

REVIEW

Chronic pain during pregnancy: a review of the literature

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Methods: A literature search was undertaken using the search engines PubMed, CINAHL, EBSCOhost, and Web of Science. Search terms used included "chronic pain" AND "pregnant OR pregnancy" OR "pregnancy complications" from inception through August 2016.

Results: The basis of this review was the 144 articles that met inclusion criteria for this review. Based on our review of the current literature, we recommend 7 guidelines for chronic pain management during and after pregnancy: 1) complete history and physical examination; 2) monitor patients for alcohol, nicotine, and substance use; 3) collaborate with patient to set treatment goals; 4) develop a management plan; 5) for opioids, use lowest effective dose; 6) formulate a pain management plan for labor and delivery; and 7) discuss reproductive health with women with chronic pain.

Conclusion: The management of chronic pain associated with pregnancy is understudied. Obstetrical providers primarily manage chronic pain during pregnancy. Some general guidelines are provided for those health care providers until more information is available.

Keywords: chronic pain, pregnancy, pregnancy complications, chronic pain in pregnancy, guidelines for pain in pregnancy, pain management in pregnancy

Introduction

Chronic pain impacts over 100 million Americans and is the most common cause of long-lasting disability. Conspicuously, women are more impacted by chronic pain disorders than men. Health care providers are frequently faced with the difficulties of managing chronic pain conditions that were present prior to, during, or after pregnancy. Pain thresholds in healthy women without chronic pain have been shown to increase throughout pregnancy peaking just prior to delivery. Animal models have shown that naltrexone, an opioid antagonist, can block these findings. It is unclear how these findings may translate to women with chronic pain or opioid exposure as such individuals often have hyperanalgesia, allodynia, and/or decreased pain thresholds at baseline.

Back pain is a frequent ailment encountered in pregnancy, and a frequently used standard textbook for obstetrics training, Williams' Obstetrics, provides few guidelines for the management of acute back pain during pregnancy.⁴ The textbook cites "evidence-based clinical research directing care is limited" and provides even less education concerning the diagnosis and/or management of chronic back pain.⁴ Health care providers caring for obstetrical patients have inadequate training and limited

Correspondence: Everett F Magann Department of Obstetrics and Gynecology, University of Arkansas for Medical Sciences, 4301 W Markham Street, Slot # 518, Little Rock, AR 72205, USA Tel +1 501 686 8345 Fax +1 501 526 7820 Email efmagann@uams.edu time to manage patients with chronic pain. Pain management teams and clinics often discharge or decline to manage women during pregnancy, and transition of care is often referred to the obstetrical team. The United States Opioid Epidemic has recently underscored the risk of prescription drug abuse for patients as well as the role that health care providers have in monitoring and educating patients for substance and alcohol use disorders. The lack of evidence-based treatment guidelines and limited formal training and expertise in the management of chronic pain places pregnant women at risk to receive suboptimal care.

The majority of published reviews have focused on the pharmacological or non-pharmacological approaches, rather than on a broader scope of how to approach chronic pain during pregnancy. The purpose of this review is to identify evidence-based clinical research for the evaluation and management of preexisting chronic pain during pregnancy, chronic pain that has been associated with pregnancy, and chronic pain in relation to mode of delivery (ie, vaginal or cesarean) as well as provide guidance to address the broader clinical care of this population. The information obtained will provide the basis for clinical guidelines for the management of chronic pain disorders by obstetrical care providers during the perinatal period. Our goal is to help guide clinicians in management of these patients; thus, an in-depth discussion of the perinatal effects of in utero pharmacological exposure is beyond the scope of the review.

Methods

A literature search was undertaken using the search engines PubMed, CINAHL, EBSCOhost, and Web of Science. Search terms used included "chronic pain/therapy" AND "pregnant OR pregnancy" OR "pregnancy complications" from inception through April 7, 2016. Abstracts were limited to those in English. There were 200 abstracts identified and reviewed by two independent reviewers. One independent reviewer then examined the full research article corresponding to chronic pain/therapy and pregnancy for relevance. References for such articles were screened for additional applicable research.

Of the 200 abstracts identified, 15 were duplicates; 3 were not available in English; 2 could not be located; and 47 were chosen for full manuscript review. After screening the reference list of each identified article, an additional 115 articles were identified, of which, 18 articles could not be located. The basis of this review includes 144 articles (Figure 1), including 47 cohort studies, 35 reviews, 28 case reports, 13 randomized clinical trials, 10 case series, 9 case control studies, 1 comparative analysis, and 1 qualitative interview.

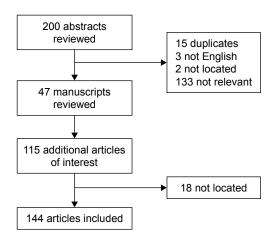


Figure I Preexisting chronic pain disorder during pregnancy.

Preexisting chronic pain disorder during pregnancy

The overall prevalence of pregnant women with preexisting chronic pain disorders is unknown. Likewise, the general course of chronic pain conditions during pregnancy has not been studied. Preexisting chronic pain disorders have a negative impact on pregnancy and are associated with increased sick leave use, insomnia, and depressive symptoms during pregnancy.^{5,6}

Numerous reviews of both the prescription and nonprescription pharmacological treatment of chronic pain have been published.^{7–13} As previously mentioned, the purpose of this review is not to discuss treatment options in-depth, but we feel it is important to discuss opioids and acetaminophen as both options have received recent scrutiny. Opioid prescriptions in the first trimester increased by 33.9% from 1995 to 2009 for non-cancer pain diagnoses indicating that pain syndromes during pregnancy are being increasingly treated with prescription opioids.¹⁴ This finding is not surprising as opioids are often second-line treatment options during pregnancy, after acetaminophen, due to the established efficacy in acute conditions and limited reproductive safety data of other pharmacological options. 15 Coinciding with the increase in prescription opioid use is an increasing rise in rates of infant with neonatal abstinence syndrome (NAS), a syndrome related to opioid exposure in utero that is characterized by difficulties with feeding and sleeping as well as central nervous system dysfunction. 16 A single source, The Canadian Guideline for the Use of Opioids in the Treatment of Chronic Noncancer Pain, has published treatment guidelines that recommend the following: 1) opioids should be tapered and discontinued or prescribed at the lowest effective dose; 2) in the postpartum period, codeine use should be avoided or limited to 4 days; 3) women should

be managed by perinatologists; and 4) women with opioid use disorders should be referred for appropriate treatment.¹⁷ The formulation of these guidelines is influenced by the potential addictive and teratogenic effects of opioids (ie, cardiovascular birth defects, neural tube defects, hydrocephaly, glaucoma, NAS, gastroschisis, and fetal death)18-21 as well as the correlation of codeine with planned cesarean delivery, postpartum hemorrhage,²² and infant death in a breastfeeding mother who is a CYP2D6 ultra-rapid metabolizer.23 No opioid is given preference in these guidelines owing to lack of data supporting individual medications. Methadone has the largest amount of reproductive safety data secondary to its use in medication-assisted treatment programs for pregnant women with opioid use disorders. Methadone is also an effective treatment for chronic pain, and a few studies have compared obstetrical and neonatal outcomes between methadone for chronic pain disorders versus opioid use disorder during pregnancy. Pregnant women receiving methadone for chronic pain had better infant growth but more neonatal morbidity and preterm birth rates compared to women with an opioid use disorder.²⁴ In two studies comparing rates of NAS with prescription opioid use and not illicit opioid use, NAS occurred in 5.6%²⁵ and 11%²⁴ of exposed neonates, rates lower than infants born to mothers in opioid treatment programs on maintenance medications (ie, methadone or buprenorphine). No evidence is available for the treatment of chronic pain during pregnancy using buprenorphine. In conclusion, reproductive safety data of opioids are limited and often conflicted. Jones et al discussed the methodological limitations of current research and the need for future research on the use of opioids in pregnancy for chronic pain disorders.9

Acetaminophen is the most common nonprescription medication recommended during pregnancy to treat pain and has been recently associated with increased risk for respiratory, testicular, attention-deficit/hyperactive-like behavioral, and autism spectrum disorders.^{24–30} These results have questioned the reproductive safety profile of acetaminophen; however, it remains one of the most commonly used medications during pregnancy, specifically for pain indications.

A variety of non-pharmacological treatments for chronic pain also exists, but data during the perinatal period are limited. Evidence supporting the use of spinal cord stimulation for chronic regional pain syndrome and other chronic pain states is primarily based on case reports and case series that demonstrate conflicting obstetrical and neonatal effects.^{31–41} Three case reports of intrathecal baclofen pumps used in pain management during pregnancy without obstetrical or neonatal complications have been reported.^{42–44}

Conservative guidelines do not recommend the use of transcutaneous electrical nerve stimulation (TENS) during pregnancy due to potential legal consequences with an adverse pregnancy outcome, but liberal guidelines suggest that the use of TENS is acceptable but should be avoided over the maternal abdomen or pelvic areas.⁴⁵ Acupuncture is also often suggested, but the literature search found no studies supporting its use.

We identified a number of articles discussing specific pain conditions during pregnancy, such as migraines, rheumatoid arthritis, sickle cell disease, Ehlers-Danlos syndrome, and vulvodynia. Each is discussed in more detail later in the paper. Additionally, our literature search found a case report of intrathecal opioids for a pregnant woman with tuberous sclerosis⁴⁶ as well as reviews of irritable bowel syndrome during pregnancy,⁴⁷ complex regional pain syndrome during pregnancy,⁴⁸ and the development of acute pain on existing chronic pain during pregnancy.⁴⁹

Headaches

Headaches and more specifically migraines are present in 16%–18% of women of reproductive age, but the prevalence during pregnancy is unknown. Most experts agree that the course of headaches (tension-type, migraines, or a combination) during pregnancy is variable. Studies have shown that migraines largely improve or remit during pregnancy with an initial occurrence during pregnancy in 1.3% of women and in the postpartum period in 4.5% of women. Pregnant women with headaches have been reported to have a greater rate of preterm delivery, gestational hypertension, and preeclampsia. St.56

Multiple review articles have been written summarizing the management of headaches, including migraines, during pregnancy, and women who are breastfeeding;^{57–59} Jürgens et al reviewed the treatment of cluster headaches during the perinatal period.60 Management includes both acute and preventative techniques. The acute management of headaches and migraines during pregnancy primarily utilizes non-pharmacological measures initially, such as relaxation techniques, appropriate duration of sleep, stress management, massages, application of cold, avoidance of triggers, and biofeedback. A combination of biofeedback, relaxation training, and physical therapy has also been successfully used in pregnancy.⁶¹ Pharmacological options, in order of level of recommendation, for acute management include acetaminophen, triptans, antiemetics, and opiates. In a large prospective, observational study, triptan (ie, sumatriptan, rizatriptan, zolmitriptan, eletriptan, naratriptan, almotriptan) use in the 6 months prior to pregnancy was associated with an increased risk of stillbirth.62 The same study found that triptan use during pregnancy was not associated with an increased rate of major congenital malformations, while use in the second and third trimester was associated with an increased risk of atonic uterus and blood loss >500 mL during labor. 62 Maternal prescription of a sumatriptan during pregnancy was not associated with increased teratogenic risk⁶³ but was associated with an increased risk of preterm delivery. 64 An examination of children with in utero exposure to triptans (ie, sumatriptan, rizatriptan, zolmitriptan, eletriptan, naratriptan, almotriptan, frovatriptan) at 3 years of age showed a positive association with externalizing behaviors but no association with altered or delayed psychomotor function, communication, or temperament development. 51,65 Metoclopramide with or without acetaminophen can also be used to treat both acute migraines as well as accompanying nausea without adverse perinatal effects. 66 Evidence supporting the use of other antiemetics for acute treatment of migraine was not found.

For migraine sufferers, preventative treatment, in order of recommendation, during the antenatal period includes tricyclic antidepressants, topiramate, magnesium, and botulinum toxin. Evidence supporting the efficacy of any option is limited during pregnancy, and treatment recommendations are based on reproductive safety data. Botulinum toxin is approved by the FDA to treat chronic migraines and has been studied in the treatment of various chronic pain disorders. ⁶⁷ The use of botulinum toxin has been discussed with recommendations that state pregnant women should make an informed decision about the use of botulinum toxin during the perinatal period after reviewing the potential risks and benefits to the pregnancy and fetus. ⁶⁷

Rheumatoid arthritis

The prevalence of rheumatoid arthritis during pregnancy is unknown. The pregnancy course of rheumatoid arthritis varies, but the majority of patients with RA improve during the antenatal period with a subsequent relapse in the post-partum period. Preconception counseling for individuals with rheumatoid arthritis is recommended given the teratogenic potential of treatment options. However, the overall impact of rheumatoid arthritis on obstetrical and neonatal outcomes is unknown. Three reviews have summarized the pharmacological management of rheumatoid arthritis during the perinatal period with a review of commonly used medications that are recommended for use during the perinatal period (ie, chloroquine, hydroxychloroquine, sulfasalazine, azathioprine, cyclosporine, corticosteroids, NSAIDS), 70-72 and glucocorticoids are the most commonly used. 33

Sickle cell disease

Although the prevalence of sickle cell disease during pregnancy is unknown, exacerbations can occur due to increased oxygen consumption and decreased functional residual capacity of the lungs during gestation, predisposing patients to sickling, vaso-occlusive pain, and sickle cell crises. 74 Early reports of pregnant women with sickle cell disease demonstrated an association with spontaneous abortions, stillbirths, and pregnancy-related maternal death;75,76 however, medical advancements have contributed to improved obstetrical and neonatal outcomes.^{77,78} Treatment during pregnancy is primarily supportive with use of medications on a case-by-case basis. Opioids have been a mainstay in the treatment of sickle cell pain crises, while hydroxyurea should be avoided due to its possible teratogenicity. 74,79 Prophylactic red blood cell transfusions have no obstetrical or neonatal advantage when compared to an as-needed use for transfusions, except for a decrease in the number of painful crises.80 Adequate hydration and avoidance of triggers are helpful in the prevention of sickle cell crises and other sequelae of sickle cell disease.

Ehlers-Danlos syndrome

Ehlers-Danlos syndrome is a heritable connective tissue disorder affecting collagen and leading to hypermobility of joints and skin and chronic debilitating musculoskeletal disorders. The prevalence of pregnant women with the syndrome is unknown. The condition can be exacerbated in women who become pregnant secondary to the additional joint laxity and increased maternal weight of pregnancy. In addition, labor can progress rapidly.⁸¹ Impact on neonatal outcomes is unknown. Management is primarily with acetaminophen, opioids, and short courses of steroids.^{7,81}

Vulvodynia

The prevalence of vulvodynia during the perinatal period is unknown. The course is also largely unknown but a few investigations, detailed later, have been completed. A qualitative study found that women with vulvodynia minimized their pain severity during preconception timing and often discontinued medications prior to conception with optimism about the possibility of improved pain during pregnancy. While pregnant, all women reported increased anxiety about pain; however, no clear consensus of pain course was found during pregnancy and in the postpartum period. In a retrospective case—cohort study, vulvodynia did not impact likelihood of pregnancy or pregnancy outcomes, but women with vulvodynia were more likely to have a cesarean section and increased postpartum pain with a vaginal delivery that resulted in a laceration or an episiotomy.

Chronic pain associated with pregnancy

Clinical practice dictates that low back pain is a common occurrence during pregnancy that begins between the fifth and seventh months, and research illustrates that low back pain and pelvic pain occur in 71.7% of pregnancies. Back pain during pregnancy is considered to be lumbar pain, pelvic pain, or a combination of both with a few studies investigating differences between the types of pain. Hip and foot pain are also common during the perinatal period but not routinely investigated. Before the types of pain are also common during the perinatal period but not routinely investigated.

Low back pain during pregnancy is associated with age;⁸⁷ Caucasian and African-American race;^{87,88} and the presence of low back pain before pregnancy, during menstruation, and during a previous pregnancy.⁸⁸ On the other hand, Hispanic ethnicity;⁸⁸ use of oral contraceptives, caffeine, tobacco, and exercise;⁸⁷ prepregnancy body weight;⁸⁷ parity;⁸⁷ and history of infertility with hormone therapy are not associated with an increased risk.⁸⁷ Low back pain causes moderate disability and about 10% of patients will take off time from work.^{87,89} The occurrence of low back pain during pregnancy increases the risk of low back pain later in life and in subsequent pregnancies.⁹⁰

The course of low back pain during the antenatal period has not been documented; however, recovery in the post-partum period occurs in 58.3% of first, 66.4% of second, and 70.7% of third pregnancies. In a prospective study investigating lumbar pain, pelvic girdle pain, and combined pain during pregnancy, results showed that 1 out of 5 women with pelvic girdle pain or combined pain had persistent pain 3 months postpartum in comparison to 1 out of 29 with lumbar pain. Pelvic pain persists at a rate of about 8.5% at 2 years postpartum, and low back pain persists at a rate of about 21.1% at 2 and 3 years postpartum.

Chronicity of antenatal back pain that continues into the postpartum period is associated with a previous history of back pain while non-pregnant or pregnant, 93 chronic pelvic girdle pain, or a combined pelvic girdle/low back pain that includes muscle weakness in the large pelvic and dorsal muscles, 94 work dissatisfaction, 91 early onset of symptoms in the pregnancy, 91,93 older age, 91,93 younger age, 89 increased sick day use during pregnancy, 89 higher parity, 89 higher body mass index, 89 higher weight gain during the perinatal period, 93 an inability to return to prepregnancy weight postpartum, 93 and lower endurance of the back flexors. 91 Physical activity level prior to pregnancy does not impact one's risk of developing chronic low back pain and pelvic pain in the postpartum period. 85

Herniated lumbar discs are one isolated cause of back pain during pregnancy, occurring at an incidence rate of 1 in 10,000 that should be recognized.⁹⁵ Warning signs include predominant radicular leg pain, motor signs of limb paresis, or abnormalities in reflex response associated with limiting sciatic pain radiation during a straight leg-raising maneuver.⁹⁵ An operative delivery is recommended.⁹⁵

Treatment of back pain during pregnancy includes both pharmacological and non-pharmacological approaches. Pharmacological options include low-dose aspirin, acetaminophen, and short use of NSAIDS.96 Non-pharmacological treatment options include patient education with or without physiotherapy, 97 mechanical supports, TENS; exercise, craniosacral therapy, osteomanipulative therapy, multimodal intervention, and acupuncture. 98 Data are conflicting regarding efficacy of the aforementioned options, and 75% of providers will not recommend any formal treatment.87,88 A 2013 Cochrane Review found moderate-quality evidence supporting acupuncture and exercise for pelvic or lumbopelvic pain and low-quality evidence supporting exercise for low back pain.⁹⁹ Meanwhile, a 2015 Cochrane Review by the same authors found low-quality evidence supporting the role of exercise in reducing low back pain and moderate-to-low quality evidence that exercise improves functional disability. 100 The latter review also found that acupuncture, craniosacral therapy, osteomanipulative therapy, and multmodal intervention may be of benefit to pregnant women with pelvic pain. 100 A systematic review for physical therapy found no strong evidence in support of therapeutic effects; 101 however, the same authors found that physical therapy is an effective treatment for pelvic girdle pain and adding specific stabilizing exercises provides a potential advantage. 102 Acupuncture is also effective when compared to physiotherapy, and both acupuncture and physiotherapy are superior to education only. 103 Pelvic support belts provided no advantage over education and exercise. 104 Likewise, there is no benefit to pelvic girdle pain with an education group, home exercise group, or supervised exercise group when participants are provided a non-elastic sacroiliac belt. 105 Group education by itself has also been found to be ineffective. 106 Case studies support the use of instrumentassisted soft tissue mobilization107 as well as acupuncture combined with opioids. 108

Chronic pain associated with mode of delivery (vaginal vs cesarean)

While some studies have shown that the risk of chronic pain after pregnancy is rare at 0.3%, ¹⁰⁹ other studies have shown a more common prevalence rate of 10%. ¹¹⁰ Studies have shown that delivery mode is not related to the risk of experiencing pain at 8 weeks, ¹¹¹ 3 months, ¹¹² and 2 years. ¹¹³ On the

other hand, mode of delivery may impact the prevalence as rates are 0.6%–22% following caesarean delivery and 1%–10% with vaginal delivery. ^{114–123} In the loan study investigating deliveries with an episiotomy, up to 12.8% of patients had chronic pain lasting a minimum of 5 months. ¹²⁴

Lavand'homme has published extensively about chronic pain after delivery and noted five key points: 1) incidence of persistent postsurgical pain after childbirth is lower than similar surgical procedures; 2) pain characteristics are poorly detailed; 3) after a cesarean delivery, pain is primarily low-intensity neuropathic pain; 4) the extent of trauma is not predictive of pain severity; and 5) the aforementioned findings suggest that protective antinociceptive mechanisms occur during the perinatal period. 115,125,126 Others have echoed the prevalence of neuropathic pain after cesarean delivery. 127

Notable risk factors for postpartum chronic pain include a history of previous chronic pain disorder and higher postoperative pain severity^{110,111,119,121,128} but not epidural analgesia.¹²⁹ Use of general anesthesia, non-private insurance status, emergency cesarean delivery, and psychological indication for cesarean delivery are risk factors noted with caesarean delivery only.^{109,117,119,130,131} In the lone genetic study, polymorphisms of the ATP-binding cassette sub-family B member-1 gene were not associated with the increased risk for development of chronic pain following a cesarean section, but a trend was found for women with the T allele of tC3435T polymorphism.¹³²

The relationship between analgesic methods (ie, intrathecal clonidine, wound infiltration, transversus abdominal plane block, intravenous ketamine, oral gabapentin, and pregabalin) and development of chronic pain following an operative abdominal delivery in the postpartum period have been discussed extensively and the following surgical techniques have been implicated as risk factors: larger surgical incisions with increased risks for lateral nerve entrapment and neuromas, uterine exteriorization and parietal peritoneal closure, 121 and repeated surgical procedures as a cause for hypersensitization. 133 Two articles compared patients based on surgical technique, Pfannenstiel or Misgav-Ladach, 134,135 and results favored the Misgav-Ladach technique. 134,135 Multiple Pfannenstiel incisions have been shown to predict chronic pain at 12 months.¹³¹ Postoperative treatment of pain also impacts the development of postpartum chronic pain as higher doses of local anesthetic and the use of anti-inflammatory medicines postoperatively decreased the incidence of chronic pain at 3 months. 130 To decrease post-cesarean pain, a double-blind placebo-controlled trial compared intravenous magnesium sulfate in low and high

doses with a control group and observed that short-term pain and long-term pain (6 weeks) were similar between the magnesium arms and a control group of women. ¹³⁶ The impact of postpartum chronic pain on neonatal outcomes or women's quality of life is largely unstudied, but one investigation found a negative impact on sexual and nonsexual activities. ¹¹⁰

Our literature search failed to find any discussion on the preferred treatment options for postpartum chronic pain with exception of the neuromas. Neuromas of the iliohypogastric and ilioinguinal nerves are often the source for chronic pain, and treatment options include nerve blocks, neurectomy, or decompression of involved peripheral nerves. ^{137,138} Although oxytocin has not been studied as a potential treatment for pain, animal studies have implicated a role of oxytocin in the alleviation of pain in the immediate postpartum period. ¹³⁹

A rare source of postpartum chronic pain is pubic symphysis diastasis. After a vaginal delivery, diagnosis is suspected with the following: pain immediately postpartum, pain exacerbation with weight bearing, symphyseal tenderness, and an interpubic gap greater than 10 mm. 140 The exact cause and prevalence are unknown. 141,142 It is thought that relaxin and progesterone levels may be associated with development of pubic symphysis diastasis; however, relaxin level in pregnancy does not correlate with either degree of symphyseal distention or pain level. 143 Evidence-based treatment is derived from a number of case reports, 142,144-146 and patients are typically managed conservatively with or without analgesia. Small studies have shown success with physical therapy or chiropractic management, 148 surgical stabilization, 144,148 and spinal cord stimulation when other treatments have failed. 149 A small retrospective chart review of 14 patients showed seven with no benefit of corticosteroid injections. 150

Summary

Chronic pain disorders are a major problem during the perinatal period, and health care professionals have received limited formal training. Previous data have largely focused on pharmacological management during pregnancy, primarily with the use of opiates. Our literature search found that the majority of information focuses on the management of preexisting chronic pain conditions during pregnancy. A moderate amount of literature describes the risk of developing a chronic pain disorder in the postpartum period due to mode of delivery; however, data detailing chronic pain conditions that develop during pregnancy are scant. Research detailing the impact of pain on obstetrical and neonatal outcomes

is largely absent. Of highest concern to many obstetrical providers is "how to" manage pregnant women with chronic pain disorders, and our literature search confirms the need to more clearly define the role(s) of the clinical care team and expand evidence-based data to guide management.

Two reasons are plausible for the limited information on pain disorders during the perinatal period. First, pregnancy in women with chronic pain is rare. This is disputed by the fact that chronic pain is the most common form of chronic illness for people under the age of 60 years, and women are disproportionally impacted. With the increasing rates of women older than 30 years giving birth, clinicians are likely to see women with complex medical problems, such as chronic pain syndromes, experiencing pregnancy. Clinicians may also discount pain complaints during the antenatal period due to the clinical belief that low back pain is a norm of pregnancy. This belief supports the next point. Second, chronic pain syndromes and their treatment during pregnancy are not studied. Past reviews as well as our current review have demonstrated previous investigations. However, data detailing efficacy of treatments, specifically non-pharmacological options, is scant. Third, the available data are difficult to capture. A limitation of our review was the inadequacy of the search terms to capture fully the articles of interest. This displays the difficulty in capturing the limited published information as well as the hazard in getting evidence-based information for busy clinicians.

We feel there is sufficient evidence to support the development of broad guidelines to address the clinical management of chronic pain disorders during the perinatal period by obstetrical care providers. Of note, pregnancy is often an exclusion criterion for many chronic pain programs, and obstetricians are deemed the "de facto" expert in all conditions during pregnancy. As such, a referral to a maternal fetal medicine physician or chronic pain provider is recommended if available. The 2016 Center for Disease Control Guideline for Prescribing Opioids for Chronic Pain¹⁵¹ and the Canadian Guideline for the Use of Opioids in the Treatment of Chronic Noncancer Pain¹⁷ provide a good starting place, but are focused on the use of opioids. Our guidelines were developed to address clinical management by obstetrical care providers of these patients as well as the use of opioids. We intend our guidelines to be a starting point for discussion and hope future research efforts will better define parameters as well as support optimal obstetrical and neonatal outcomes. Guidelines are presented in the following text alongside a level of evidence supporting them (Table 1).

Table I Level of evidence

Level	Evidence
I	Evidence from a systematic review or meta-analysis of all
	relevant RCTs (randomized controlled trials) or evidence-
	based clinical practice guidelines based on systematic
	reviews of RCTs or three or more RCTs of good quality
	that have similar results.
II	Evidence obtained from at least one well-designed RCT
	(eg, large multi-site RCT).
III	Evidence obtained from well-designed controlled trials
	without randomization.
IV	Evidence from well-designed case-control or cohort studies.
V	Evidence from systematic reviews of descriptive and
	qualitative studies (meta-synthesis).
VI	Evidence from a single descriptive or qualitative study.
VII	Evidence from the opinion of authorities and/or reports of
	expert committees.

Abbreviation: RCT, randomized controlled trial.

Guideline #1:At the initial obstetrical appointment, an appropriate history and physical examination should be completed (Level VII)

Helpful information includes detailed information about the cause of chronic pain; pre-conception treatment history with response; current treatment plan and provider; history of chronic pain during previous pregnancies; and an assessment of comorbid conditions that may complicate treatment, such as depression. Obtaining medical records from previous providers can be helpful to obtain complete and accurate health information. This information will help guide the clinician in the management plan. If collaborating with other providers (ie, chronic pain management teams), it is pertinent to delineate responsibilities (eg, who will prescribe pain medications).

Guideline #2:All patients should be monitored for alcohol, nicotine, and illicit drug use disorders continuously throughout the perinatal period (Level VII)

The presence of an alcohol, nicotine, or substance use disorder complicates the management of chronic pain as these individuals are at potential risk for overdoses and abuse of prescription medications. Monitoring includes screening patients; obtaining a urine drug screen at the initial appointment and periodically thereafter; and checking the state's prescription drug monitoring program, if available, at each appointment. Screening can be completed via forms or conversation. Available screening forms include the 4Ps and Car, Relax, Alone, Forget, Friends, Trouble screen (CRAFFT, for women aged 26 years or younger) questionnaires. There is no clear consensus which screening form is optimal for pregnancy. Urine drug screens provide an objective

and secondary assessment for illicit drug use and should be obtained with consent from the patient. Identification of substance use during pregnancy has potential legal complications, and thus some providers may avoid obtaining urine drug screens. We feel that urine drug screens are a vital aspect of chronic pain management as they help identify exposures that pose a teratogenic risk to fetus as well as maternal risk of overdosing and substance use. Checking the state's prescription drug monitoring program helps verify the patient's history at the initial visit, provides a continuous evaluation of appropriate medication use, and aids in the assessment of a substance use disorder. Any patient with a confirmed or suspected substance use disorder should be referred for an evaluation and/or treatment with an addiction expert. We also advise patients to be cognizant of their inherent concerns for abuse and addiction, including how these concerns may negatively impact their treatment decisions.

Guideline #3: Collaborate with the patient to establish a treatment goal (Level VII)

Being free of pain is often not an achievable treatment goal without significant safety hazards during the pregnancy. This will often require providers to be comfortable with a patient's discomfort. We recommend establishing an objective goal, such as an appropriate pain level or allowing a patient to work full time.

Guideline #4: A management plan should be formulated that minimizes in utero exposures while maximizing obstetrical and fetal health (Level IV)

Treatments include both non-pharmacological and pharmacological treatments. Non-pharmacological options include acupuncture, chiropractic care, TENS units, spinal cord stimulation, exercise, and transcranial magnetic stimulation. Given the limited amount of information detailing safety and efficacy, we caution the use of non-pharmacological treatment options. Providers should document that the patient has received previous benefit as well as a discussion of the unknown risks of the treatment to both mom and fetus. Pharmacological management should focus on minimizing exposures to the fetus. This can occur by simplification of medication regimens and elimination of medications that pose limited efficacy and/or excessive risk to mom and/or the unborn child (eg, acetaminophen). For example, a patient who is receiving treatment with oxycodone-acetaminophen and morphine prn would be treated with oxycodone monotherapy at effective doses. Some medications may need to be tapered to prevent withdrawal and/or rebound pain. Opioid prescriptions should be tapered to discontinuation if possible as eliminating and/or preventing NAS is a laudable goal. We recognize that not all patients can be managed without opioids and recommend to prescribe them safely (see guidelines in the following text). Antiepileptic drugs are often used to manage chronic pain conditions and require additional folic acid supplementation (eg, 4 mg PO daily) during pregnancy to decrease the risk of neural tube defects. Folic acid supplementation should also be considered with any in utero exposure to antiepileptic drugs during the antepartum period.

Guideline #5: For opioid prescriptions, clinicians should prescribe at the lowest effective dose and quantity (Level IV)

We recommend responsible prescribing of opioid medications, and encourage pill counts at each appointment to verify compliance with the prescribed regimen. While the authors agree that excessive use of opioids should be avoided due to overall risk of high doses of opioid, such as death or iatrogenic overdose, opioids should be utilized to effect if they cannot be eliminated from use. In addition, no data demonstrate improved obstetrical or neonatal outcomes with lower doses compared to high doses.

Guideline #6: A pain management plan for labor and delivery should be formulated in collaboration with the patient (Level V)

Patients with chronic pain disorders often express fear and anxiety of labor and delivery due to the anticipation of pain; thus, these patients benefit from reassurance and recognition of their pain tolerance, hyperanalgesia, and increased analgesic requirements as well as a discussion of pain management at delivery. For patients requiring specialized services from anesthesiology, they may benefit from a consultation prior to delivery. Following a vaginal delivery, patients will benefit from scheduled acetaminophen and ibuprofen. Some patients may require opioids following a vaginal delivery, and almost all will require opioids following an operative delivery. When providing opioids following delivery, patients should receive 3–5 days of higher doses of opioid medications with strict dosage instructions. In the acute postpartum period, a plan for transitioning the patient back to the prior chronic pain management team should be formulated.

Guideline #7: Clinicians should discuss reproductive health with any woman with a chronic pain condition and provide counseling as appropriate (Level VII)

As half of all pregnancies in the United States are unplanned, this discussion is appropriate at any time during the perinatal period as well as during annual evaluations of non-pregnant patients. Topics include appropriate contraceptive methods

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as well as a risk-benefit discussion of medications during the perinatal period. For women using medications with potential teratogenicity, a daily prenatal vitamin (with additional folic acid if an antiepileptic medication is prescribed) may also be helpful. In general, women of reproductive age managed on antiepileptic drugs should receive folic acid supplementation (eg, minimum of 1 mg PO daily).

Conclusion

The course and management of chronic pain disorders during the perinatal period is understudied, and obstetrical care providers are typically responsible for pain management. In recognition of the need for more data, we have provided general guidelines for the clinical leadership of these patients during the perinatal period. Cohort studies investigating the treatment of various chronic pain disorders provide the highest level of supporting evidence (ie, Level IV). The lack of well-designed randomized clinical trials as well as systematic reviews and meta-analyses prohibits the practice of evidence-based medicine. Future research should address this need by focusing on the prevalence, course, and optimal management of chronic pain disorders during the antenatal and postpartum period.

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Author contributions

All authors contributed toward data analysis, drafting and critically revising the paper, gave final approval of the version to be published, and agree to be accountable for all aspects of the work.

Disclosure

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