

[CASE REPORT]

Group A Streptococcal Peritonitis and Toxic Shock Syndrome in a Postmenopausal Woman

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Abstract:

We herein report the case of a 66-year-old woman presenting with symptoms of gastroenteritis. Computed tomography showed small-bowel dilation without ischemic signs. After admission, she went into shock and was treated for sepsis of unknown origin. She was later diagnosed with group A streptococcal peritonitis due to an ascending vaginal infection. This case highlights the importance of considering Group A *Streptococcus* (GAS) infection as a cause of peritonitis in postmenopausal women.

Key words: Group A Streptococcus, streptococcal toxic shock syndrome, primary peritonitis

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Introduction

Although Group A *Streptococcus* (GAS) infections of the female reproductive tract are occasionally seen in premenopausal women, cases in postmenopausal women are rare. We describe a case of GAS peritonitis in a previously healthy postmenopausal woman to raise awareness about this potentially fatal condition.

Case Report

A 66-year-old woman presented to our emergency department with abdominal pain, vomiting, and watery diarrhea of 2 days in duration. Two days prior to her admission to our hospital, she visited her community gynecologist after experiencing clear vaginal discharge of 3 days in duration. She underwent a gynecological examination, including a Pap smear and vaginal swab for culture. She had no invasive procedures such as biopsy. The patient noted that the pelvic examination that was performed by the community gynecologist was rather more painful than usual.

The patient had no significant medical history other than an appendectomy at 55 years of age. She had undergone treatment for dental cavities a few weeks prior to her admission. She was not on any medication. She did not report a history of using tobacco, alcohol, or illicit drugs. She was a nurse and worked at a community clinic once a week. She had no symptoms of upper respiratory infection and had not been exposed to patients with such symptoms. The patient had no history of sexually transmitted diseases and had been abstinent from sexual intercourse for 10 years.

On admission, the patient was alert and fully oriented, but was clearly in distress. A physical examination revealed the following findings: systolic blood pressure, 89 mmHg; pulse, 93 beats/min; respiratory rate, 30 breaths/min; and body temperature, 38.8°C. Her abdomen was soft and seemed to show signs of intermittent guarding. She also showed a translucent vaginal discharge.

The laboratory data showed leukocytosis $(9,400/\mu L)$, elevated C-reactive protein (15.2 mg/dL) and creatinine (1.69 mg/dL; normal range, 0.46-0.79 mg/dL), and metabolic acidosis (pH 7.29, pCO₂ 39.8 mmHg, HCO₃ 18.5 mEq/L). Her liver function test results were within the normal ranges. Computed tomography (CT) showed small-bowel dilation without ischemic signs, and mild ascites in the pelvis. The patient's uterus seemed large for a postmenopausal woman and showed irregular enhancement, but this was not deemed to be significant at the time (Figure).

She was admitted for close monitoring with a diagnosis of enteritis of unknown origin. We initially infused her with large amounts of intravenous fluids. Eight hours after admission, her condition showed rapid deterioration, and she developed septic shock with disseminated intravascular coagu-

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Figure. Computed tomographic scan of the pelvis showing irregularly enhanced uterus (white arrow).

lation [systemic inflammatory response syndrome (SIRS) score 3, 0.18 γ noradrenaline required to maintain mean arterial pressure >65 mmHg, lactate 2.34 mmol/l, platelet 116,000/ μ L, prothrombin time-international normalized ratio (PT-INR) 1.7, fibrinogen degradation products (FDP) 46 μ g/mL]. As peritoneal infection following a vaginal procedure was suspected, intravenous meropenem (2 g/day) was initiated, along with thrombomodulin.

Based on the suspicion of an ascending infection from the genital tract and the observation of uterine enlargement on CT, the gynecology department was consulted. As the patient was in a critical condition, antibiotic treatment was initiated before the gynecological examination. A pelvic examination revealed normal leucorrhea with mild cervical motion tenderness. Pelvic inflammatory disease was thought to be unlikely as there was dissociation between the severity of the findings from the pelvic examination and her overall status. Some white blood cells were seen on Gram staining of the vaginal discharge, but there were no signs of bacteria or fungi.

On the 2nd day of hospitalization, the blood cultures taken on admission grew Gram-positive *Streptococci*. Cardiologists were consulted to rule out infectious endocarditis, but transthoracic echocardiography showed no vegetation. Repeat CT showed uterine enlargement, cervical wall thickening, and a pelvic abscess, strongly suggesting an ascending infection from the vagina.

By the 3rd day of hospitalization, the patient's condition slowly began to improve. The blood cultures grew GAS, and the vaginal swab taken at the community gynecologist 2 days before admission was also reported to grow GAS. The vaginal culture taken after admission remained negative. Mtyping of the isolates was not performed. No other organisms were isolated. She was diagnosed with primary GAS peritonitis caused by ascending infection from the vagina. As the formation of the pelvic abscess suggested a polymicrobial infection, ampicillin/sulbactam (12 g/day) and clindamycin (1,800 mg/day) were administered. Intravenous antibiotics were continued for 16 days until the pelvic abscess decreased in size on a follow-up CT examination. The patient eventually made a complete clinical recovery and was discharged on oral antibiotics. The antibiotics were continued until the complete resolution of the pelvic abscess was confirmed on follow-up.

Discussion

Although GAS is not considered to be a part of the normal flora of the female genital tract, we found 86 cases (65 reports) of primary peritonitis caused by GAS (1-65), most of which were in women (female:male ratio approximately 7:1), suggesting an ascending genital route of infection. GAS peritonitis has been reported to occur in premenopausal women, mostly in association with procedures such as intrauterine device (IUD) insertion (14, 24, 64).

Streptococcal infection in a postmenopausal woman in the absence of any precipitating factor is rare. It is suggested that physiologically, the cervical mucus of postmenopausal women is more tenacious and serves as a mechanical barrier to ascending infections (66). Decreased sexual activity, the likelihood of high-risk procedures such as abortions as well as the use of IUDs may be the reason that primary peritonitis is rare in postmenopausal women. However, this would also mean that postmenopausal women that are sexually active or who have transvaginal procedures are at risk of primary GAS peritonitis. It is not possible to say whether the Pap smear played a role in the development of peritonitis in the present case, as the procedure is generally considered to be noninvasive. Considering that our patient had an increase in vaginal discharge before the gynecological examination, we believe that the patient would have developed peritonitis regardless of the procedure. If the Pap smear were the cause of the patient's peritonitis, this would be the first reported case.

As most reports did not clearly state whether the women were menopausal, we divided the reported cases into two groups (Table). Ascending infection from the vagina was suspected in 22 of the cases in women of <50 years of age, nearly half of whom had a precipitating factor. The entry site remained unclear in most women of \geq 50 years of age, but six cases were thought to have been associated with an ascending infection from the vagina, none of which had precipitating factors.

In our case, the age and the fact that the patient waited two days before visiting the emergency department may explain the severity. Vaginal atrophy and the loss of vaginal acidity brought about by menopause are other possible risk factors for vaginal infections in postmenopausal women (59, 67).

The symptoms of GAS infection tend to be poorly localized and mimic common diseases. More than half of the 28 cases in which ascending infection from the vagina was suspected, presented with symptoms that were similar to infectious enterocolitis (for example, abdominal pain, nausea and diarrhea). We noticed that a pelvic examination was not mentioned in many of the cases. Considering that it is diffi-

Age	<50	≥50
Number of patients	55	20
Possible entry site	vagina(total): 22	vagina(total): 6
	vagina(IUD/abortion): 10	vagina(IUD/abortion): 0
	URI: 14	URI: 2
	other: 5	other: 1
	unclear: 14	unclear: 11
Pelvic examination	20	9
Surgical debridement	42	11
Outcome	well: 49	well: 16
	dead: 4	dead: 3
	unknown: 2	unknown: 1
Mortality		
with surgery	4.8% (2/42)	0% (0/11)
without surgery	15.4% (2/13)	33% (3/9)

Table.GAS Peritonitis in Women.

GAS: Group A *Streptococcus*, IUD: intrauterine device, URI: upper respiratory tract infection

cult to deduce the portal of entry based on the symptoms, careful history taking and a thorough physical examination, including a pelvic examination, should be performed.

Rimawi et al. stated that surgical therapy to remove the source of GAS and its toxin production is beneficial in the treatment of streptococcal toxic shock syndrome (STSS) (68). This theory may explain the high rate of exploratory laparotomy and drainage in GAS peritonitis (53 cases). Venkataramanasetty et al. and Gisser et al. reported no improvement in their patients despite early aggressive management with intravenous antibiotics, intravenous fluids, and IUD removal (30, 48). In both cases, improvement was not seen until the patients' infected tissues were debrided. Among the patients who received surgical treatment, the mortality rate was 3.7% (2 out of 53 patients), while the mortality rate among the patients who did not receive surgical treatment was 22.7% (5 out of 22 patients). These results support Rimawi's theory.

However, some patients with GAS peritonitis can be treated with appropriate antibiotics. In our case, CT-guided pelvic abscess drainage was considered, but because the improvement of the abscess was confirmed on follow-up CT, antibiotics were continued, which eventually led to complete resolution of the abscess. The patient in our case is the oldest reported woman with a suspected ascending vaginal infection caused by GAS who survived without surgical debridement. The need for surgical intervention should be carefully considered depending on the severity of the peritonitis and the clinical course after the prompt initiation of proper antibiotic therapy. To aid in the diagnosis, less invasive methods such as paracentesis of ascites, Gram staining, and rapid antigen detection tests may also be beneficial.

Conclusion

Primary GAS peritonitis in postmenopausal women is

rare. As prompt treatment with appropriate antibiotics may prevent the need for surgical intervention, careful history taking and a thorough physical examination are required. Ascending infection from the vagina should always be considered in postmenopausal women if there is no other obvious cause, even when the patient is sexually inactive.

The authors state that they have no Conflict of Interest (COI).

References

- Sanchez NC, Lancaster BA. A rare case of primary group A streptococcal peritonitis. Am Surg 67: 633-634, 2001.
- Thomas D, Perpoint T, Dauwalder O, et al. *In vivo* and *in vitro* detection of a superantigenic toxin Vbeta signature in two forms of streptococcal toxic shock syndrome. Eur J Clin Microbiol Infect Dis 28: 671-676, 2009.
- Kinsella A, Kavanagh DO, McGiobuin S, Schlaffer K, Evoy D. Primary peritonitis from an insect bite. Ir Med J 102: 87-88, 2009.
- **4.** Gavala A, Klimopulos S, Exarchos D, et al. Persistent primary peritonitis due to group A *Streptococcus* and *E. coli*. Intensive Care Med **28**: 1829-1831, 2002.
- Kanetake K, Hayashi M, Hino A, et al. Primary peritonitis associated with streptococcal toxic shock-like syndrome: report of a case. Surg Today 34: 1053-1056, 2004.
- Gribbin JC, Cox CJ. Spontaneous bacterial peritonitis in a healthy adult male. Aust N Z J Surg 60: 723-725, 1990.
- Gelshorn C, Piffaretti JC, Haldimann B, Martinoli S. [Diarrhea and peritonitis in infection caused by type A beta hemolytic streptococcus]. Helv Chir Acta 60: 931-934, 1994.
- Hoshino N, Hasegawa H. Two cases of primary peritonitis due to *Streptococcus pyogenes*. Nihon Fukubu Kyukyu Igakukai Zasshi (J Abdom Emerg Med) 30: 697-701, 2010 (in Japanese, Abstract in English).
- Kubota T, Morimoto Y, Imaoka K. Group A streptococcal primary peritonitis with abdominal compartment syndrome. Geka (Surgery) 70: 911-915, 2008 (in Japanese, Abstract in English).
- Tanaka H, Masahisa Y. A case of primary peritonitis and empyema associated with streptococcal toxic shock-like syndrome. Nihon

Kyukyu Igakukai Zasshi (J Jpn Assoc Acute Med) **24**: 357-362, 2013 (in Japanese, Abstract in English).

- Monneuse O, Tissot E, Gruner L, et al. Diagnosis and treatment of spontaneous group A streptococcal peritonitis. Br J Surg 97: 104-108, 2010.
- Goepel JR, Richards DG, Harris DM, Henry L. Fulminant Streptococcus pyogenes infection. Br Med J 281: 1412, 1980.
- Troillet N, Leuenberger A, de Werra P, Praz G. [Invasive Streptococcus pyogenes infection (beta-hemolytic Streptococcus of group A)]. Schweiz Med Wochenschr 124: 1064-1069, 1994.
- Scott RB. Critical illnesses and deaths associated with intrauterine devices. Obstet Gynecol 31: 322-327, 1968.
- **15.** Gendron N, Joubrel C, Nedellec S, et al. Group A *Streptococcus* endometritis following medical abortion. J Clin Microbiol **52**: 2733-2735, 2014.
- 16. Saha P, Morewood T, Naftalin J, Hopkins S. Acute abdomen in a healthy woman: primary peritonitis due to group A *Streptococcus*. J Obstet Gynaecol 26: 700-701, 2006.
- Legras A, Lodico R, Ferre R, Valleur P, Pautrat K. Primary peritonitis due to *Streptococcus* A: laparoscopic treatment. J Visc Surg 148: e315-e317, 2011.
- Malota M, Felbinger TW, Ruppert R, Nussler NC. Group A *Strep-tococci*: a rare and often misdiagnosed cause of spontaneous bacterial peritonitis in adults. Int J Surg Case Rep 6C: 251-255, 2015.
- Manalo R, Mirza H, Opal S. *Streptococcus pyogenes* tuboovarian abscess: a potential sexually transmitted disease? Sex Transm Dis 29: 606-607, 2002.
- Khoury GA, Wall RA. Streptococcal peritonitis associated with the cathartic colon. Br J Surg 69: 327, 1982.
- 21. Haap M, Haas CS, Teichmann R, Horger M, Raible A, Lamprecht G. Mystery or misery? Primary group A streptococcal peritonitis in women: case report. Am J Crit Care 19: 454-458, 2010.
- Brosseau JD, Mazza JJ. Group A streptococcal sepsis and arthritis. Origin from an intrauterine device. JAMA 238: 2178, 1977.
- 23. van Lelyveld-Haas LE, Dekkers AJ, Postma B, Tjan DH. An unusual cause of a spontaneous bacterial peritonitis in a young healthy woman. N Z Med J 121: 82-85, 2008.
- 24. Marshall BR, Hepler JK, Jinguji MS. Fatal Streptococcus pyogenes septicemia associated with an intrauterine device. Obstet Gynecol 41: 83-87, 1973.
- **25.** Monif GR, Williams BT, Dase DF. Group A *Streptococcus* as a cause of endometritis/salpingitis/peritonitis in a nongravid female. Obstet Gynecol **50**: 509-510, 1977.
- 26. Okumura K, Schroff R, Campbell R, Nishioka L, Elster E. Group A streptococcal puerperal sepsis with retroperitoneal involvement developing in a late postpartum woman: case report. Am Surg 70: 730-732, 2004.
- 27. Park JY, Moon SY, Son JS, Lee MS, Jung MH. Unusual primary peritonitis due to *Streptococcus pyogenes* in a young healthy woman. J Korean Med Sci 27: 553-555, 2012.
- 28. Jarvis J, Trivedi S, Sheda S, Frizelle FA. Primary peritonitis in adults: is it time to look for a better diagnostic classification? ANZ J Surg 76: 127-129, 2006.
- **29.** Golden GT, Stevenson TR, Ritchie WP Jr. Primary peritonitis in adults. South Med J **68**: 413-414, 1975.
- **30.** Venkataramanasetty R, Aburawi A, Phillip H. Streptococcal toxic shock syndrome following insertion of an intrauterine device--a case report. Eur J Contracept Reprod Health Care **14**: 379-382, 2009.
- Fikrig E, Worthington MT, Lefkowitz LB Jr. Septic shock and acute respiratory distress syndrome after salpingitis caused by *Streptococcus pyogenes* group A. South Med J 82: 634-635, 1989.
- 32. Brase R, Kuckelt W, Manhold C, Bohmert F. [Spontaneous bacterial peritonitis without ascites]. Anasthesiol Intensivmed Notfallmed Schmerzther 27: 325-327, 1992.
- 33. Ohnishi K, Nakanishi M, Ogaki Y, et al. A case of group A strep-

tococcal infection presented as pelvic inflammatory disease. Tokyo Sanka Fujinka Gakkai Kaishi (Tokyo J Obstet Gynecol) **61**: 419-423, 2012 (in Japanese).

- 34. Doloy A, Godin C, Decousser JW, Panel P, Greder-Belan A, Doucet-Populaire F. Primary peritonitis due to *Streptococcus pyogenes* with reduced susceptibility to fluoroquinolones. Diagn Microbiol Infect Dis 62: 447-449, 2008.
- 35. Kobayashi M. Primary group A streptococcal peritonitis complicated by acute renal failure. Nihon Kyukyu Igakukai Zasshi (J Jpn Assoc Acute Med) 11: 118-122, 2000 (in Japanese, Abstract in English).
- 36. Barham WB, Haberberger RL, Decker CF. Group A streptococcal sepsis secondary to acute salpingitis. Clin Infect Dis 16: 444-445, 1993.
- 37. Borgia SM, Low DE, Andrighetti S, Rau NV. Group A streptococcal sepsis secondary to peritonitis and acute pelvic inflammatory disease. Eur J Clin Microbiol Infect Dis 20: 437-439, 2001.
- 38. Tsuruta H, Fukui M, Kooguchi K, Shimosato G, Beppu S. A case of streptococcal toxic shock syndrome presented as primary peritonitis. Nihon Shuchu Chiryo Igakukai Zasshi (J Jpn Soc Intensive Care Med) 10: 213-214, 2003 (in Japanese).
- 39. Souma Y. A case of primary peritonitis caused by group A betahemolytic streptococcal pelvic peritonitis. Nihon Gekakei Rengo Gakkaishi (J Jpn Coll Surg) 30: 215-219, 2005 (in Japanese).
- Graham JC, Moss PJ, McKendrick MW. Primary group A streptococcal peritonitis. Scand J Infect Dis 27: 171-172, 1995.
- Garvey P, Ledger WJ. Group a streptococcus in the gynecologic patient. Infect Dis Obstet Gynecol 5: 391-394, 1997.
- **42.** Moskovitz M, Ehrenberg E, Grieco R, et al. Primary peritonitis due to group A *Streptococcus*. J Clin Gastroenterol **30**: 332-335, 2000.
- 43. Fox KL, Born MW, Cohen MA. Fulminant infection and toxic shock syndrome caused by *Streptococcus pyogenes*. J Emerg Med 22: 357-366, 2002.
- **44.** Tilanus AM, de Geus HR, Rijnders BJ, Dwarkasing RS, van der Hoven B, Bakker J. Severe group A streptococcal toxic shock syndrome presenting as primary peritonitis: a case report and brief review of the literature. Int J Infect Dis **14** Suppl 3: e208-e212, 2010.
- **45.** Nogami Y, Tsuji K, Banno K, et al. Case of streptococcal toxic shock syndrome caused by rapidly progressive group A hemolytic streptococcal infection during postoperative chemotherapy for cervical cancer. J Obstet Gynaecol Res **40**: 250-254, 2014.
- 46. Casadevall A, Pirofski LA, Catalano MT. Primary group A streptococcal peritonitis in adults. Am J Med 88: 63N-64N, 1990.
- 47. Mourton S, Rich W. Group A streptococcal toxic shock syndrome after an office endometrial biopsy: a case report. J Reprod Med 51: 665-668, 2006.
- 48. Gisser JM, Fields MC, Pick N, Moses AE, Srugo I. Invasive group A *Streptococcus* associated with an intrauterine device and oral sex. Sex Transm Dis 29: 483-485, 2002.
- 49. Lamb EK, Anasti JN, Leonetti HB. Group A *Streptococcus* causing PID from an initial pharyngeal infection. A case report. J Reprod Med 44: 639-641, 1999.
- Auskalnis S, Bogusevicius A, Maleckas A, Butrimavicius S, Toker I. [Primary peritonitis caused by group A beta-hemolytic *Strepto-coccus*]. Medicina (Kaunas) 40: 969-974, 2004.
- Cho EE, Fernando D. Fatal streptococcal toxic shock syndrome from an intrauterine device. J Emerg Med 44: 777-780, 2013.
- Christen RD, Moser R, Schlup P, Neftel KA. Fulminant group A streptococcal infections. Report of two cases. Klin Wochenschr 68: 427-430, 1990.
- 53. Van Den Bossche MJ, Devriendt D, Weyne L, Van Ranst M. [Primary peritonitis combined with streptococcal toxic shock syndrome following an upper respiratory tract infection caused by *Streptococcus pyogenes*]. Ned Tijdschr Geneeskd 152: 891-894,

2008.

- 54. Imamura S, Aso M, Mii S, Sakata H, Kato H. A case of fulminant infection with *Streptococcus pyogens* presented with primary peritonitis. Nihon Rinsho Geka Gakkai Zasshi (J Jpn Surg Assoc) 64: 2879-2882, 2003 (in Japanese, Abstract in English).
- **55.** Brivet FG, Smadja C, Hilbert U, et al. Usefulness of abdominal CT scan in severe peritoneal sepsis linked to primary peritonitis. Scand J Infect Dis **37**: 76-78, 2005.
- 56. Bibler MR, Rouan GW. Cryptogenic group A streptococcal bacteremia: experience at an urban general hospital and review of the literature. Rev Infect Dis 8: 941-951, 1986.
- 57. Hikone M, Kobayashi K, Washino T, et al. Streptococcal toxic shock syndrome secondary to group A *Streptococcus* vaginitis. J Infect Chemother 21: 873-876, 2015.
- 58. Paulson Q, Douglass E, Moreno A, Aydelotte J. Group A streptococcal toxic shock syndrome secondary to necrotizing pelvic inflammatory disease in a postmenopausal woman. IDCases 5: 21-23, 2016.
- Paraskevaides EC, Wilson MC. Fatal disseminated intravascular coagulation secondary to streptococcal cervicitis. Eur J Obstet Gynecol Reprod Biol 29: 39-40, 1988.
- 60. Matsuyoshi T, Imamura T, Sasaki N, et al. A case of streptococcal toxic shock-like syndrome presented as primary peritonitis. Nihon Shuchu Chiryo Igakukai Zasshi (J Jpn Soc Intensive Care Med) 23: 61-62, 2016 (in Japanese).
- 61. Ugurlu G, van der Houwen C, Brandenburg A, Schreuder I, Bogchelman D. *Streptococcus pyogenes* vaginitis in a postmenopausal woman. Eur J Obstet Gynecol Reprod Biol 138: 115-116,

2008

- 62. Yamada K. A case of primary peritonitis due to group A *Streptococcus*. Kensei Byoin Iho (Med J Kensei) 36: 13, 2013 (in Japanese).
- **63.** Kouijzer IJ, Polderman FN, Bekers EM, Bloks PH, Schneeberger PM, de Jager CP. Initially unrecognised group A *Streptococcal* pelvic inflammatory disease in a postmenopausal woman. Neth J Med **72**: 494-496, 2014.
- **64.** Ledger WJ, Headington JT. Group a beta-hemolytic *Streptococcus*. Obstet Gynecol **39**: 474-482, 1972.
- **65.** Vuilleumier H, Halkic N. Streptococcal toxic shock syndrome revealed by a peritonitis. Case report and review of the literature. Swiss Surg **7**: 25-27, 2001.
- 66. Jackson SL, Soper DE. Pelvic inflammatory disease in the postmenopausal woman. Infect Dis Obstet Gynecol 7: 248-252, 1999.
- 67. Verstraelen H, Verhelst R, Vaneechoutte M, Temmerman M. Group A streptococcal vaginitis: an unrecognized cause of vaginal symptoms in adult women. Arch Gynecol Obstet 284: 95-98, 2011.
- **68.** Rimawi BH, Soper DE, Eschenbach DA. Group A streptococcal infections in obstetrics and gynecology. Clin Obstet Gynecol **55**: 864-874, 2012.

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