

Stenotrophomonas Maltophilia Peritonitis in a Child: Case Report and Review of the Literature

Bir Çocuk Olguda Stenotrophomonas Maltophilia Peritoniti: Olgu Sunumu ve Literatür Derlemesi

Olgu Sunumu
Case Report

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ABSTRACT

Stenotrophomonas maltophilia peritonitis has been only occasionally reported in patients undergoing continuous ambulatory peritoneal dialysis (CAPD). Because this microorganism has multi-drug resistance, its treatment is hard and long-term. The treatment might not be successful despite all the efforts and the process of peritoneal dialysis, and may terminate with loss of the catheter. In the present paper, S. maltophilia peritonitis developed in a 6-year-old girl patient, who underwent peritoneal dialysis due to bilateral dysplastic kidney, suffered from episodes of peritonitis frequently and required hospitalization, was presented with literature data. Even though the case received multiple antibiotic treatment and underwent endoluminal brushing (EB), the success of treatment could not be achieved. To the best of our knowledge, this patient is the youngest case in the literature.

Keywords: *Stenotrophomonas maltophilia, peritonitis, child*

Öz

Stenotrophomonas maltophilia peritoniti, sürekli ayaktan periton diyalizi (SAPD) geçiren hastalarda nadiren bildirilmiştir. Mikroorganizma çoklu ilaç direncine sahip olduğu için tedavisi uzun ve zordur. Tüm çabalara ve periton diyalizi sürecine rağmen tedavi başarılı olmayabilir, kateter kaybı ile sona erebilir. Bu yazıda, bilateral displastik böbrek nedeniyle periton diyalizi uygulayan, peritonit ataklarından sıkıntı çeken ve hastaneye yatırılan 6 yaşındaki bir kız hastada gelişen S. maltophilia peritoniti, literatür verileri ile sunuldu. Olguya çoklu antibiyotik tedavisi uygulanmış ve endoluminal fırçalama (EB) yapılmış olmasına rağmen tedavinin başarısı sağlanamamıştır. Bildiğimiz kadarıyla, bu hasta literatürdeki en genç vakadır.

Anahtar kelimeler: *Stenotrophomonas maltophilia, peritonit, çocuk*

Received/Geliş: 12.07.2019
Accepted/Kabul: 07.10.2019
Published Online: 05.01.2021

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INTRODUCTION

Stenotrophomonas maltophilia (*S. maltophilia*) an opportunistic pathogen is a free living, motile, aerobic, oxidase-negative, glucose non-fermentative, gram-negative, multidrug-resistant bacillus prevalent particularly among inpatients⁽¹⁾. It can be frequently isolated from water, earth, animals, plants, and hospital equipments⁽²⁾. *S. maltophilia* is the only one species of the genus

Stenotrophomonas which is known to infect humans. It was isolated from pleural fluid in 1943 by Edward for the first time and named as *Bacterium brokeri*^(3,4). Hugh and Ryschenkow reclassified and named it as *Pseudomonas maltophilia* in 1961. Twenty years later, Swings et al., named *P. maltophilia* as *Xanthomonas maltophilia*. Finally, Palleroni and Bradbury gave the last and up-to-date name, *S. maltophilia*, to the microorganism in 1993⁽³⁾. *S. maltophilia* leads to numerous

Cite as: Alaygut D, Alparslan C, Sarıtař S, et al. *Stenotrophomonas maltophilia peritonitis in a child: Case report and review of the literature*. Tepecik Eđit. ve Arařt. Hast. Dergisi. 2020;30(3):316-21.



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different infections such as bacteremia, endocarditis, respiratory tract infections, meningitis, urinary tract infections, skin and soft tissue infections, mastoiditis, bone and joint infections, peritonitis, typhlitis and biliary sepsis, wound infections, and central venous catheter-related infections in immunocompromised people⁽⁴⁾. Risk factors are considerably variable and Table 1 shows these risk factors⁽¹⁾. *S. maltophilia* was previously reported to be the cause of both peritonitis and infections at the exit site in patients undergoing chronic peritoneal dialysis. This microorganism, which is resistant and hard to control via medical treatment, is substantially important because it may lead to catheter loss and paves the way for the growth of other opportunist microorganisms. In this paper, a 6-year-old child who underwent chronic peritoneal dialysis and had to start the hemodialysis program by losing peritoneal dialysis catheter because of *S. maltophilia* growth was presented with the literature data. To the best of our knowledge, this case is the youngest one in the literature.

Table 1. Risk factors for *S. maltophilia* infections.

Malignancy, particularly hematological malignancy
Human immunodeficiency virus (HIV)
Cystic fibrosis
Intravenous drug abuse
Surgical and accidental trauma
Prolonged hospitalization
Admission to ICU and mechanical ventilation
Vascular catheters and urinary catheters
Corticosteroids and immunosuppressive therapy
Prior treatment with broad-spectrum antibiotics
Gastrointestinal tract colonization and mucositis
Hematopoietic stem cell transplantation
Travel to hospital by air

CASE REPORT

The 6-year-old girl patient who had the end-stage renal failure due to bilateral dysplastic kidney and was in peritoneal dialysis program (CAPD) for 2 years, was hospitalized in the service with pre-diagnosis of peritonitis upon lack of appetite, vomiting, stomach ache, fever, and cloudy dialysis fluid 3 days before her admission. From her history, it was found out that she was previously treated 5 times

due to peritonitis and she had the last peritonitis 3 months previously. Some parameters measured were as follows: body weight: 12.9 kg (3p), height: 90 cm (3p), heart rate: 94/min, respiratory rate: 20/min, blood pressure: 129/100 (>95.p/>95.p) mmHg, and body temperature: 38°C. Besides abdominal distension and sensitivity were detected on physical examination. Some biochemical values were as follows: white blood cell /WBC):13.100 U/L, C-reactive protein 164.7 mg/L (N: 0-5), procalcitonin 9.4 ng/mL (N:0-0.1), creatinine: 4.6 mg/dL, urea:86 mg/dL, K 6.9 mg/dL, and serum albumin 3.1 mg/dl. Peritoneal fluid was cloudy and microscopic examination contained >1000 cell/mm³. Empirical antibiotic treatment was initiated with intraperitoneal doses of cefazolin (loading dose: 500 mg/L, and maintenance dose: 125 mg/L) and ceftazidime (loading dose: 500 mg/L, and maintenance dose: 125 mg/L) before obtaining blood and peritoneal fluid cultures of the case. *S. maltophilia* grew in peritoneal fluid culture. Microorganism was sensitive to TMP-SMX, levofloxacin, and tigecycline, moderately sensitive to ceftazidime, and resistant to other antibiotics. Cefazolin treatment was terminated and trimethoprim- sulfamethaxazole (co-trimoxazole) was started at systemic renal dose (5 mg/kg/day), and intraperitoneally (TMP-SMX loading dose: 320/160 mg, maintenance dose: 80/400 mg). The case whose cell count was monitored daily underwent peritoneal endoluminal brushing process twice and cell count declined down to 20 /mm³. However, the case was accepted to be resistant peritonitis because the same microorganism grew again in the culture of the control peritoneal fluid which was sent 2 more times while the patient was under treatment. Systemically administered TMP-SMX was stopped and levofloxacin (8 mg/kg/day) was started intravenously. Intraperitoneal ceftazidime and TMP-SMX was continued. Fluconazole was also started as antifungal because of multiple antibiotic treatment. WBC was not identified in peritoneal fluid on the 5th day of levofloxacin treatment. Thrombocytopenia developed on the 8th day of treatment, and pancytopenia on the day 10 which were

Table 2. Demographic and clinical characteristics of patients with Stenotrophomonas maltophilia infection (peritonitis and/ or exit site infection) in literature.

Patient no	Age	Gender	RD	RF/ co-morb.	Period of dialysis (months)	No of previous infection	Time of the last infection	Type of infection	Treatment	Infection duration (weeks)	Outcome	Ref.
1	61	F	DN	MI	32	3	20 mo.ago	P	Cefaz (IP) + Tob(IP) Ceft (IP) + Amk (IP) Ceft (IP) + Amk (IP) + Pip (IV)	23	Catheter removal	Baek et al 5
2	34	F	DN	PH	24	1	18 mo ago	P	Cefaz (IP) + Tob (IP)	3	Continued PD	Baek et al 5
3	48	F	DN	No	15	2	1 mo.ago	P	Van (IP) + Ceft (IP)+ Cjpr (IP) + Tmp-Smx (IP) + Ceft (IP) Amphotericin (IV)	5	Catheter removal, HD	
4	62	M	RT	CVA	12	0	7 mo.ago	ESI	Ceft(IV,IP) + Tob (IP)	2	Continued PD	Baek et al 5
5	50	M	No data	COPD	CHF	5	23 mo.ago	ESI	1. Cjpr (PO) + Cefac (PO) 2. Cjpr (PO) + Amxcv (PO)	1 st : 2 2 nd : 2	Continued PD	Baek et al 5
6	54	F	Alport dis.	No	23	0	-	P	Ceft (IP) + Van (IP) Tmp-Smx (IP) + Amk (IP)	2	Continued PD	Baek et al 5
7	No data	No data	No data	No data	No data	0	-	P	→ Van (IP) + Imip (IP) → Ceft (IP) + Gent (IP) → Cjpr	No data	Catheter removal	Machuca E et al 21
8	No data	No data	No data	No data	No data	1	8 wk. ago	P	Van (IP) + Imip (IP) Ceft Cjpr Tmp-Smx	No data	Catheter removal	Szeto et al 6
9	No data	No data	No data	No data	No data	3	2 wk.ago	P	Van (IP) + Imip (IP) Ceft (IP) + Neti (IP)	No data	Catheter removal	Szeto et al 6
10	No data	No data	No data	No data	No data	1	7 wk ago	P	Van (IP) + Imip (IP) Ceft (IP) Cjpr (IP)	No data	Catheter removal	Szeto et al 6
11	No data	No data	No data	No data	No data	2	2 wk ago	P	Van (IP) + Imip (IP) Ceft (IP) + Neti (IP)	No data	Catheter removal	Szeto et al 6
12	No data	No data	No data	No data	No data	0	-	P	Van (IP) + Imip (IP) Ceft (IP) + Ampicillin (IP)	No data	Catheter removal	Szeto et al 6
13	54	M	Calsineurine toxiity	No data	No data	No data	No data	P	Van (IV) + Ceft (IP) Tmx –Smx (IP)	No data	HD	Szeto et al 6
14	57	F	DN	No	36	0	-	P	Van (IP) + Ceft (IP) Ceft (IP) + Levofl (IP)	2	Continued PD	Beatriz Millan-Diaz et al 12
15	63	M	DN	No	43	4	No data	P	Van (IP)+ Gent (IP) Ceft (IP) + Cipro (IP)	No data	Catheter removal, HD	Azak A etal 11
16	65	F	DN	No	19	2	No data	P	Van (IP) +Gent (IP) Ceft (IP)+ Cipro (IP)	No data	Catheter removal, HD	N.Al-Hilali et al 13
17	40	M	DN	No	17	0	-	ESI	Ceft (IP) + Amic (IP)	No data	Replaced, Continued PD	N. Al-Hilali et al 13
18	56	M	DN	No	12	1	No data	ESI	Tmp-Smx (oral)	No data	Not replaced, Continued PD	N. Al-Hilali et al 13
19	35	F	No data	No data	64	No data	No data	ESI	Tmp-Smx (oral)	30 day	Continued PD	N. Al-Hilali et al 13
20	75	M	No data	No data	11	No data	No data	ESI, P	Parenteral antibiotic	455 days	Catheter removal	Dapena F et al 8
21	30	F	No data	No data	20	No data	No data	ESI	Tmp-Smx (oral)	500 days	Continued PD, Granuloma	Dapena F et al 8
22	37	M	No data	No data	59	No data	No data	ESI	Tmp-Smx (oral)	7 day	Continued PD	Dapena F et al 8
23	39	M	No data	No data	53	No data	No data	ESI	Tmp-Smx (oral)	120 days	Continued PD	Dapena F et al 8
24	36	F	No data	No data	36	No data	No data	ESI	Tmp-Smx (oral)	7 day	Continued PD	Dapena F et al 8
25	74	F	No data	No data	32	No data	No data	ESI	Tmp-Smx (oral)	24 days	Continued PD	Dapena F et al 8
26	39	M	No data	No data	8	No data	No data	ESI	Parenteral antibiotics	45 days	Continued PD	Dapena F et al 8
27	60	M	CPN	No data	96	Repeatedly	12 mo ago	P	Ceft (IP) + Tmp-Smx (IV)	3	Continued PD	Dapena F et al 8
28	64	F	PKD	No data	120	2	84 mo ago	P	Ceft (IP) + Amc (IP) Tmp-Smx (IV)	6	Continued PD	Tzanetou K et al 10
29	64	F	CGN	No data	96	0	No data	P	Van (IP) + Ceft (IP) Amc (IP)+Cjpr (IV)+Tmp-Smx (IV)	No data	Continued PD	Tzanetou K et al 10
30	40	M	CGN	No data	96	1	No data	P	Tmp-Smx (IV)	No data	Continued PD	Tzanetou K et al 10
31	No data	No data	N	No data	96	0	No data	P	Tmp-Smx (IV) + ticarcillin-clavulonate (IV)	No data	Cathetere replacement	Tzanetou K et al 10
32	61	M	No data	No data	60	1.6 *	No data	P	No data	No data	Continued PD	Tzanetou K et al 10
33	64	M	No data	No data	9	1.3*	No data	P	No data	No data	Continued PD	Taylor G et al 9
34	52	F	No data	No data	26	0.0*	No data	P	No data	No data	Catheter removal	Taylor G et al 9
35	19	F	No data	No data	68	0.9*	No data	P	No data	No data	Catheter removal	Taylor G et al 9
36	16	F	No data	No data	6	2.0*	No data	P	No data	No data	Cathetere removal HD	Taylor G et al 9
37	43	F	No data	No data	99	0.7*	No data	P	No data	No data	Continued PD	Taylor G et al 9
38	16	M	No data	No data	1	0.0*	No data	P	No data	No data	Cathetere removal, renal failure resolved	Taylor G et al 9
Our case	6	F	DK	RH	26	5	3	P	Ceph (IP) + Ceft (IP) Tmp-Smx (IP) + Ceft (IP) + Tmp-Smx (IV) + endoluminal brushing (2 times) Tmp-Smx (IP) + Ceft (IP) + Levofloxacin (IV)	4	Catheter removal, HD	Taylor G et al 9

Amk: Amikasin, Ceft: Ceftazidime, CVA: Cerebrovascular accident, COPD: Chronic obstructive pulmonary disease, CGN: Chronic glomerulonephritis, CHF: Congestive heart failure, DN: Diabetic nephropathy, DK: Dysplastic kidney, ESI: Exit-site infection, MI: Myocardial infarction, N: Nephrolithiasis, P: Peritonitis, PH: Panhypopituitarism, PKD: Polycystic kidney disease, RD: Renal disease, RH: Recurren hospitalization, RT: Renal tuberculosis, RF: Risk factors, Van: Vancomisin

evaluated as the side effect of levofloxacin. Treatment was terminated. Bacterial growth was not observed in the culture of the control peritoneal fluid. Intraperitoneal treatment was completed within 21 days and then discontinued. The patient who re-applied to the clinic after 24 hours with the complaints of deteriorated general condition, widespread abdominal distension, vomiting, and high fever which were evaluated as being compatible with sepsis and peritonitis. Peritoneal fluid was cloudy and a gelatin-like structure was forming in a short time. In the microscopic examination >1000 cell/mm³ were determined. Vancomycin, meropenem, and fluconazole were started systemically. Catheter was removed and hemodialysis treatment was started. The case was discharged after completion of systemic treatments within 21 days.

DISCUSSION

S. maltophilia has been continuing to confront us as a nosocomial pathogen with ever-increasing prevalence. It is more frequent especially in immunosuppressed individuals and Table 1 shows risk factors. Even though patients undergoing peritoneal dialysis were not listed in this table, both diseases leading to comorbidities in these people and negative effects of uremia on immune system make them risky. In their study, Baek et al. ⁽⁵⁾ indicated that even though it was not statistically significant, serum albumin, hemoglobin, creatinine and BUN values of 5 patients were lower than those in other CAPD patients and uremia, malnutrition, and anemia were nonspecific suppressive factors of immune function. Our CAPD patient had malnutrition. Furthermore, she had additional risk factors as recent history of peritonitis, use of broad-spectrum antibiotic, and frequent hospitalizations. The last antibiotic treatment was the most important risk factor reported in occurrence of peritonitis ⁽⁶⁾. Particularly, use of broad-spectrum antibiotics is a risk factor for the growth of opportunistic and multiple-drug resistant organisms. It is stated to occur when imipenem is used frequently ⁽⁷⁾. The use

of imipenem is not known because our case had been treated in another center previously. However, it was found out that she had peritonitis frequently and the last episode of peritonitis was experienced 3 months ago.

When we searched the literature with the key words: "*S. maltophilia*, *Xanthomonas maltophilia* and *P. Maltophilia* and *peritonitis*", we reached 38 cases with peritonitis and exit site infection (ESI) Age, gender, underlying renal disorders, risk factors, comorbid diseases, length of dialysis treatment (in months), the number of previous episodes of peritonitis, the time elapsed since the last peritonitis, the type of infection, treatments provided, duration of infection, and clinical results of the cases were reviewed and shown in Table 2. Firstly, our case was remarkable because she was the youngest one identified in the literature. The biggest patient series with 8 patients was reported by Dapena F et al., ⁽⁸⁾ and they compared ESI associated with *X. Maltophilia* in 8 CAPD patients with ESI and 15 patients with *P. Aeruginosa*-associated ESI (Table 2 Patient no 19-26). One of these cases (No 20) had concurrent peritonitis. Except for two patients, oral TMP-SMX treatment was administered and no patients experienced catheter loss except for one. While the patient number 19 experienced 3 episodes, the patient number 25 experienced 4 episodes. When compared with pseudomonas-associated ESI, patients with *X. Maltophilia* had a better prognosis.

The other large series with 7 patients was reported by Taylor G et al. ⁽⁹⁾. All of these patients had peritonitis. It was reported that two patients received an immunosuppressive treatment. One of these patients received immunosuppressive treatment due to cardiac transplantation, and the other one used cyclosporine due to Wegener granulomatosis. Peritoneal dialysis of four patients was stopped. All patients had catheter loss in 6-patient series of Szeto et al. ⁽⁶⁾. Average age of these patients was 52, and they had chronic glomerulonephritis (n=5), and

polycystic kidney (n=1). One of 5 patients reported by Tzanetou K et al. ⁽¹⁰⁾ had catheter loss. Baek et al. ⁽⁵⁾, reported a total of 5 cases including 2 patients with ESI and 3 patients with peritonitis, and 2 patients had catheter loss. Azak A et al. ⁽¹¹⁾, reported a single case, Beatriz Millan Diaz et al. ⁽¹²⁾, reported single case, and N.Al-Hilali et al. ⁽¹³⁾ reported 4 cases.

S. maltophilia is known to be multi-drug resistant and is the reason of catheter loss in patients undergoing peritoneal dialysis. Being resistant to numerous drugs also makes it harder to select antibiotics. The organism was reported to be resistant to imipenem and meropenem from carbapenems, tobramycin and gentamicin from aminoglycosides, amoxicillin, clavulanic acid, a majority of cephalosporins (except for ceftazidime), quinolones, and numerous antipseudomonal penicillin ^(3,4,14,15). In the study of CANWARD conducted between 2007 and 2011 in Canada, 22.746 clinical isolates were evaluated, 1.4% consisted of *S. maltophilia* and in vitro activity for tigecycline was found to be good ⁽¹⁶⁾. Different studies reported that the use of following combinations would create a synergistic effect: TMP-SMX and ticarcillin-clavulanate, TMP-SMX and ceftazidime, ticarcillin-clavulanate and levofloxacin, ticarcillin-clavulanate and aztreonam, ceftazidime-ciprofloxacin ^(17,18). IP treatment was changed to treatment with ceftazidime and TMP-SMX because resistance to empirically started cefazolin was also observed in our case. Additional systemic treatment with TMP-SMX was started. However, systemic treatment was replaced with levofloxacin as culture-negativity could not be shown. Patient had thrombocytopenia (platelets 38.000 mm³) which was considered to develop in association with the drug treatment and then pancytopenia. Drug-associated thrombocytopenia may lead to mild to severe thrombocytopenia and it was also reported with fluoroquinolone group of drugs and recovery was observed when the drug was discontinued ⁽¹⁹⁾. Thrombocytopenia of our case recovered as well by discontinuing medication (it

was discontinued on the 10th day of treatment). Bacterial growth was not observed in the culture at the end of treatment.

Differently from the cases in the literature, we additionally applied endoluminal brushing (EB) to our patient two times as the peritoneal fluid contained greater number of resistant cells. Biofilm layer occurring on the catheter in resistant and persistent peritonitis is known to have a crucial role. Antibiotic efficacy is considered to be decreased by reducing this biofilm layer via EB ⁽²⁰⁾. EB was successful for two of the three pediatric cases who had resistant peritonitis and were reported by us as a referral center and there was no loss of catheter. However, in none of these cases *S. Maltophilia* was grown ⁽²⁰⁾. Nevertheless, we think that EB should be tried as a part of treatment before removing catheter. But, we could not succeed with this case.

Consequently, *S. Maltophilia* is a microorganism whose infections lead to generally poor prognosis in patients undergoing chronic peritoneal dialysis. Immune dysfunction created by uremia, use of antibiotics and repetitive hospitalizations due to recurrent peritonitis pose a risk in these patients. Although ESI alone is less risky in terms of catheter loss, paying attention to resistance pattern of the microorganism while selecting an antibiotic should not be forgotten.

Conflict of Interest: None.

Informed Consent: None.

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