

doi: 10.14735/amcsnn2020570

Posterior reversible encephalopathy syndrome associated with *Neisseria elongata* meningitis

Syndrom reverzibilní encefalopatie v zadní cirkulaci ve spojitosti s meningitidou vyvolanou *Neisseria elongata*

Dear editor,

Posterior reversible encephalopathy syndrome (PRES) is a well-described clinical-radiological entity that refers to reversible subcortical vasogenic brain edema mostly affecting the posterior cerebral hemispheres [1]. The posterior predilection might be caused by relatively lower sympathetic vascular innervation of the region [2]. The association between infection and PRES is well recognised as the 3rd or the 4th common predisposing factor, but PRES related to meningitis is extremely rare [2,3]. We present a case of PRES associated with *Neisseria elongata* meningitis.

A previously healthy 35-year-old woman was admitted to the emergency department with headache, nausea, vomiting, fever, sleepiness and loss of consciousness. She had no previous complaints and the symptoms occurred within the last few days

but were gradually getting worse. She was referred to the intensive care unit upon recognition of fever (38.4 °C), somnolence and meningeal irritation signs such as neck stiffness. Laboratory tests showed leucocytosis (WBC 34,790/μL) and increased C-reactive protein (173 mg/L) and procalcitonin (1.39 ng/mL) levels. Lumbar puncture was purulent, cerebrospinal fluid (CSF) had the following composition: 377.94 mg/dL protein, 30 mg/dL glucose (concomitant blood glucose 181 mg/dL), and 1,800 WBC with abundant neutrophils. Cultures of CSF, blood, nasopharynx and urine were taken. Antibiotic therapy with meropenem 6g/day and vancomycin 2g/day empirically was started with the diagnosis of bacterial meningitis. Her consciousness improved quickly while her headache persisted and the synchronous neurological examination was

Redakční rada potvrzuje, že rukopis práce splnil ICMJE kritéria pro publikace zasílané do biomedicínských časopisů.

The Editorial Board declares that the manuscript met the ICMJE "uniform requirements" for biomedical papers.

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Accepted for review: 11. 5. 2020

Accepted for print: 8. 10. 2020

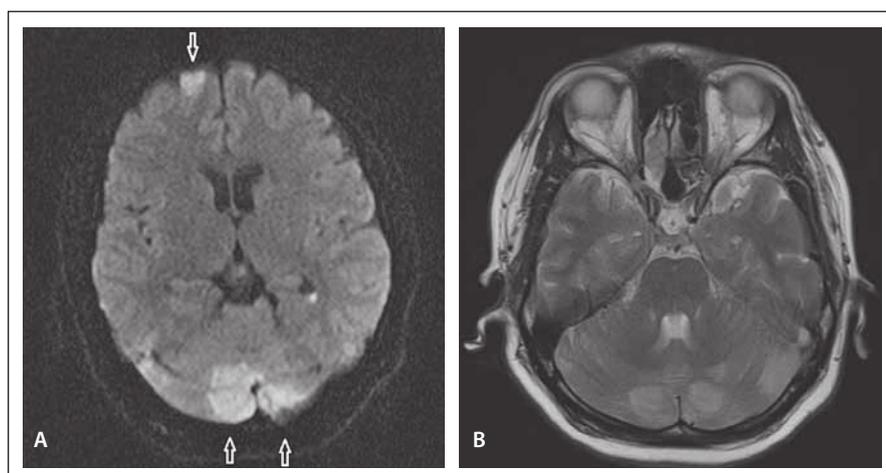


Fig. 1. (A) Diffusion weighted MRI – focal areas of restriction in the left superior frontal region (arrow) and in both cerebellar hemispheres (arrows). (B) MRI T2-WI sequence – hyper-intense lesions in both cerebellar hemispheres.

Obr. 1. (A) Difuzí vážená MR – ložiska restrikce nahoře frontálně vlevo (šipka) a v obou mozečkových hemisférách (šipky). (B) T2-WI sekvence na MR – hyperintenzní léze v obou mozečkových hemisférách.

normal except for bilateral papilledema and facial asymmetry. Diffusion weighted MRI showed focal areas of restriction in the left superior frontal, left posterior parietal; right temporo-occipital regions, both lateral ventricles (linear lesion) and both cerebellar hemispheres (Fig. 1A). Related hyperintense lesions on T2-WI MRI sequences were found especially in both cerebellar hemispheres (Fig. 1B). MR venography was normal.

After 3 days of culture from CSF, small (1 mm diameter) colonies of a non-motile, Gram negative coccobacillus grew that was facultative anaerobic, oxidase positive and

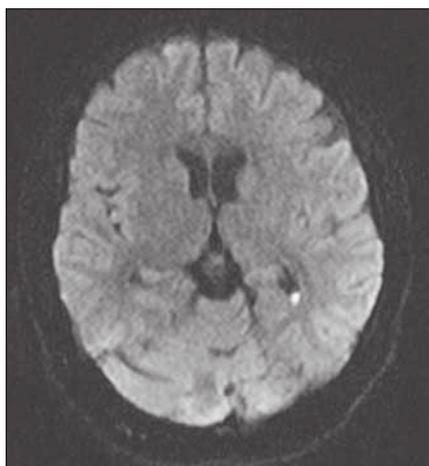


Fig. 2. Control diffusion weighted MRI after 10 days. Focal lesions related to edema were largely resolved.

Obr. 2. Kontrolní difúzí vážená MR po 10 dnech. Ložiskové léze v důsledku edému z velké části vymizely.

catalase negative and identified as *Neisseria elongata* by automated mass spectrometry microbial identification system (Vitek MS; bioMérieux, France). Unfortunately, further testing for determination of subspecies was not available.

In search for the Marcy l'Etoile, predisposing cardiac disease, trans-esophageal echocardiography was found to be normal. Visual examination and EEG were also non-pathologic. Kept on antibiotic treatment for bacterial meningitis and symptomatic treatment for headache, the patient recovered symptomatically and radiologically. After 10 days, the cerebral lesions were mostly resolved (Fig. 2) and completely disappeared at the 8th week on control MRI studies confirming the diagnosis of PRES associated with meningitis.

Various definitions of PRES share common basic features that are present in more than 90% of the cases, namely: specific neurological symptoms (headache, nausea, vomiting, convulsions), predisposing factors, specific radiologic pattern (focal areas in the parieto-occipital regions of hyperintense somewhat symmetrical signals on FLAIR [fluid attenuated inversion recovery] sequences),

related vasogenic brain edema, reversibility and exclusion of other patterns of encephalopathy and edema [1,2].

The exact pathophysiological mechanism leading to PRES remains conflicting. There are 2 main theories; the vasogenic theory implies the loss of autoregulation in cerebral blood flow due to sudden severe increase in blood pressure leading to cerebral hyperperfusion, vascular injury and damage to the blood-brain-barrier and resultant vasogenic edema. However, like our patient, more than 1/3 of the cases were found to be normotensive, thus an alternative mechanism based on endothelial dysfunction-injury or vasoconstriction causing hypoperfusion and ischemia was put forward [3]. The endothelial injury might be cytotoxic (immune suppressants-toxins), immunogenic (autoimmune diseases) or neuropeptide related (infection-sepsis) considering common risk factors such as arterial hypertension, renal diseases- transplantation-immune suppression, sepsis-infection, preeclampsia and autoimmune diseases [3,4]. Infection-sepsis was reported as an important predisposing factor in 20–25% of the cases [3,5].

The PRES was most probably related to cerebral infection in our patient who had clear initial clinical features like fever, meningeal irritation signs, leucocytosis, increased C-reactive protein and procalcitonin levels, purulent lumbar puncture with abundant neutrophils and so on. Thus, antibiotics were started and any additional vascular imaging was not performed. However, vascular imaging methods could be helpful to understand the blood flow conditions during the PRES as loss of cerebral small artery and precapillary sphincter autoregulation, leading to vasodilatation and finally to hyperperfusion and vasogenic edema, which might be the mechanism in normotensive patients.

N. meningitidis normally colonizes the nasopharynx asymptotically and the ability to invade the blood stream for the life-threatening meningitis, *N. meningitidis*' virulence is related to acquisition-possession of a polysaccharide capsule probably after genetic transport from other *Neisseria* species [6].

Capsule genes were identified in nonpathogenic *Neisseria* species, including *Neisseria subflava* and *Neisseria elongata* showing potential for human disease [6]. *Neisseria elongata* (family *Neisseriaceae*) is also a commensal of the oropharynx and subspecies *nitroreducens* is a rarely reported cause of septicemia and endocarditis usually in patients with cardiac valve disease or history of a dental procedure-infection [7,8]. PRES related to meningitis is extremely rare and to our knowledge this is the first case of *N. elongata* meningitis associated PRES [2,9]. CNS involved-complicated *N. elongata* infection is also a very rarely reported event [7]. As mentioned before, cardiac valve disease or dental procedure history is commonly present in *N. elongata* infections. Though our patient did not have any cardiac disease, common dental procedures might be a probability.

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