Predicting spinal hypotension during Caesarean section

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Abstract:
Hypotension under spinal anaesthesia for Caesarean section remains a common problem with attendant maternal and foetal morbidity attached to it. This review examines some of the issues surrounding the prediction of spinal hypotension, including concerns with current evidence, debate regarding the mechanism of hypotension and the utility of prediction in this group of patients. It will then cover some of the more conventional and established preoperative predictors of hypotension. Particular attention will be paid to the assessment of autonomic function and some of the novel methods being used as predictors of severe maternal hypotension. The implications of autonomic dysfunction and areas for future research are discussed.

Keywords: Spinal anaesthesia, hypotension, Caesarean section, prediction.

Introduction
Hypotension under spinal anaesthesia for Caesarean section is a common and important problem, with significant maternal and foetal implications. The ability to predict which patients are at particular risk for severe hypotension would enable anaesthetists not only to prepare appropriately but potentially also to individualise treatment. In the setting where junior doctors are asked to anaesthetise patients for Caesarean section, the early identification of patients at risk might also allow for timely referral to regional centres where appropriate expertise is available. This review will deal predominantly with the issues surrounding the prediction of spinal hypotension in the patient for Caesarean section, with a particular focus on those methods which assess the autonomic nervous system.

Why is this an important question?
Spinal hypotension during Caesarean section is common and up to 80% of anaesthetics require the use of vasopressor to treat it.1 The exact incidence of hypotension varies considerably with the definition, as will be seen later.2 Under these circumstances, one could argue that the prediction of hypotension might be accomplished simply through the decision to administer a spinal anaesthetic. In addition, the treatment of hypotension is so effective, that some editorials have even questioned the necessity of hypotension, and there is little consensus on what this definition should be. Klohr and colleagues demonstrated that in studies involving hypotension during Caesarean section between 1999 and 2009, there were 15 different definitions across 63 studies.2 No definition was used in more than a quarter of papers, with the most common definition being a reduction in baseline blood pressure of more than 80%. In applying these definitions to a cohort of their own patients, they found that the incidence of hypotension could vary between 7.4% and 74.1% simply by altering the criteria for inclusion.

Problems with the evidence
The incidence of hypotension during Caesarean section under spinal anaesthesia is significantly influenced by the definition of hypotension, and there is little consensus on what this definition should be. Klohr and colleagues demonstrated that in studies involving hypotension during Caesarean section between 1999 and 2009, there were 15 different definitions across 63 studies.2 No definition was used in more than a quarter of papers, with the most common definition being a reduction in baseline blood pressure of more than 80%. In applying these definitions to a cohort of their own patients, they found that the incidence of hypotension could vary between 7.4% and 74.1% simply by altering the criteria for inclusion.

Another problem with much of the evidence is the small numbers of patients in the various cohorts. While many of the studies demonstrate a statistically significant difference in either an intervention or for a predictive factor, the law of large numbers suggests that these studies may be operating within areas of maximum variability. The results of these studies are thus susceptible to increased fragility. It is further apparent that patients who would seem most likely to suffer a catastrophic hypotensive episode under spinal anaesthesia are unlikely to receive this form of anaesthetic. Patients with extreme tachycardia, overt sepsis or cardiac conditions that preclude significant elevation of the cardiac output are examples of patients that would not usually receive a spinal anaesthetic. One must therefore be mindful of the population to which the
majority of these studies apply: the otherwise healthy patient for elective or emergency Caesarean section under spinal anaesthesia.

**What is the mechanism of spinal hypotension?**

In appreciating which factors are likely to predict spinal hypotension, it would seem important that an understanding of the mechanism of hypotension under spinal anaesthesia is necessary. There is a significant body of recent work which has looked at the haemodynamic changes which occur under spinal anaesthesia and the effects of various treatment regimens on these changes.

It has long been held that the dominant mechanism of hypotension in the patient for Caesarean section is caval compression. However treatments based on the caval compression theory have not proved effective and the principles behind this theory have been challenged in a fairly recent editorial. This editorial suggests that a reduction in arterial sympathetic tone is more likely to be the dominant mechanism of spinal hypotension, and this has subsequently been supported by studies which used continuous cardiac output monitors during spinal anaesthesia. The typical response to spinal anaesthesia is therefore hypotension due to decreased systemic vascular resistance and a resultant increased response to spinal anaesthesia is hypotension due to caval compression. However treatments based on the caval compression theory have not proved effective and the principles behind this theory have been challenged in a fairly recent editorial. This editorial suggests that a reduction in arterial sympathetic tone is more likely to be the dominant mechanism of spinal hypotension, and this has subsequently been supported by studies which used continuous cardiac output monitors during spinal anaesthesia. The typical response to spinal anaesthesia is therefore hypotension due to decreased systemic vascular resistance and a resultant increased heart rate, although a small proportion of patients may respond with hypotension and bradycardia. It has also been noted that pre-eclamptic patients are relatively resistant to the effects of spinal anaesthesia which several authors propose is due to circulating vasoconstrictors offsetting the effects of the loss of arterial vascular tone. This group of patients may thus comprise a group to whom more conventional risk factors for hypotension do not apply.

**What are the proven predictors of spinal hypotension?**

**Caval compression**

While caval compression may not be the dominant or most common cause of spinal hypotension during Caesarean delivery, it seems that a subset of susceptible patients are at particular risk. Studies using the supine stress test to identify patients at risk have shown that a positive stress test is predictive of hypotension under anaesthesia. This test involves the measurement of heart rate and blood pressure in both the supine and the lateral position, and observing the changes that occur. Interestingly, while the supine hypotensive syndrome is thought to exist in only approximately 8% of patients, the supine stress test was positive in a far greater proportion in these studies, ranging from 36% to 52%.

**Factors related to the technique of spinal anaesthesia**

There are a number of factors which are related to the way in which the anaesthetic is given rather than specific to individual patients. One study showed that a slower speed of injection resulted in a lower incidence of hypotension and in addition, the hypotension was of delayed onset, shorter duration and required less ephedrine. A recent review has also confirmed the more obvious assumption that a lower dose of spinal anaesthetic reduces the incidence of hypotension, as demonstrated in a number of prospective trials.

**Maternal Body Mass Index**

Some studies have suggested that patients with an increased body mass index (BMI) are at increased risk for the development of hypotension under spinal anaesthesia for Caesarean delivery. Few studies have been designed to specifically examine this question, and the BMI cut-offs in these studies varied from 25 to 29. Other studies which use multi-variate analysis to detect risk factors for hypotension have also confirmed a raised BMI as a risk factor. Interestingly, one study showed that an inadequate weight gain (less than 11kg compared to antenatal weight) was a risk factor for altered heart rate variability patterns and hypotension under spinal anaesthesia. Patients with an abnormal antenatal BMI were excluded from this study.

**Maternal heart rate**

The use of baseline heart rate as a predictor for the development of hypotension under spinal anaesthesia has somewhat surprisingly shown conflicting results. One study showed a significantly increased incidence of hypotension in the group with a higher baseline heart rate. However, a study with a very similar design did not show baseline heart rate as a good predictor of spinal hypotension. It should be remembered however, that these studies specifically study healthy patients and women with higher baseline heart rates may well have been excluded from these studies.

One of the mechanisms thought to be responsible for catastrophic hypotension under spinal anaesthesia is the Bezold-Jarisch reflex, a cardiac depressor reflex involving bradycardia, vasodilatation and hypotension. A review on the subject highlighted that the term has come to refer to perioperative bradycardia associated with hypotension. The risk factors for the development of this reflex have been studied in the general population and it appears that patients who are healthy and of younger age are at increased risk, along with those who have a higher level of spinal blockade. However, there seems to be little evidence which can reliably predict the onset of this reflex in the obstetric population for spinal anaesthesia.

**Other predictors**

One of the factors that repeatedly comes up in multi-variate analysis is advanced age. This has been shown to be a predictor of spinal hypotension during Caesarean section in a few studies which assessed multiple risk factors, with one study suggesting that an age greater than 35 being the cut-off. It has also been shown that increased preoperative positional blood pressure change correlates with an increased incidence of spinal hypotension and higher ephedrine requirements. The authors postulated that an increased level of sympathetic activity was the mechanism behind this phenomenon. A higher level of preoperative anxiety has been shown to be predictive of an increased incidence of hypotension under spinal anaesthesia. The authors also postulated that this was due to an increased level of sympathetic activity in the preoperative phase.

There have been some novel methods developed recently which show promise as immediate preoperative predictors...
of spinal hypotension. One of these is the use of a perfusion index derived from a pulse oximeter, which effectively assesses peripheral perfusion dynamics and vascular tone, and which has been shown to predict spinal hypotension.44 More studies will be required to determine the place of this monitor in routine clinical practice.

Assessment of autonomic function

Given that hypotension is almost inevitable with spinal anaesthesia, it would potentially be more useful to screen for patients with an inability to compensate for this hypotension. While it seems that the sympathetic induction by spinal anaesthesia is the dominant cause of hypotension, it is also apparent that the compensation required in order to maintain blood pressure is dependent on a functioning autonomic nervous system (ANS). Assessment of the ANS could therefore provide clinicians with useful information regarding the ability of patients to cope with haemodynamic instability. However, the assessment of autonomic function is complex and varies from bedside tests described over thirty years ago35 to the more advanced techniques used today. One of these methods is the assessment of heart rate variability (HRV) in patients preoperatively, which is then subjected to statistical analysis. A variety of different statistical tests may be performed, including time domain, frequency domain and non-linear analysis. A task force has attempted to provide guidelines for this use of this technology, its related equipment and the statistics employed, although it is now somewhat outdated.36

Chamchad and colleagues were the first to demonstrate that the use of heart rate variability techniques could be used to predict spinal hypotension in patients undergoing elective Caesarean section.37 They used a variable called the point correlation dimension to assess patients as either low or high risk for the development of hypotension. Point correlation dimension accurately predicted which women were likely to get hypotension and performed better than conventional markers such as baseline heart rate. Hanss’ group then used frequency domain analysis of the HRV to predict the onset of spinal hypotension during Caesarean delivery, using a retrospective model to identify the at-risk group and then a prospective group to confirm this.38 They used a variable called the LF/HF ratio, which is thought to reflect sympathetic versus parasympathetic balance.39 This is supported by work in the obstetric patient which shows decreasing LF/HF values in patients undergoing spinal, and therefore a presumed lower sympathetic outflow.40 They demonstrated that patients with a higher balance, and thus higher sympathetic tone, were particularly susceptible to spinal hypotension.41 Importantly, assessments of heart rate variability were predictive of hypotension if performed on the same day of surgery, but not if performed the day before. Hanss’ group then used these variables to successfully guide prophylactic treatment in the patients that were at risk for the development of severe hypotension.42 There has also been further work outside of the obstetric setting which showed that altered heart rate variability predicted hypotension under spinal anaesthesia, both with the use of the LF/HF ratio, and with measurements of entropy.42

While HRV shows promise as a tool for evaluating perioperative risk, there are also several issues which have been raised.43 Among them are concerns about the effects of heart rate on variability, the types of analysis techniques utilized and the lack of understanding around the mechanisms of altered heart rate variability.44 In addition, the techniques for assessing heart rate variability lack standardization and are not yet available as monitor which is easily utilized by the anaesthetist. Currently it remains predominantly a research tool, although both the software and equipment could easily be incorporated into a bedside monitor.

The implications of autonomic dysfunction

There are thus several studies which have shown that alterations in autonomic function lead to an increased risk of hypotension under spinal anaesthesia. It should be remembered that autonomic neuropathy in anaesthesia is associated with a host of other potential complications and perioperative problems.44 Studies in the non-obstetric population have shown that patients with autonomic dysfunction may be at an increased risk of cardiac adverse events,45 cardiovascular instability46 and hypothermia.47 Admittedly these studies applied to patients with specific pathology and potentially complex disease processes. It should be appreciated however, that abnormalities in the autonomic nervous system have at least the potential to affect a multitude of other systems, including the cardiac, gastrointestinal and renal systems. Issues like altered temperature regulation and an impaired stress response are highly relevant in the perioperative period and would certainly apply to Caesarean sections. Studies which address the risk of complications linked to autonomic dysfunction in patients with altered heart rate variability would be useful in testing this hypothesis.

Conclusion

There are thus a number of proven predictors for the development of hypotension during spinal anaesthesia in the obstetric population, but much of the evidence is hampered by lack of consensus regarding definitions and by small study sizes. While a degree of hypotension is almost universal, it is specifically the patient with severe hypotension or catastrophic collapse that would derive the most benefit from advance warning. In addition, patients with an inability to compensate secondary to autonomic dysfunction could be at risk for a number of other complications which have yet to be clearly elucidated. Further work in this area should concentrate on trying to identify these patients in particular.

Declaration

The author declares that he has no financial or personal relationship(s) which may have inappropriately influenced him in writing this paper.

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