

## Nitazene Dependence Causes Real Withdrawal. Standard Buprenorphine Doses May Fall Short.

### Clinical Scenario

A 22-year-old man with no history of opioid use presents to the emergency department in opioid withdrawal after vaping a product sold online as a cannabinoid; standard-dose buprenorphine fails to control his symptoms. Toxicology later confirms protonitazene in both his blood and the vape liquid.

### The Pearl

Nitazenes are benzimidazole synthetic opioids whose potency can exceed fentanyl, and regular use can produce a genuine physical dependence with a withdrawal syndrome that frontline protocols were not built to treat. A 2026 case report described exactly this: a previously opioid naive young man whose chronic protonitazene vaping led to dependence, then **required buprenorphine doses higher than standard induction levels call for, plus adjunctive diazepam and olanzapine, before withdrawal was controlled.**<sup>1</sup>

This matters because a routine induction or taper calibrated for heroin, oxycodone, or fentanyl withdrawal may leave a nitazene-dependent patient undertreated and at high risk of returning to an unpredictable illicit supply.

### Why It Matters

A New South Wales surveillance cohort identified 27 laboratory-confirmed nitazene exposures over nearly seven years: 20 were unintentional poisonings, and **7 presented in acute withdrawal, a pattern the authors flagged as a novel clinical observation.**<sup>2</sup> Vaping was the most common route, and protonitazene, protonitazepyne, and isotonitazene were the most frequently detected compounds; patients, and even their suppliers, may not know nitazenes are present at all.

A parallel review found no published studies on methadone, buprenorphine, or naltrexone effectiveness specifically in nitazene-dependent patients, leaving clinicians to extrapolate from protocols not built for this potency class.<sup>3</sup> Missing this risks undertreated withdrawal and a faster return to use.

**Bottom Line:** If a patient reports vaping or using a product they did not know contained nitazenes and presents in withdrawal that is not responding to a standard buprenorphine dose, escalate dosing and add adjunctive medication early rather than waiting for a slow taper to catch up.

### References

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2. Roberts DM, Tisdell B, Sajeev MF, Jiranantakan T, Harvey C, Brown JA. Clinical experiences with the nitazene class of synthetic opioids: a cohort study. *Ann Emerg Med.* 2025;86(5):475-483.
3. Nielsen S, Silva JP, Jones JD, et al. Behavioural effects and naloxone effectiveness with new synthetic opioids. *Drug Alcohol Rev.* 2025;45(1):e70040.

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