBL-producing organisms. Patients at risk of recurrence include those with large pus collection or the presence of enterococci. At-risk patients should be monitored vigilantly. Early diagnosis and intervention are crucial to improving clinical outcomes.

#### Contributors

All authors designed the study, acquired the data, analysed the data, drafted the manuscript, and critically revised the manuscript for important intellectual content. The authors had full access to the data, contributed to the study, approved the final version for publication, and take responsibility for its accuracy and integrity.

## **Conflicts of interest**

As an editor of the journal, PLS was not involved in the peer review process. Other authors have no conflict of interest to disclose.

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### Data availability

All data generated or analysed during the present study are available from the corresponding author upon reasonable request.

#### Ethics approval

The present study was approved by the Central Institutional Review Board of Hospital Authority (reference: IRB-2024-276). All patients were treated in accordance with the tenets of the Declaration of Helsinki.

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