TO THE EDITOR: In their review, “Cerebrospinal fluid secretion by the choroid plexus,” Damkier, Brown, and Praetorius (3) aimed to present recent advances in understanding the regulation of cerebrospinal fluid (CSF) secretion and limitations of current understanding of CSF formation and choroid plexus (CP) physiology. However, this review seems rather incomplete because an important part of CSF physiology, which disagrees with the classic view (2, 11, 13), had not been discussed. In this letter, we want to draw attention to only few of the many argumentations against the classic hypothesis. Due to the space limitation, the presented materials of CP here is very simplified.

In the light of classic hypothesis, CSF is presented as the so-called “third circulation” where CSF circulates as a slow river from the brain ventricles through entire CSF system to the cortical subarachnoid space to eventually be passively absorbed into superior sagittal sinus through arachnoid granulations. The CP represents a key organ in classic CSF physiology and is presented as a powerful biological pump responsible for CSF secretion (3, 14). Thereby, CP is considered to be responsible also for CSF pressure regulation (9) as well as for the development of hydrocephalus (as reviewed in Ref. 11).

CP IN CSF Secretion

According to the classic hypothesis, there is no doubt that CP is the main place of CSF formation and a biological pump by which CSF is exclusively secreted (14). By definition, CSF formation is production of CSF volume rate per unit of time. Since the volume of CSF represents the volume of water (because 99% of CSF is water), the CP should be the main place of water entry into CSF system. But both experimental (4, 5, 12) and clinical data (1, 10) have clearly shown that there is no significant contribution of CPs to water influx into the CSF system. Consequently, if the CP does not behave as a biological water pump (99% of CSF volume), it cannot be regarded as a CSF pump.

CP in Development of Hydrocephalus

According to the classic hypothesis, there is no doubt that hydrocephalus would develop in a situation where the CSF system is blocked between the site of secretion and the site of absorption. Due to continuous CSF secretion by CP (biological pump) and the inability of CSF absorption, hydrocephalus would occur in front of the blockade. But the facts are that, even with CSF pathways completely blocked, hydrocephalus would not develop every time (6, 11, 12). Furthermore, when CPs where removed in hydrocephalic patients, hydrocephalus still persisted, and shunt should have been reinstalled (11, 12). On the other hand, hydrocephalus might develop although blockade between the site of CSF secretion and absorption does not exist (8). Consequently, blockade of CSF circulation as well as CP functioning does not represent a key factor in hydrocephalus development.

All of these facts cannot be explained by the classic hypothesis but are easily explicable and fit very well in a new hypothesis of CSF physiology (2, 7, 11, 13) stating that CSF formation and absorption (CSF exchange) are constant and omnipresent processes in the CSF system, mainly as a consequence of water filtration between the capillaries and interstitial fluid of the central nervous system, and only partially in the CPs. There-
fore, when elaborating the CP functions, it would be necessary to discuss the new approach to the CSF physiology, even if authors disagree with this new hypothesis. It is the only way to elucidate the true role of CP in the nervous system.

**DISCLOSURES**

No conflicts of interest, financial or otherwise, are declared by the author(s).

**REFERENCES**