

Pros and Cons of Using BioFire FilmArray in the Diagnosis of Meningitis/Encephalitis

Marta Valentim

Department of Internal Medicine, Hospital Distrital de Santarém, Portugal

***Correspondence to:** Dr. Marta Valentim, Department of Internal Medicine, Hospital Distrital de Santarém, Portugal.

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Abbreviations (if used)

ME: Meningitis/Encephalitis

CSF: Cerebrospinal fluid

FDA: Food and Drug Administration

HIV: human immunodeficiency virus

Introduction

Meningitis/Encephalitis (ME) is a life-threatening entity, hence a rapid diagnosis and treatment is required to minimize the morbidity and mortality [1]. In United States there are approximately 4.100 cases of bacterial meningitis and 20.000 of encephalitis per year, with 12% and 7% of mortality associated, respectively [1].

Nowadays we have several diagnostic procedures for identify different microorganisms in the cerebrospinal fluid (CSF) responsible for ME: the gram stain (the current gold standard), the culture, the antigen detection and the different molecular methods [1]. All these methods can be useful to achieve the correct microbiological diagnostic, but there are two main problems common to all of them: first the lack of

sensitivity and specificity, and second, it takes time to get the results, in case the test achieve to detect the microorganism responsible for the ME [1].

The FilmArray ME Panel (BioFire, Salt Lake City, UT) were approval by Food and Drug Administration (FDA) in 2015 [2]. It has the capacity to detect multiple pathogens simultaneously. The results are obtained in one hour, using about 200 μ L of CSF sample obtained by lumbar puncture [3].

Panel Targets

The FilmArray ME Panel was designed to detect the most common pathogens that cause community-acquired or perinatally CNS infections [1], and also pathogens prevalent in immunocompromised patients [3] (Table 1).

Table 1: CSF FilmArray Panel Targets

Bacterial	Virus	Fungal
<i>Escherichia coli</i> K1	Herpes simplex virus 1	<i>Cryptococcus neoformans</i>
<i>Haemophilus influenzae</i>	Herpes simplex virus 2	<i>Cryptococcus gattii</i>
<i>Listeria monocytogenes</i>	Varicella-zoster virus	
<i>Neisseria meningitidis</i>	Cytomegalovirus	
<i>Streptococcus agalactiae</i>	Enterovirus	
<i>Streptococcus pneumoniae</i>	Human herpesvirus 6	
	Human parechovirus	

Pros

Analyzing the advantages of using this recent approved method, it is clear that the diagnostic speed is a huge improvement over other methods that we currently have, since it will allow us to identify the etiology and beginning an early and adequate treatment [3]. It is described in the literature that this method allows the detection of the microorganism responsible for EM, up to 22.9% in cases whose CSF cultures have been negative [2].

Another advantage associated with the use of FilmArray is the ease in the diagnostic identification of viral etiologies, which are generally difficult to obtain accurate and fast results by using other methods [2]. The widespread use of this technique will allow the suitability in the management, prescription and suspension, of the antibiotic therapy [3].

It is described that in the pediatric population it was possible to discontinue the antibiotic treatment during the first 24 hours, in 41.7% of the patients with positivity in the FilmArray for viral agents [3,4].

The relevance of the constant emergence of new diagnostic techniques is none other than the contribution, together with other tests and clinical manifestations, of adequate decision-making in specific cases by the clinical team [3,5].

Associated with a compatible clinic this panel has increased the detection of rare microbiological etiologies of ME [3]. Another positive point, is that it can help in the diagnosis of specific subpopulations such as pediatrics or patients infected with the human immunodeficiency virus (HIV) where the immune reconstitution inflammatory syndrome sometimes makes it difficult to differentiate it from CNS infections [6].

Finally, we should point out that the economic impact of the implementation of these techniques in the daily practice is barely described or studies performed. Soucek *et al.* perform a cost-effective study about the implementation of the FilmArray in a community hospital, in which they conclude that it would be justified the use from the second year, thinking about the cost reduction of the hospital stay, the use of ant biotherapy or the costs associated with morbidity/mortality [3,7].

Cons

The main points against the use of the FilmArray focus on that these techniques can never replace other diagnostic methods; because of the design it currently presents, it does not allow the detection of all the microorganisms responsible for ME [2].

Furthermore, despite the lack of literature in these areas, some studies as well as sporadic case reports agree with the fact, that the results obtained by FilmArray can delay and contribute to errors in the definitive diagnosis, due to false-positive [8] and false-negative [3] results. Taking into account how the technique is performed, it is related with a high risk of contamination of the sample which would increase the false-positive results. A clinical trial showed a high number of false positives secondary to contamination by *Streptococcus pneumoniae* [1,3].

The main problem with false-positive results are the delay to achieve the real diagnosis and the increase in the number of errors committed, as well as the deterioration in the patients prognosis [3,8].

Although only an article talks about the cost-effectiveness of the implementation of this procedure, the costs associated with the use are high [3], being necessary an infrastructure adapted to perform the technique with the greatest rigor possible, as well as trained staff and specific protocols for its implementation [3].

Conflicts of Interests

The authors declare that they have no competing interests.

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