Postdural puncture headache (PDPH) is a major complication of neuraxial anesthesia that can occur following spinal anesthesia and with inadvertent dural puncture during epidural anesthesia. Obstetric patients are considered at increased risk for this condition because of their sex, young age, and the widespread use of neuraxial blocks. Inadvertent dural puncture during epidural anesthesia is a more common cause of PDPH than spinal anesthesia because of the use of small, pencil-point needles for spinal anesthesia in this population. During epidural placement, inadvertent puncture of the dura mater occurs at a rate of 1.5% (95% CI 1.5–1.5%), and more than half (52.1%; 95% CI: 51.4–52.8%) of these patients develop PDPH [1]. A more recent study showed that the incidence of PDPH after dural puncture with an epidural needle could be as high as 76–85% [2]. Although PDPH usually resolves spontaneously, it has the potential to cause significant morbidity in obstetric patients. It can also interfere with the mother’s ability to take care of herself or her baby, and may extend the length of hospital stay or evolve into chronic headache. In a retrospective case–control study, Webb et al. [3] reported that chronic headache occurred in 28% of obstetric women after inadvertent dural puncture with a 17-gauge Tuohy needle. Preventing PDPH should be the primary goal of clinicians dealing with this population. In fact, proper attention to procedure-related factors can significantly reduce its incidence. Occasionally, inadvertent dural puncture and PDPH are unavoidable complications. Therefore, anesthesiologists need to be familiar with treatment modalities and prevention. This review summarizes current concepts regarding the diagnosis, risk factors, pathophysiology, prevention, and treatment of PDPH.

Key Words: Epidural blood patch, Postdural puncture headache.

Introduction

Postdural puncture headache (PDPH) is a common complication after inadvertent dural puncture. Risks factors include female sex, young age, pregnancy, vaginal delivery, low body mass index, and being a non-smoker. Needle size, design, and the technique used also affect the risk. Because PDPH can be incapacitating, prompt diagnosis and treatment are mandatory. A diagnostic hallmark of PDPH is a postural headache that worsens with sitting or standing, and improves with lying down. Conservative therapies such as bed rest, hydration, and caffeine are commonly used as prophylaxis and treatment for this condition; however, no substantial evidence supports routine bed rest and aggressive hydration. An epidural blood patch is the most effective treatment option for patients with unsuccessful conservative management. Various other prophylactic and treatment interventions have been suggested. However, due to a lack of conclusive evidence supporting their use, the potential benefits of such interventions should be weighed carefully against the risks. This article reviews the current literature on the diagnosis, risk factors, pathophysiology, prevention, and treatment of PDPH.
Clinical Symptoms and Diagnosis

PDPH presents as a dull throbbing pain with a frontal-occipital distribution. Typically, the headache is aggravated by sitting or standing, and is reduced by lying down [4]. The diagnosis should be questioned in the absence of a postural component of the headache. At least partial relief should occur when the patient assumes the supine position. According to the International Classification of Headache Disorders criteria for the diagnosis of PDPH, headache develops within 5 days after dural puncture and disappears spontaneously within 1 week, or up to 48 h after an epidural blood patch. The headache may be accompanied by neck stiffness, tinnitus, hypoacusia, photophobia, and nausea [5]. However, recent studies indicate that PDPH occurs within 3 days after dural puncture [6-8], and up to 29% of patients have headache as the only symptom [6]. Rarely, the headache may last for months or even years [9]. Postpartum headaches are very common, developing in 39% of women, and tension and migraine headaches are more common in pregnant than non-pregnant women [10]. Therefore, when diagnosing PDPH in obstetric patients, it is important to exclude other causes of headache, such as functional headaches. Less frequent complications of accidental dural puncture include reversible encephalopathy [11-13], pneumocephalus [14], and subdural hematoma [15,16]. Therefore, when neurological signs or changes in headache characteristics such as non-postural headache occur, serious etiologies should be excluded such as subdural hematoma, cerebral thrombosis, and reversible encephalopathy [17].

Pathophysiology

In 1899, August Bier pioneered work on spinal anesthesia and first described PDPH, suggesting that it might be caused by cerebral spinal fluid (CSF) loss. Although the precise mechanism of this condition remains unknown, the postulated cause of the headache is reduced CSF pressure due to loss of CSF in the epidural space through the dural puncture site [18]. Decreased CSF pressure creates a loss of the cushioning effect normally provided by intracranial fluid. The resulting traction placed on intracranial pain-sensitive structures elicits pain [18]. A second possible cause is distension of the cerebral blood vessels. With a sudden drop in CSF pressure, vasodilation of the intracranial vessels occurs to maintain a constant intracranial volume, resulting in a pathophysiology similar to vascular headache [19]. The beneficial effects of vasoconstrictor drugs such as caffeine and theophylline in PDPH support this mechanism.

Risk Factors

Patient factors

The well-known risk factors of PDPH include young age, female sex, and pregnancy. Young adults are at higher risk of developing this condition than older individuals (14% vs. 7%) [20] because with increasing age, the dura may be less elastic and less likely to gape. Women, particularly during pregnancy, are considered at increased risk for PDPH [6,21]. Its high incidence may be attributed to increased estrogen levels, which influence the tone of the cerebral vessels, thereby increasing the vascular distension in response to CSF hypotension [6,21,22]. Another risk factor for PDPH is vaginal delivery. Pushing efforts during the second stage can increase the size of the dural hole and CSF loss. However, two retrospective studies examining whether second-stage pushing increased the incidence and severity of PDPH or the need for an epidural blood patch reported conflicting results [23,24].

Some evidence suggests that morbidly obese patients have a decreased incidence of PDPH [25-27]. The increase in epidural pressure observed in obese compared to thin patients may lessen the pressure gradient from the intrathecal space to the epidural space, decreasing the loss of CSF. However, again, retrospective studies have revealed conflicting results [28]. In a retrospective review of the records of 125 parturient women with accidental dural puncture or PDPH, Miu et al. [28] found no evidence that women with a high body mass index (BMI) are less likely to develop PDPH or require an epidural blood patch. In contrast, in a retrospective review of 518 parturient women who experienced accidental dural puncture, Peralta et al. [29] reported that the incidence of PDPH in parturients with a BMI ≥ 31.5 kg/m² (39%) was lower than that in those with a BMI < 31.5 kg/m² (56%; difference = −17%, 95% CI −7% to −26%, P = 0.0004). However, the BMI did not affect the severity of PDPH or the need for an epidural blood patch.

Needle design, size, and direction

The type and size of needle are also important factors in PDPH, given that research clearly demonstrates that larger dural tears result in a higher incidence of this condition. Cutting nee-
dles (Quincke needles) are associated with a higher incidence of PDPH compared to blunt or pencil-point needles (Sprotte and Whitacre needles). Schmittner et al. [31] and Gisore et al. [32] confirmed the significantly lower incidence of PDPH with pencil-point needles compared to Quincke cutting needles in similar studies (1.7% vs. 6.6%, P = 0.02 and 4.5% vs. 24.2%, P = 0.042). A modification of the Quincke (Atracan) needle is also available, with a cutting point and a double bevel to cut a small dural hole and then dilate it. Several studies have confirmed that the bigger the needle, the greater the incidence of PDPH [19,33-35]. With Quincke needles, the incidence and severity of PDPH is directly related to the size of the needle. A similar effect may occur with pencil-point needles.

Another needle-related factor that results in a reduced incidence of PDPH is the orientation of the needle bevel parallel to the long axis of the spine, resulting in decreased disruption of dural fibers [36]. This longitudinal bevel orientation separates the dural fibers rather than cuts them, which facilitates closure of the dural hole on withdrawing the needle. A longitudinal orientation of the needle bevel has been confirmed to reduce the risk of PDPH compared to perpendicular insertion [37,38].

Regarding the direction of the bevel of a Tuohy needle, Norris et al. [37] reported a decreased incidence of PDPH when the bevel entered the epidural space along the long axis of the spine and then was rotated 90 degrees before inserting the catheter compared to keeping a perpendicular approach.

**Prophylactic Interventions**

**Conservative**

When unintentional dural puncture occurs, several conservative therapies are commonly used, such as hydration and bed rest. These therapies aim to decrease CSF loss through the dural hole and restore CSF with additional fluid intake [39]. These methods are simple and do not have any serious adverse effects, but no conclusive evidence supports their use for the prevention of PDPH [40,41]. Particularly during the puerperium, early ambulation is recommended due to the risk of deep vein thrombosis and pulmonary embolism because of hypercoagulability. The authors of a 2016 Cochrane review [40] found no evidence suggesting that routine bed rest after a dural puncture is beneficial for preventing PDPH (risk ratio [RR]: 1.16, 95% CI: 1.02–1.32). Furthermore, bed rest likely increased PDPH (RR: 1.24, 95% CI: 1.04–1.48) compared to early ambulation. There was also no evidence to support a benefit of prophylactic fluid supplementation (RR: 1, 95% CI: 0.59–1.69). Several prophylactic drugs have been studied, but their clinical effectiveness has not been established. For example, epidural morphine [42] and intravenous cosyntropin [43] have shown promising effects in the prevention of PDPH after dural puncture. However, neither is recommended routinely due to side effects and limited evidence.

**Invasive**

**Prophylactic epidural blood patch**

A prophylactic epidural blood patch can be performed through the epidural cathether, which is re-sited after inadvertent dural puncture, just before the epidural catheter is removed. Autologous blood injected into the epidural space is thought to seal the dural defect. Therefore, the use of a prophylactic epidural blood patch might prevent PDPH and the need for further treatments in that regard. However, recent systemic reviews with a meta-analysis of four randomized controlled studies have not conclusively supported the use of a prophylactic epidural blood patch for preventing PDPH [44,45]. Three of the four randomized controlled studies analyzed in these systemic reviews found a statistical difference in PDPH incidence. However, the study using a sham procedure to exclude the placebo effect did not show any difference. In a randomized controlled study published in 2014, Stein et al. [46] found a significant reduction in PDPH in 11 of 60 patients who received a prophylactic epidural blood patch compared to the 39 of 49 patients who did not (P < 0.0001). This study was flawed because no sham epidural blood patch was used. Although a prophylactic epidural blood patch was effective for some studies, methodological limitations were identified in these studies such as problems with randomization, blindness, and small sample size. A prophylactic epidural blood patch should be performed after full recovery of sensation to prevent the inhibition of coagulation by local anesthetics [47] or accidental total spinal anesthesia [48]. In addition, the occurrence of pain can be a signal to stop the injection of blood. In this context, the procedure is usually performed at least 5 h after the last dose of epidural anesthetic. Risks associated with prophylactic epidural blood patches include common transient complications such as backache and rare complications such as a neurological deficit or infection.

**Intrathecal catheter placement**

The placement of an intrathecal catheter through the dural puncture hole for up to 24 h has been proposed as a preventive measure for PDPH. Theoretically, this intervention prevents CSF leakage by sealing the dural hole, and maintaining the catheter for up to 24 h results in a localized inflammatory response that encourages hole closure [49]. This procedure allows immediate analgesia during labor through the intrathecal catheter, negating the risk of repeated dural punctures [50–52]. Systematic reviews [44,45,53] have indicated that intrathecal catheter placement
does not significantly reduce the incidence of PDPH. However, this intervention does reduce the overall severity of PDPH and the need for an epidural blood patch. Similarly, in a retrospective study of 218 inadvertent dural puncture cases, Bolden and Gebre [54] compared the incidence of PDPH and the need for an epidural blood patch between a re-sited epidural group and a spinal catheter group. There were no differences in incidence between the groups (68.0 vs. 55.9%, odds ratio [OR]: 1.7, 95% CI: 1.0–2.9, P = 0.07). The need for an epidural blood patch was significantly reduced in the spinal catheter group compared to the re-sited epidural group (52.0% to 20.3%, OR: 4.2, 95% CI: 2.4–7.6, P < 0.001). The addition of intrathecal saline to the intrathecal catheter reduced the need further, from 55.2% to 25.0% (OR: 3.7, 95% CI: 1.2–11.2, P = 0.02). Intrathecal catheterization has potential risks such as meningitis or abscess [55], arachnoiditis, and cauda equina syndrome [56,57].

**Epidural saline administration**

Saline injected into the epidural space may decrease CSF loss by reducing the pressure gradient between the epidural and subarachnoid spaces. Therefore, epidural saline has been used to prevent PDPH with variable success. However, a systematic review and meta-analysis of epidural saline failed to demonstrate the prophylactic effects of epidural saline on the incidence of PDPH or the need for an epidural blood patch [44]. A prophylactic epidural blood patch, intrathecal catheter placement, and epidural saline injection are not considered routine preventative therapies due to lack of substantial evidence. A careful decision regarding the use of these prophylactic interventions is needed because of this lack of evidence and the potential risks.

**Invasive**

Patients who do not respond to conservative treatment within 48 h require more aggressive interventions. An epidural blood patch is considered the treatment of choice for moderate and severe PDPH, with success rates of 61–98% [71-75]. Based on the success rates, an epidural blood patch seems to be less effective for the obstetric population. Stride and Cooper [23] reported 64% complete relief of PDPH after the first blood patch. This lower success rate may be related to the large-gauge needle puncture of the dura during epidural analgesia.

The postulated mechanism of action of the epidural blood patch involves sealing the dural puncture site. Magnetic resonance imaging studies have found that epidural blood adheres to the thecal sac, resulting in clot formation for up to 18–24 h [76]. The pain relief is often rapid after application of a blood patch, although the CSF volume is not restored immediately. Therefore, the sudden relief cannot be explained by the puncture site closure. Carrie [77] hypothesized that the epidural injection of blood increases the lumbar CSF pressure, subsequently restor-
ing the intracranial CSF pressure and resulting in reflex cerebral vasoconstriction.

Observational studies report that an epidural blood patch within 24–48 h of dural puncture is less effective [72,78,79]. Early onset PDPH is likely to be more severe; therefore, selection bias is possible. Alternatively, if the therapeutic benefit of an epidural blood patch is due to clot formation at the dural puncture site, factors inhibiting clot formation result in unfavorable effects. First, the initial high CSF flow may displace the clot. Second, if the epidural blood patch is performed too early, clot degradation may occur and prevent hole closure [80].

Although the optimal volume of blood required for a successful epidural blood patch is not known, volumes ranging from 5 to 30 ml have been reported [44,72]. Paech et al. [72] randomized 121 patients to receive 15, 20, or 30 ml autologous blood during an epidural blood patch. The efficacy of the epidural blood patch was similar in all of the three groups, with a success rate of 70%. However, the authors found that 19% and 46% of those randomized to the 20 ml and 30 ml volumes, respectively, failed to receive the assigned volume because of injection pain. Because the epidural space varies in capacity and compliance, patients experienced pain during injection before a volume of 30 ml was administered. Similarly, a recent retrospective review by Booth et al. [81] reported that increasing the blood volume up to 30 ml did not reduce the need for a repeated epidural blood patch, when the final volume was primarily determined by back pain during injection.

The epidural injection of colloids is a useful alternative to blood if an epidural blood patch is contraindicated. Dextran-40 [82,83] or hydroxyethyl starch [84] have been described with various success rates. Colloids are thought to result in increased epidural pressure and decreased CSF leakage. Complications include transient discomfort and a burning sensation. Although evidence is limited, colloid solutions can be suggested in patients who refuse an epidural blood patch or when an epidural blood patch is ineffective.

The greater occipital nerve, which is derived from the dorsal root of the second cervical nerve, is the main sensory nerve in the occipital region. Greater occipital nerve blocks have been used for the treatment of different types of headache. A study of greater occipital nerve block for the treatment of PDPH showed beneficial effects in reducing pain severity, although the evidence is limited. Several authors have suggested its use as an alternative to an epidural blood patch since it is less invasive and leads to prompt symptom relief [85,86].

**Conclusions**

Since PDPH can occur after puncture of the dura matter, the most effective method for decreasing its incidence is to develop techniques that minimize dural hole formation during spinal block and that prevent inadvertent dural puncture during epidural block. With better awareness of the risk factors related with PDPH, it is important for clinicians to be vigilant when performing spinal or epidural blocks. In reality, inadvertent puncture of the dura mater still occurs and PDPH continues to be a problem in these patients. Although various prophylactic interventions after inadvertent dural puncture have been suggested based on risk factors and pathophysiologic concerns, there is still insufficient evidence of their benefits. For the treatment of PDPH, an epidural blood patch is most effective treatment modality, with a high rate of success. Several other treatment modalities for PDPH are available, but high-level evidence supporting their efficacy is still needed. Overall, further investigation including large qualified trials is warranted, and careful attention should be paid to these issues until substantial evidence is available.

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Postdural puncture headache

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