

SPECIAL ARTICLES

Chronic Opioid Therapy and Sleep: An American Academy of Sleep Medicine Position Statement

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There is a complex relationship among opioids, sleep and daytime function. Patients and medical providers should be aware that chronic opioid therapy can alter sleep architecture and sleep quality as well as contribute to daytime sleepiness. It is also important for medical providers to be cognizant of other adverse effects of chronic opioid use including the impact on respiratory function during sleep. Opioids are associated with several types of sleep-disordered breathing, including sleep-related hypoventilation, central sleep apnea (CSA), and obstructive sleep apnea (OSA). Appropriate screening, diagnostic testing, and treatment of opioid-associated sleep-disordered breathing can improve patients' health and quality of life. Collaboration among medical providers is encouraged to provide high quality, patient-centered care for people who are treated with chronic opioid therapy.

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INTRODUCTION

The American Academy of Sleep Medicine (AASM) is the leading professional society dedicated to the promotion of sleep health. The AASM improves sleep health and fosters high-quality, patient-centered care through advocacy, education, strategic research, and practice standards. The AASM endeavors to advance sleep health policy that improves the health and well-being of the general public.

While fatigue and disturbed sleep are common symptoms of chronic pain, there is also a complex relationship among opioids, sleep and daytime function. Chronic use of opioid therapy is known to disrupt sleep architecture and sleep stage distribution. These opioid-related sleep disturbances in turn can cause or contribute to disturbed sleep, fatigue and daytime sleepiness.

Another adverse effect of opioid use is respiratory depression.² It is important for medical providers to be aware of the risk of sleep-disordered breathing in chronic opioid therapy patients.^{3–6} Types of sleep-disordered breathing that can be associated with chronic opioid use include both isolated symptoms (eg, snoring), as well as sleep-related breathing disorders characterized by abnormalities of respiration during sleep, including sleep-related hypoventilation, central sleep apnea (CSA), and obstructive sleep apnea (OSA).^{7,8} Effective testing is available for sleep-disordered breathing associated with opioid therapy, and treatment can improve patients' health and well-being.

While opioid therapy can contribute to sleep disruption and sleep-disordered breathing, there is one sleep disorder for which narcotics may be prescribed as a therapy. Current guidelines state that opioid therapy can be an effective treatment option for patients with severe, refractory restless legs syndrome (RLS) in whom other therapies have failed to control symptoms.⁹

POSITION

It is the position of the AASM that:

- Medical providers need to be aware that chronic opioid therapy can alter sleep architecture and cause respiratory depression, increasing the risk for sleepdisordered breathing.
- Appropriate screening and diagnostic testing can identify sleep-related hypoventilation, CSA, and OSA in people treated with chronic opioid therapy.
- The treatment of opioid-associated sleep-disordered breathing can improve patients' health and well-being.
- In patients with chronic pain, collaboration among primary care providers, pain medicine specialists, and sleep specialists is encouraged to provide high quality, patient-centered care for people who are treated with chronic opioid therapy.

 Medical providers need to be aware that the cautious and judicious use of low-dose opioid therapy is a treatment option for severe, refractory RLS.

DISCUSSION

The use of opioid medications has soared since 1990 and has received increasing attention given its overall impact on our nation's social welfare and public health. ^{10,11} Nearly 92 million Americans use prescription opioids, 11.5 million misuse them, and 1.9 million have opioid use disorder. ¹² Since the late 1990s an epidemic of drug overdose deaths has developed in the U.S. In 2017 alone, approximately 68% of the more than 70,200 overdose deaths involved either a prescription opioid, an illegal opioid such as heroin, or illicitly manufactured fentanyl. ¹³

Although nonpharmacologic therapy and nonopioid pharmacologic therapy are preferred treatment options for chronic pain, current clinical guidelines from the American Academy of Pain Medicine, American Pain Society, and the Centers for Disease Control and Prevention acknowledge that long-term prescription opioids remain a viable treatment option for some patients with chronic, noncancer pain when the expected benefits are anticipated to outweigh the potential harms. ^{14,15} These clinical guidelines define chronic opioid therapy as the use of opioids on most days for at least three months.

Regular, chronic use of opioids is known to interfere with sleep via several mechanisms. It can lead to changes in sleep architecture including a reduction in sleep efficiency as well as slow wave sleep. ¹⁶ There also may be a reduction in rapid eye movement (REM) sleep. ¹⁷ These effects of chronic opioid ingestion on sleep architecture appear to be dose dependent and lead to poor sleep quality, resulting in daytime sleepiness. ^{1,16–19}

In addition to causing changes in sleep architecture, multiple studies have demonstrated that chronic use of opioids is associated with an increase in sleep-disordered breathing.^{20,21} The use of chronic opioids can result in isolated symptoms (eg, snoring) and/or more severe abnormalities of respiration during sleep, including sleep-related hypoventilation, CSA and OSA.^{7,8} Sleep-related hypoventilation involves chronic hypoventilation and hypercapnia due to impaired respiratory drive and alterations in carbon dioxide (CO₂) and oxygen (O₂) chemosensitivity. While both OSA and CSA have been noted, studies have generally demonstrated a higher prevalence of CSA with chronic opioid use.^{20–22} In CSA, the repetitive cessation of breathing is associated with absent or reduced respiratory effort due to diminished central respiratory drive, which may be further impaired in the setting of chronic opioid use.²³ When CSA is caused by an opioid or other respiratory depressant, ataxic breathing can occur.²⁴ OSA is characterized by repetitive episodes of complete or partial upper airway obstruction, with subsequent reductions in blood oxygen saturation, and may be terminated by brief arousals from sleep.²³ While untreated CSA and OSA can be detrimental to a patient's health and well-being, effective treatments are available, including different modalities of positive airway pressure (PAP) therapy, such as continuous PAP (CPAP), bilevel PAP (BPAP), and adaptive servo-ventilation (ASV).

Polysomnography (PSG) is one type of sleep test that is indicated for the diagnosis of sleep-related breathing disorders. Although a home sleep apnea test (HSAT) may be used for the diagnosis of OSA in uncomplicated adult patients presenting with signs and symptoms that indicate an increased risk of moderate-to-severe OSA, diagnosis by HSAT has not been substantiated for the evaluation of sleep-disordered breathing in patients taking opioid medications and those who are at increased risk for nonobstructive sleep-disordered breathing. Also, the use of arterial PCO₂ or surrogate (ie, end-tidal PCO₂ or transcutaneous PCO₂) during polysomnography is required for the diagnosis of sleep-related hypoventilation. 23,27

Although opioid therapy can contribute to sleep disruption, daytime symptoms and sleep-disordered breathing, narcotics are also prescribed in rare cases of RLS, a sleep-related movement disorder that is associated with disturbed sleep.²³ Opioids can be especially useful in patients with severe, refractory RLS, who are unresponsive or intolerant to other recommended therapies such as iron, dopamine agonists, antiepileptic medications, or benzodiazepenes.^{9,28–30} The effective opioid doses for RLS are significantly lower than those used for chronic pain; therefore, treatment tends to be well-tolerated and less likely to produce dependence in people with RLS.^{29–31} To promote the assessment of the long-term safety and effectiveness of opioid medications for RLS, a national RLS Opioid Registry has been created.³²

CONCLUSIONS

Chronic opioid therapy requires careful clinical attention because of the serious risks of addiction, abuse, and overdose. While fatigue and disturbed sleep are common symptoms of chronic pain, there is a complex relationship among opioids, sleep and daytime function that can also contribute to fatigue and disturbed sleep. Therefore, it is important for medical providers to be aware that chronic opioid use is associated with changes in sleep architecture and an increased risk of respiratory depression during sleep. Appropriate screening and diagnostic testing with inlaboratory PSG can identify sleep-related hypoventilation, CSA and OSA in people treated with chronic opioid therapy. Although the best therapeutic option for opioid-associated sleep-disordered breathing may be withdrawal of opioids, providers are often faced with the challenge of effectively managing the underlying disorder while also ensuring patient safety, which requires collaborative care. The treatment of sleep-disordered breathing with an effective PAP modality in these patients can improve their health and well-being.

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DISCLOSURE STATEMENT

The authors are the 2018–2020 board of directors of the AASM. This position statement is intended by the AASM to help physicians and other medical providers make decisions about the appropriate care of patients treated with chronic opioid therapy. It is to be used for educational and informational purposes only.