# Massive diarrhoea and sepsis due to an infection with Neisseria meningitidis serogroup W

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# ABSTRACT

Invasive meningococcal disease is associated with significant mortality. Classic presentation consists of high fever, headache and neck stiffness. *Neisseria meningitidis* serogroup W may present with atypical symptoms, which complicates recognition. Furthermore, it is associated with a high case fatality rate.

#### KEY WORDS

Invasive meningococcal disease, *Neisseria meningitidis* serogroup W, diarrhoea, gastroenteritis.

## INTRODUCTION

Bacterial meningitis in adults is associated with significant morbidity and mortality. The most common causative agents are *Streptococcus pneumoniae* and *Neisseria meningitidis*. The classic clinical presentation of bacterial meningitis includes high fever, neck stiffness, headache and confusion. Even when diagnosed early and treated adequately, morbidity and mortality of bacterial meningitis remain high.<sup>1-4</sup>

Thirteen subtypes of *Neisseria meningitidis* have been identified, based on the antigenic structure of their polysaccharide capsule. The most prevalent serogroups in the Netherlands include B, C, W, and Y. Patients with meningitis caused by *Neisseria meningitidis* serogroup W (MenW) may have an atypical presentation with gastrointestinal complaints including profound vomiting and diarrhoea.<sup>5,6</sup>

Several countries have reported an increasing incidence of MenW carriage and infections.<sup>45,7-9</sup> In this case report,

#### What was known on this topic?

*Neisseria meningitidis* is known to cause meningitis, sepsis, and rarely, shock or multiple organ dysfunction syndrome. Classical signs and symptoms are high fever, neck stiffness, headache, and confusion. Even when treated promptly, these infections are correlated with high morbidity and mortality.

#### What does this add?

This case of an infection caused by *Neisseria meningitidis* serogroup W, presenting with massive diarrhoea without meningeal symptoms, clearly illustrates the atypical clinical presentation of this serotype. Timely recognition and prompt treatment of this serious infection is warranted. Invasive infection with Neisseria meningitidis serogroup W should be included in the differential diagnosis of every patient presenting with a septic shock and gastrointestinal complaints.

we present a patient with a severe infection due to MenW with massive diarrhoea as the main presenting symptom.

## CASE REPORT

A 23-year-old previously healthy female presented with complaints of nausea, vomiting, profound diarrhoea and generalised pain and cramps in her legs, especially on the soles of her feet, two days prior to her visit. On the second day of complaints, she became seriously ill and was admitted at the emergency department. Physical examination in the emergency department showed a

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pulse rate of 140 beats per minute, a blood pressure of 93/73 mmHg, a temperature of 35.8 °C, a respiratory rate of 38 breaths per minute, and an oxygen saturation of 95% with 3 liters of oxygen per minute by nasal canula. She was initially lethargic, with a Glasgow Coma Scale score of E4M4V1 without signs of neck stiffness. Her legs were cold and mottled without petechial rash. A chest X-ray showed no abnormalities. Laboratory results revealed lactic acidosis (lactate 9.0 mmol/l, pH 7.04, pO2 4.9 kPa, bicarbonate 10.0 mmol/l), leukopenia (0.8\*109/l), elevated CRP (332 mg/l), thrombocytopenia (5\*10<sup>9</sup>/l), and signs of disseminated intravascular clotting (DIC; platelet count 8\*109/l, PT 29.3 sec., D-dimer 10 mg/l, Fibrinogen 3.8 g/l). The differential diagnosis consisted of gastroenteritis with sepsis (after eating fish soup), atypical meningitis, or a toxic shock syndrome. A lumbar puncture was not performed because of severe thrombocytopenia and, according to the neurologist, her case presentation not being primarily neurological. Antibiotic treatment containing ceftriaxone (2 g IV twice a day) and dexamethasone (10 mg IV four times a day), was switched to a broader regimen consisting of meropenem (2 g IV twice a day), clindamycin (600 mg IV four times a day), and intravenous immunoglobulins (50 g IV once a day) because of deterioration within hours into a septic shock, for which noradrenalin and vasopressin infusion was required.

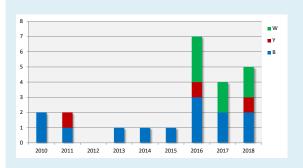
Ultimately, she developed multiple organ dysfunction syndrome (MODS), with mechanical ventilation because of exhaustion; septic cardiomyopathy, confirmed by transthoracic echocardiography for which inotropic agents were initiated; and fulminant DIC with petechiae on the extremities and haematemesis. In addition, because of acute kidney injury, renal replacement therapy was started. After two days of antibiotic treatment, blood cultures showed N. meningitidis, and subsequently antibiotic therapy was switched to benzylpenicillin (2 million IV six times a day). Our patient stabilised and inotropic agents could be decreased within the next days. She was weaned off the ventilator after a tracheostomy was placed. The antibiotic treatment with benzylpenicillin was continued for a total duration of one week. The Netherlands Reference Laboratory for Bacterial Meningitis determined that the isolated N. meningitidis was serogroup W.

After initial good clinical improvement, our patient unfortunately developed gradual demarcation of the lower extremities due to sepsis and DIC. After an ICU period of 22 days, she was discharged to the general surgery ward for a bilateral lower leg and right thumb amputation. She rehabilitates with orthopaedic prostheses.

# **Table 1.** Characteristics of invasive meningococcalinfections at Erasmus University Medical Centrebetween January 2010 and October 2018

	Number of patients	Percent (%) of patients
Total number of patients	23	100%
Age of the patients (years)		
< 5	8	35%
5-14	2	9%
15-24	4	17%
25-44	3	13%
45-64	5	22%
> 65	I	4%
<b>Culture sample positive for</b> <i>N</i> . <i>meningitidis</i>		
Blood	22	96%
Cerebrospinal fluid	4	17%
Serogroup		
В	13	57%
W	7	30%
Y	3	13%
Outcome		
Dead	5	22%
Alive	18	78%

**Figure 1.** Number of patients with invasive meningococcal disease in the Erasmus University Medical Centre between January 2010 and October 2018 differentiated by serogroup



X-axis denotes year of invasive *N. meningitidis* infection; y-axis denotes number of meningococcal infections. \* The year 2018 consists of data up to October.

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#### DISCUSSION

We present a case of a young patient who developed septic shock and MODS due to an invasive infection with *Neisseria meningitidis*, for which the isolate was determined to belong to serogroup W (MenW). The initial presenting symptom of our patient was massive diarrhoea.

The incidence of invasive MenW infections is increasing worldwide, which is consistent with the increasing numbers in our academic hospital. The characteristics and subtypes of invasive meningococcal disease, defined as *N. meningitidis* isolated from blood and/or cerebrospinal fluid (CSF) at our medical centre are illustrated in table I and figure I. MenW infections are associated with a high case fatality rate,<sup>6</sup> which was also seen in our centre. In the last eight years, 23 patients were identified, of which five patients (22%) died as a result of the infection (serogroup W:2, serogroup B:I, serogroup Y:I).

The introduction of a vaccine against *N. meningitidis* serogroup C (MenC) in 2002 resulted in a rapid decline of the incidence of MenC in the Netherlands.<sup>10</sup> The growing number of MenW infections urges the need for vaccination against MenW. In 2018, the MenC vaccine, as part of the national vaccination program, will be replaced by the quadrivalent MenACWY-vaccine in the Netherlands.<sup>11</sup>

In summary, *Neisseria meningitidis* is known for causing sepsis and meningitis. Currently, the incidence of infections due to MenW is increasing in the Netherlands as it is worldwide. This serogroup can present atypically, as this case also illustrates, with massive diarrhea,<sup>5,6</sup> and highlights the difficulty of timely recognition of the disease which is pivotal for guidance of prompt treatment. Invasive infection with *Neisseria meningitidis* serogroup W should be included in the differential diagnosis of every patient presenting with a severe septic shock and gastrointestinal complaints.

#### DISCLOSURES

All authors declare no conflicts of interest. No funding or financial support was received.

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#### REFERENCES

- Bijlsma MW, Brouwer MC, Kasanmoentalib ES, et al. Community-acquired bacterial meningitis in adults in the Netherlands 2006-14: a prospective cohort study. Lancet Infect Dis. 2016; 23;388:343-55.
- Thigpen MC, Whitney CG, Messonnier NE, et al. Emerging Infections Programs Network. Bacterial meningitis in the United States, 1998-2007. N Engl J Med. 2011;364:2016-25.
- Stoof SP, Rodenburg GD, Knol ML, et al. Disease Burden of Invasive Meningococcal Disease in the Netherlands Between June 1999 and June 2011: A subjective Role for Serogroup and Cloncal Complex. Clin Infect Dis. 2015;61:1281-92.
- Rosenstein NE, Perkins BA, Stephens DS, Popovic T, Hughes JM. Meningococcal disease. New Engl J Med. 2001;344:1378-88.
- Wunderink HF, Vlasveld IN, Knol MJ, Van der Ende A, Van Essen HER, Kuijper EJ. Gastro-intestinale klachten bij meningokokkeninfectie. Ned Tijdschr Geneeskd. 2017;161:D1456.
- Campbell H, Parikh SR, Borrow R, Kaczmarski E, Ramsay ME, Ladhani SN. Presentation with gastrointestinal symptoms and high case fatality associated with group W meningococcal disease (MenW) in teenagers, England, July 2015 to January 2016. Euro Surveill. 2016;21:pii = 30175.
- Oldfield NJ, Cayrou C, AlJannat MAK, et al. Rise in Group W Meningococcal Carriage in University Students, United Kingdom. Emerg Infect Dis. 2017;23:1009-11.
- Martin NV, Ong KS, Howden BP, et al. Communicable Diseases Network Australia MenW Working Group. Rise in invasive serogroup W meningococcal disease in Australia 2013-2015. Commun Dis Intell Q Rep. 2016;24;40:E454-E459.
- Whittaker R, Dias JG, Ramliden M et al. The epidemiology of invasive meningococcal disease in EU/EEA countries, 2004-2014. Vaccine. 2017;35:2034-41.
- Bijlsma MW, Brouwer MC, Spanjaard L, Van de Beek D, Van der Ende A. A decade of herd protection after introduction of meningococcal serogroup C conjugate vaccination. Clin Infect Dis. 2014;59:1216-21.
- Knol MJ, Ruijs W, Antonise-Kamp L, De Melker HE, Van der Ende A. Implementation of Men ACWY vaccination because of ongoing increase in serogroup W invasive meningococcal disease, the Netherlands, 2018. Euro Surveill. 2018;23:pii=18-00158.

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