



OFFICIAL PUBLICATION OF
THE MINISTRY OF HEALTH,
BRUNEI DARUSSALAM

Brunei International Medical Journal

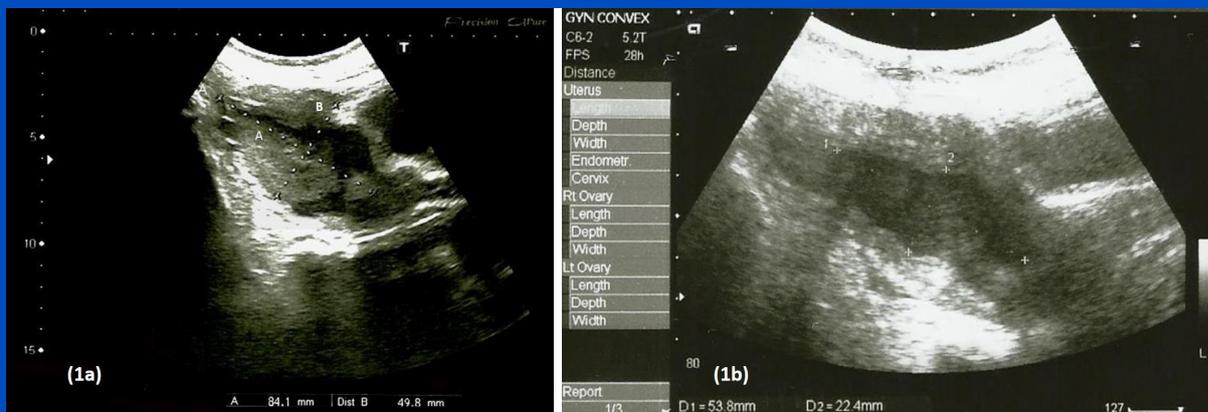
Volume 15

16 April 2019 (10 Sya'ban 1440H)

RESISTANT PYOMETRA IN POST-MENOPAUSAL WOMAN WITH POORLY CONTROLLED DIABETES MELLITUS: A CASE REPORT.

WM Nazlee WZ, Rosnani Z, Imran A

Family Medicine Department, School of Medical Science, Health Campus, Universiti Sains Malaysia, 16150 Kubang Kerian, Kelantan, Malaysia



ABSTRACT

Pyometra is a rare disease defined by accumulation of pus inside uterine cavity. Poorly controlled diabetes mellitus can predispose post-menopausal women to develop pyometra as diabetes is known to suppress patients' immune system and impair the natural drainage of the uterus. We reported here the case of a 66-year-old post-menopausal woman with poorly controlled diabetes who came with a complaint of vaginal whitish brown discharge for one week. The diagnosis of pyometra was made on ultrasound scanning, which is challenging especially in primary care where ultrasound scan may not be widely available. Patient was managed with hysteroscopy endometrial curettage and antibiotics by the admitting gynecology team. Result of biopsy was reported as chronic inflammation but patient refused for further surgical intervention. She was managed conservatively and her diabetes was better controlled on insulin treatment with improvement of her vaginal discharge. This case highlights the important of thorough assessment in post-menopausal women presented with vaginal discharge especially if they have condition which suppressed their immune system. Early diagnosis of pyometra may prevent serious complication such as spontaneous uterine rupture and severe sepsis.

Keywords: Diabetes Mellitus; Diabetes Complications; Post-menopausal; Pyometra; Vaginal discharge.

Brunei Int Med J. 2019;15:53-57

Brunei International Medical Journal (BIMJ)

Official Publication of the Ministry of Health, Brunei Darussalam

EDITORIAL BOARD

Editor-in-Chief	William Chee Fui CHONG
Sub-Editors	Vui Heng CHONG Ketan PANDE
Editorial Board Members	Nazar LUQMAN Muhd Syafiq ABDULLAH Alice Moi Ling YONG Ahmad Yazid ABDUL WAHAB Jackson Chee Seng TAN Dipo OLABUMUYI Pemasiri Upali TELISINGHE Roselina YAAKUB Pengiran Khairol Asmee PENGIRAN SABTU Dayangku Siti Nur Ashikin PENGIRAN TENGAH

INTERNATIONAL EDITORIAL BOARD MEMBERS

Lawrence HO Khek Yu (Singapore)	Surinderpal S BIRRING (United Kingdom)
Emily Felicia Jan Ee SHEN (Singapore)	Leslie GOH (United Kingdom)
John YAP (United Kingdom)	Chuen Neng LEE (Singapore)
Christopher HAYWARD (Australia)	Jimmy SO (Singapore)
Jose F LAPENA (Philippines)	Simon Peter FROSTICK (United Kingdom)

Advisor

Wilfred PEH (Singapore)

Past Editors

Nagamuttu RAVINDRANATHAN
Kenneth Yuh Yen KOK

Proof reader

John WOLSTENHOLME (CfBT Brunei Darussalam)

Aim and Scope of Brunei International Medical Journal

The Brunei International Medical Journal (BIMJ) is a six monthly peer reviewed official publication of the Ministry of Health under the auspices of the Clinical Research Unit, Ministry of Health, Brunei Darussalam.

The BIMJ publishes articles ranging from original research papers, review articles, medical practice papers, special reports, audits, case reports, images of interest, education and technical/innovation papers, editorials, commentaries and letters to the Editor. Topics of interest include all subjects that relate to clinical practice and research in all branches of medicine, basic and clinical including topics related to allied health care fields. The BIMJ welcomes manuscripts from contributors, but usually solicits reviews articles and special reports. Proposals for review papers can be sent to the Managing Editor directly. Please refer to the contact information of the Editorial Office.

Instruction to authors

Manuscript submissions

All manuscripts should be sent to the Managing Editor, BIMJ, Ministry of Health, Brunei Darussalam; e-mail: editor-in-chief@bimjonline.com. Subsequent correspondence between the BIMJ and authors will, as far as possible via should be conducted via email quoting the reference number.

Conditions

Submission of an article for consideration for publication implies the transfer of the copyright from the authors to the BIMJ upon acceptance. The final decision of acceptance rests with the Editor-in-Chief. All accepted papers become the permanent property of the BIMJ and may not be published elsewhere without written permission from the BIMJ.

Ethics

Ethical considerations will be taken into account in the assessment of papers that have experimental investigations of human or animal subjects. Authors should state clearly in the Materials and Methods section of the manuscript that institutional review board has approved the project. Those investigators without such review boards should ensure that the principles outlined in the Declaration of Helsinki have been followed.

Manuscript categories

Original articles

These include controlled trials, interventional studies, studies of screening and diagnostic tests, outcome studies, cost-effectiveness analyses, and large-scale epidemiological studies. Manuscript should include the following; introduction, materials and methods, results and conclusion. The objective should be stated clearly in the introduction. The text should not exceed 2500 words and references not more than 30.

Review articles

These are, in general, invited papers, but unsolicited reviews, if of good quality, may be considered. Reviews are systematic critical assessments of

literature and data sources pertaining to clinical topics, emphasising factors such as cause, diagnosis, prognosis, therapy, or prevention. Reviews should be made relevant to our local setting and preferably supported by local data. The text should not exceed 3000 words and references not more than 40.

Special Reports

This section usually consist of invited reports that have significant impact on healthcare practice and usually cover disease outbreaks, management guidelines or policy statement paper.

Audits

Audits of relevant topics generally follow the same format as original article and the text should not exceed 1,500 words and references not more than 20.

Case reports

Case reports should highlight interesting rare cases or provide good learning points. The text should not exceed 1000 words; the number of tables, figures, or both should not be more than two, and references should not be more than 15.

Education section

This section includes papers (i.e. how to interpret ECG or chest radiography) with particular aim of broadening knowledge or serve as revision materials. Papers will usually be invited but well written paper on relevant topics may be accepted. The text should not exceed 1500 words and should include not more than 15 figures illustration and references should not be more than 15.

Images of interest

These are papers presenting unique clinical encounters that are illustrated by photographs, radiographs, or other figures. Image of interest should include a brief description of the case and discussion with educational aspects. Alternatively, a mini quiz can be presented and answers will be posted in a different section of the publication. A maximum of

three relevant references should be included. Only images of high quality (at least 300dpi) will be acceptable.

Technical innovations

This section include papers looking at novel or new techniques that have been developed or introduced to the local setting. The text should not exceed 1000 words and should include not more than 10 figures illustration and references should not be more than 10.

Letters to the Editor

Letters discussing a recent article published in the BIMJ are welcome and should be sent to the Editorial Office by e-mail. The text should not exceed 250 words; have no more than one figure or table, and five references.

Criteria for manuscripts

Manuscripts submitted to the BIMJ should meet the following criteria: the content is original; the writing is clear; the study methods are appropriate; the data are valid; the conclusions are reasonable and supported by the data; the information is important; and the topic has general medical interest. Manuscripts will be accepted only if both their contents and style meet the standards required by the BIMJ.

Authorship information

Designate one corresponding author and provide a complete address, telephone and fax numbers, and e-mail address. The number of authors of each paper should not be more than twelve; a greater number requires justification. Authors may add a publishable footnote explaining order of authorship.

Group authorship

If authorship is attributed to a group (either solely or in addition to one or more individual authors), all members of the group must meet the full criteria and requirements for authorship described in the following paragraphs. One or more authors may take responsibility 'for' a group, in which case the other group members are not authors, but may be listed in an acknowledgement.

Authorship requirement

When the BIMJ accepts a paper for publication, authors will be asked to sign statements on (1) financial disclosure, (2) conflict of interest and (3) copyright transfer. The correspondence author may sign on behalf of co-authors.

Authorship criteria and responsibility

All authors must meet the following criteria: to have participated sufficiently in the work to take public responsibility for the content; to have made substantial contributions to the conception and de-

sign, and the analysis and interpretation of the data (where applicable); to have made substantial contributions to the writing or revision of the manuscript; and to have reviewed the final version of the submitted manuscript and approved it for publication. Authors will be asked to certify that their contribution represents valid work and that neither the manuscript nor one with substantially similar content under their authorship has been published or is being considered for publication elsewhere, except as described in an attachment. If requested, authors shall provide the data on which the manuscript is based for examination by the editors or their assignees.

Financial disclosure or conflict of interest

Any affiliation with or involvement in any organisation or entity with a direct financial interest in the subject matter or materials discussed in the manuscript should be disclosed in an attachment. Any financial or material support should be identified in the manuscript.

Copyright transfer

In consideration of the action of the BIMJ in reviewing and editing a submission, the author/s will transfer, assign, or otherwise convey all copyright ownership to the Clinical Research Unit, RIPAS Hospital, Ministry of Health in the event that such work is published by the BIMJ.

Acknowledgements

Only persons who have made substantial contributions but who do not fulfill the authorship criteria should be acknowledged.

Accepted manuscripts

Authors will be informed of acceptances and accepted manuscripts will be sent for copyediting. During copyediting, there may be some changes made to accommodate the style of journal format. Attempts will be made to ensure that the overall meaning of the texts are not altered. Authors will be informed by email of the estimated time of publication. Authors may be requested to provide raw data, especially those presented in graph such as bar charts or figures so that presentations can be constructed following the format and style of the journal. Proofs will be sent to authors to check for any mistakes made during copyediting. Authors are usually given 72 hours to return the proof. No response will be taken as no further corrections required. Corrections should be kept to a minimum. Otherwise, it may cause delay in publication.

Offprint

Contributors will not be given any offprint of their published articles. Contributors can obtain an electronic reprint from the journal website.

DISCLAIMER

All articles published, including editorials and letters, represent the opinion of the contributors and do not reflect the official view or policy of the Clinical Research Unit, the Ministry of Health or the institutions with which the contributors are affiliated to unless this is clearly stated. The appearance of advertisement does not necessarily constitute endorsement by the Clinical Research Unit or Ministry of Health, Brunei Darussalam. Furthermore, the publisher cannot accept responsibility for the correctness or accuracy of the advertisers' text and/or claim or any opinion expressed.

RESISTANT PYOMETRA IN POST-MENOPAUSAL WOMAN WITH POORLY CONTROLLED DIABETES MELLITUS: A CASE REPORT.

WM Nazlee WZ, Rosnani Z, Imran A

Family Medicine Department, School of Medical Science, Health Campus, Universiti Sains Malaysia, 16150 Kubang Kerian, Kelantan, Malaysia

ABSTRACT

Pyometra is a rare disease defined by accumulation of pus inside uterine cavity. Poorly controlled diabetes mellitus can predispose post-menopausal women to develop pyometra as diabetes is known to suppress patients' immune system and impair the natural drainage of the uterus. We reported here the case of a 66-year-old post-menopausal woman with poorly controlled diabetes who came with a complaint of vaginal whitish brown discharge for one week. The diagnosis of pyometra was made on ultrasound scanning, which is challenging especially in primary care where ultrasound scan may not be widely available. Patient was managed with hysteroscopy endometrial curettage and antibiotics by the admitting gynecology team. Result of biopsy was reported as chronic inflammation but patient refused for further surgical intervention. She was managed conservatively and her diabetes was better controlled on insulin treatment with improvement of her vaginal discharge. This case highlights the important of thorough assessment in post-menopausal women presented with vaginal discharge especially if they have condition which suppressed their immune system. Early diagnosis of pyometra may prevent serious complication such as spontaneous uterine rupture and severe sepsis.

Keywords: Diabetes Mellitus; Diabetes Complications; Post-menopausal; Pyometra; Vaginal discharge.

INTRODUCTION

Pyometra is an accumulation of pus in uterine cavity. It is a rare condition in post-menopausal women who present with vaginal discharge. The diagnosis is quite challenging especially in primary care settings with limited resources. Ultrasound pelvic plays an important role in diagnosing pyometra. The most common cause of pyometra is malignant disease of genital tract.¹⁻³ However, poorly controlled diabetes mellitus (DM) could predispose patient to pyometra due to impairment of patient's immune system and interference with the natural drainage of

uterus.⁴ In a menopause patient, endometrium loses its resistance and the drainage of the cervical canal is impaired which can precipitate the formation of pyometra.⁵ Early diagnosis and management are important as it can prevent patient from developing severe sepsis and uterine perforation. We reported a case of 66-year-old post-menopausal women who had poorly controlled DM presenting with a week history of vaginal discharge and was diagnosed to have pyometra on ultrasound scan. This case serves as a reminder especially to the primary care doctors to include rare diseases as possible differentials and to broaden the horizon of both the possible diagnosis and the scope of management in patient unwilling to conform to conventional management.

Correspondence: Rosnani Zakaria MAFP (Malaysia), FRACGP (Australia), Family Medicine Department, School of Medical Science, Health Campus, Universiti Sains Malaysia, 16150 Kubang Kerian, Kelantan, Malaysia.
Mobile no: +60199866763; Fax No: 09-7653370
email: rosnani@usm.my

CASE REPORT

A 66-year-old lady with underlying DM for 10 years and menopause for 16 years was referred to our family medicine clinic by a private general practitioner (GP) for management of ovarian cyst. She had an episode of non-itchy brownish vaginal discharge for one week. The discharge was non-smelly and of a small amount, staining her panty liner. She had no fever, constitutional symptoms, abdominal pain or history of post-menopausal bleeding. The GP did a pelvic ultrasound which showed a mass containing fluid which he thought was an ovarian cyst. She had poorly controlled DM type 2 with HbA1c of 15.4% (good control: <7.0%) and her fasting blood sugar was 20 mmol/L (normal <7 mmol/L). Clinically she looked well, afebrile and her body mass index (BMI) was 18.9m/kg (weight 41kg, high 1.47m). Her abdomen was soft with no mass palpable. On bimanual examination, her uterus was enlarged and bulky (about 8 weeks in size). Per speculum examination showed normal cervix with a closed os and presence of copious yellowish-greenish discharge pooled at posterior fornix. Pelvic ultrasound performed at our clinic showed an enlarged uterus (9.1cmx4.7cm) with a collection inside uterine cavity (3.5cmx7.2cm). The myometrium thickness was 1.5cm (Figure 1). During this time, high vaginal swab was taken and sent for culture and sensitivity which came back as mixed

growth. Pipelle sampling was also done and it was consistent with abscess tissue and no malignant cell seen.

She was admitted to the gynecology ward and was started on intravenous (i.v) cefuroxime 750mg three times per day (tds) and intravenous metronidazole 500mg tds. Basal bolus subcutaneous (s/c) insulin was started in the ward to control her diabetes. On day four of admission, she underwent hysteroscopy with direct punch biopsy and endometrial curettage under general anesthesia. Hysteroscopic findings was that of hyperemic area at endocervical and endometrium with thick white discharge and slough at left ostium and lower part of uterus. Swab and discharge culture grew *Klebsiella pneumonia* that was sensitive to cefuroxime and cotrimoxazole. Punched biopsies taken at three different areas of the thickened endometrium showed multiple fragments of micro-collagen tissue infiltrated with lymphocyte that was consistent with chronic inflammation and infection on histopathological examination. Two other biopsy taken by endocervical and endometrial curettage shows acute and chronic inflammation. The discharge was also sent for *Mycobacterium tuberculosis bacilli (MTB)* culture and acid-fast bacilli (AFB) smear which was negative.

She was treated with 7 days of i.v

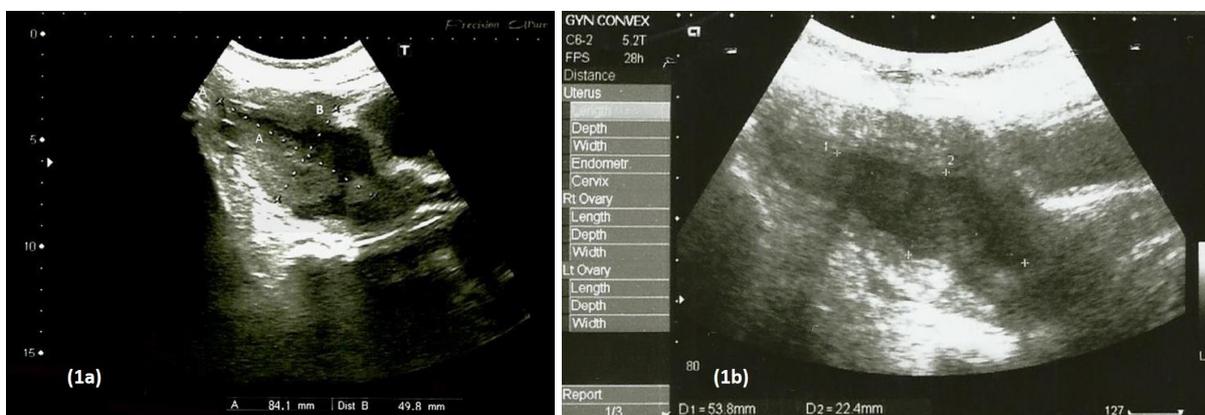


Figure 1: (a) Pelvic ultrasound showed an enlarged uterus measuring 9.1cmx4.7cm with a collection inside uterine cavity (3.5cmx7.2cm). The myometrium thickness was measured at 1.5cm. (b) Pelvic ultrasound performed 1 month post discharged showed residual pyometra after the hysteroscopy endometrial curettage.

cefuroxime and metronidazole and was discharged well after 10 days of hospitalisation. She was discharged with oral antibiotic to be completed for another 4 days and s/c insulin for control of her DM. At follow-up in the gynaecology clinic, she was noted to be well but still had small amount of per vaginal discharge, her blood glucose level varies between 6 mmol/L to 8 mmol/L in the morning indicating good controlled of her DM. Repeat ultrasound scan showed presence of small collection (3.0x2.0 cm) still persistent within uterine cavity (Figure 1b). She was counselled for hysterectomy for unresolved pyometra with her family member, however she declined surgery and was discharged back to our outpatient clinic. At one year follow-up, she was relatively well and still carry on with her usual routine even though she still complained of occasional vaginal discharge. Her diabetes is also better controlled on s/c insulin. Her fasting capillary blood sugar range from 4-7 mmol and her random capillary blood sugar between 7-11mmol. She is however more motivated and adhered to her diabetic treatment better than before.

DISCUSSION

Pyometra is a rare and potentially life-threatening disease. It is more common in women who had malignant uterine lesion (incidence 1.5-4.0%) compare to 0.2% in post-menopausal woman.^{1,3} Even though it makes up only 0.038% of all gynecologic admissions, it is an important cause of uterine rupture.⁶⁻⁸ Classically, pyometra is defines as an accumulation of pus inside uterine cavity and patient may present with post-menopausal bleed, vaginal discharge, intraabdominal pain, intraabdominal mass or hypoalbuminemia.⁷⁻⁹ Yildizhan et al. in 2006 summarised 22 cases of spontaneous ruptured pyometra and found that the most common presenting symptoms were abdominal pain (95.5%), vomiting (41.0%), nausea (9.1%), and fever (9.1%).⁷ He also reported

that only 35% of cases of spontaneous rupture of pyometra were associated with malignancy (Cervical, endometrial and colorectal malignancy). However up to 50% women with pyometra may be asymptomatic.

Ultrasound is usually the initial and reliable imaging study to detect pyometra especially in primary care setting. It shows collection of fluid inside the uterine cavity (enlarged uterus with hypoechoic lesion inside uterine cavity). However, sometimes it is difficult to distinguish it from intrauterine malignancy, hence the uterine biopsy or pipelle sampling may be needed. If perforation of pyometra is suspected, Computed Tomography scan of abdomen is a better diagnostic imaging tool.^{3,6,7}

Vaginal swab culture is important to be done in patient with pyometra even though cultures were positive in only 50% of all cases.¹ Reported organisms cultured from pyometra were *Bacteroides fragilis*, *E. coli* and *Enterococcus* species.¹⁰ *Klebsiella Pneumonia* is gram-negative bacilli bacteria that are commensals in intestines, nasopharynx and skin and it rarely causes infection in genital tract. It usually causes pneumonia, urinary tract infection and wound infection. Poor control of DM predisposes patient to be more susceptible to infection. It is due to low immune response and increased adherence of microorganism towards diabetics mucosal or epithelial cells.^{4,5} Some microorganisms become virulent and active in high glucose environment especially candida and *E. coli* species.⁵

It is important to rule out *Tuberculosis bacilli* (TB) as one of the causative organisms in patient with pyometra especially in immunocompromised patient in TB endemic countries. History of TB contact or previous TB infection should be asked, and sample of endometrial tissue should be routinely sent for AFB stain and MTB PCR to diagnose it.¹¹ Patients diagnosed with TB pyometra response well to

prolong treatment with anti-TB medications usually for 6 months.¹¹

Since most cases (75%) of pyometra have been reported to be associated with some form of malignancy, tissue biopsies should be routinely sent for histopathological examination.^{1,3,7} In our case, multiple punch biopsies of three different sites of the thickened endometrium, one at the endocervix as well as tissues from the endometrial curettage were all sent for histopathological examination of which all were reported as chronic inflammatory changes with no evidence of malignancy.

In uncomplicated cases, pyometra can be managed by dilatation and drainage of uterine cavity. Wide spectrum antibiotics covering for both anaerobic and aerobic bacteria are the antibiotics of choice. Hysterectomy is the treatment of choice especially in cases of spontaneous perforation of pyometra and in refractory cases.^{9,12} In patients who presented with spontaneous perforation of pyometra, exploratory laparotomy needs to be done urgently.^{3,6,7} The mainstay management of our patient was to maintain better diabetic control which was achieved with s/c insulin injection.

CONCLUSION

Pyometra is a rare condition due to collection of pus in the uterus and is commonly associated with uterine malignancy. Tissue diagnosis to exclude uterine malignancy needs to be done and when exclude, causes such as poorly controlled diabetes as with this case can be considered. For unresolved pyometra, hysterectomy should be considered but if patient refused, long-term follow up with good diabetic controlled should be carried out to monitor for complications such as sepsis.

ACKNOWLEDGEMENT

We thanked Associate Prof Dr Adibah Ibrahim and

Dr Rahimah Abdul Rahim our Obstetric and Gynaecologist, for their expertise in co-management of the patient and to Hospital Universiti Sains Malaysia where the patients' main care is provided.

CONFLICT OF INTEREST

No potential conflict of interest relevant to this article was reported.

REFERENCES

- 1: Lui MW, Cheung VY, Pun TC. Clinical Significance of Pyometra. *J Reprod Med.* 2015;60(7-8):329-32.
- 2: Kerimoglu OS, Pekin A, Yilmaz SA, Bakbak BB, Celik C. Pyometra in elderly post-menopausal women: a sign of malignancy. *Eur J Gynaecol Oncol.* 2015;36(1):59-61.
- 3: Geranpayeh L, Fadaei-Araghi M, Shakiba B. Spontaneous Uterine Perforation due to Pyometra Presenting as Acute Abdomen. *Infect Dis Obstet Gynecol.* 2006;2006:1-2. [Accessed on 2018 Jul 18]. Pdf available at <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1581470/pdf/IDOG2006-60276.pdf>
- 4: Knapp S. Diabetes and Infection: Is There a Link?-A Mini-Review. *Gerontology.* 2013;59:99-104. [Accessed on 2018 Jul 18]. Pdf available at <https://www.karger.com/Article/Pdf/345107>
- 5: Geerlings SE, Hoepelman AI. Immune dysfunction in patients with diabetes mellitus (DM). *FEMS Immunol Med Microbiol.* 1999;26(3-4):259-65.
- 6: Bhikaji Shitut P, Wahane A, Dhakne M, Karale V, Prof A, Resident S. An interesting case of a pyometra in a postmenopausal woman. *Sch J Med Case Rep.* 2016;4(1):44-6. [Accessed on 2019 Jan 29]. Pdf available at <http://saspjournals.com/wp-content/uploads/2016/02/SJMCR-4144-46.pdf>
- 7: Yildizhan B, Uyar E, Sişmanoğlu A, Güllüoğlu G, Kavak ZN. Spontaneous perforation of pyometra. *Infect Dis Obstet Gynecol.* 2006;2006:1-3. 26786. [Accessed on 2018 Jul 18]. Pdf available at <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC1581463/pdf/IDOG2006-26786.pdf>
- 8: Ou YC, Lan K-C, Lin H, Tsai CC, ChangChien CC. Clinical characteristics of perforated pyometra and impending perforation: Specific issues in gynecological emergency. *J Obstet Gynaecol Res.* 2010;36(3):661-6.
- 9: Gupta B, Agarwal R, Radhakrishnan G, Delhi N. Spontaneous perforation by pyometra — an

- acute emergency. *SAJOG.* 2011;17(1):4–5. [Accessed on 2018 Jul 18]. Pdf available at https://www.researchgate.net/profile/Bindiya_Gupta/publication/279507186_Spontaneous_perforation_by_pyometra_-_an_acute_emergency/links/5697a63d08ae34f3cf1f07ea/Spontaneous-perforation-by-pyometra-an-acute-emergency.pdf?origin=publication_detail
- 10] Ikeda M, Takahashi T, Kurachi H. Spontaneous Perforation of Pyometra: A Report of Seven Cases and Review of the Literature. *Gynecol Obstet Invest.* 2013;75(4):243–9.
- 11] Lee DH, Cho DH, Lee JH, Kim YR. A case of postmenopausal pyometra caused by endometrial tuberculosis. *Korean J Obstet Gynecol.* 2012;55(6):429-432. [Accessed on 2018 Jul 18]. Pdf available from: <https://synapse.koreamed.org/Synapse/Data/PDFData/2021KJOG/kjog-55-429.pdf>
- 12] Gami N, Mundhra R, Guleria K, Arora VK, Garg S. Recurrent pyometra and xanthogranulomatous salpingitis: A rare pathologic association in a postmenopausal lady. *J Midlife Health.* 2014;5(3):156–8. [Accessed on 2019 Mar 18]. Full text available at <https://www.ncbi.nlm.nih.gov/pmc/articles/>
-