

VIVITROL (naltrexone extended release)

Effective Date: 1/28/14

Date Developed: 1/28/14 by Robert Sterling, MD

Last Approval Date: 1/26/16, 1/24/17, 1/23/18, 1/22/19,
2/18/20, 2/2/21, 8/3/21, 2/1/22, 1/31/23, 2/13/24,
ARCHIVED 2/18/25

Naltrexone (a pure opioid antagonist) is a derivative of oxymorphone similar in structure to naloxone and nalorphine (a morphine derivative); it acts as a competitive antagonist at opioid receptor sites, showing the highest affinity for mu receptors. Endogenous opioids are involved in modulating the expression of alcohol's reinforcing effects. Naltrexone also modifies the hypothalamic-pituitary-adrenal axis to suppress alcohol consumption.

Authorization:

Alcohol use disorder: Treatment of alcohol use disorder.

Opioid use disorder, mild to severe: For the blockade of the effects of exogenously administered opioids.

Dosing

Alcohol use disorder:

Oral: 50 mg once daily; may increase to 100 mg once daily after 1 week based on response and tolerability

IM: 380 mg once every 4 weeks.

Opioid use disorder

IM: 380 mg once every 4 weeks.

NOTE: Oral naltrexone tablets have not been shown to be more effective than placebo for opioid use disorder due to poor patient compliance

NOTE: Do not initiate therapy until patient has passed a naloxone challenge test or is opioid-free (including tramadol) for at least 7 to 10 days after last opioid use
PRECAUTIONS: injection site reactions; worsening depression; hypersensitivity reactions; unintended withdrawal symptoms if given prior to several days of abstinence; hepatic impairment; significant drug

interactions exist, requiring dose/frequency adjustment or avoidance. Consult drug interactions database for more information.

REFERENCES

1. Kleber HD, "Naltrexone," *J Subst Abuse Treat*, 1985, 2(2):117-22.
2. Mitchell JE, "Naltrexone and Hepatotoxicity," *Lancet*, 1986, 1(8491):1215.
3. O'Connor PG and Kosten TR, "Rapid and Ultrarapid Opioid Detoxification Techniques," *JAMA*, 1998, 279(3):229-34.
4. Ahamad K, Korthuis PT, Lum PJ, Johnson C, Wood E. A delayed injection-site reaction in a patient receiving extended-release naltrexone. *Subst Abus*. 2016;37(2):278-280..2016.113891
5. American Society of Addiction Medicine (ASAM). The ASAM national practice guideline for the treatment of opioid use disorder: 2020 focused update. *J Addict Med*. 2020;14(2S)(suppl 1):1-91. doi:10.1097
6. Croop RS, Faulkner EB, Labriola DF. The safety profile of naltrexone in the treatment of alcoholism. Results from a multicenter usage study. The Naltrexone Usage Study Group. *Arch Gen Psychiatry*. 1997;54(12):1130-1135. doi:10.1001/archpsyc.1997
7. Fishman M. Precipitated withdrawal during maintenance opioid blockade with extended release naltrexone. *Addiction*. 2008;103(8):1399-1401.
8. Garbutt JC, Kranzler HR, O'Malley SS, et al; Vivitrex Study Group. Efficacy and tolerability of long-acting injectable naltrexone for alcohol dependence: a randomized controlled trial. *JAMA*. 2005;293(13):1617-1625.
9. Gastfriend DR. Intramuscular extended-release naltrexone: current evidence. *Ann N Y Acad Sci*. 2011;1216:144-166.
10. Vivitrol (naltrexone) [prescribing information]. Waltham, MA: Alkermes Inc; ~~March 2021~~ September 2022
11. Gowing L, Ali R, White JM. Opioid antagonists with minimal sedation for opioid withdrawal. *Cochrane Database Syst Rev*. 2017;5(5):CD002021. doi:10.1002/14651858.CD002021.pub4
12. American Society of Addiction Medicine (ASAM). The ASAM national practice guideline for the treatment of opioid use disorder: 2020 focused update. *J Addict Med*. 2020;14(2S)(suppl 1):1-91.
13. Gubitosa JC, Terwilliger T, Ukazu A, Gordon E. Naltrexone-associated non-ST-elevated myocardial infarction. *Cureus*. 2020;12(10):e11198. doi:10.7759/cureus.11198
14. Towers CV, Katz E, Weitz B, Visconti K. Use of naltrexone in treating opioid use disorder in pregnancy. *Am J Obstet Gynecol*. 2020;222(1):83.e1-83.e8. doi:10.1016/j.ajog.2019.07.037.
15. Miotto K, McCann MJ, Rawson RA, Frosch D, Ling W. Overdose, suicide attempts and death among a cohort of naltrexone-treated opioid addicts. *Drug Alcohol Depend*. 1997;45(1-2):131-134.

Revision History:

Date Reviewed/No Updates: 1/13/15 by C. Sanders, MD

Date Approved by P&T Committee: 1/27/15

Date Reviewed/No Updates: 1/26/16 by C. Sanders, MD; R. Sterling, MD

Date Approved by P&T Committee: 1/26/16

Date Reviewed/No Updates: 1/24/17 by C. Sanders, MD; R. Sterling, MD

Date Approved by P&T Committee: 1/24/17

Date Reviewed/No Updates: 1/23/18 by C. Sanders, MD; R. Sterling, MD

Date Approved by P&T Committee: 1/23/18

Date Reviewed/No Updates: 1/22/19 by C. Sanders, MD; R. Sterling, MD

Date Approved by P&T Committee: 1/22/19

Date Reviewed/No Updates: 2/18/20 by H. Taekman, MD; R. Sterling, MD

Date Approved by P&T Committee: 2/18/20

Date Reviewed/No Updates: 2/2/21 by H. Taekman, MD; R. Sterling, MD

Date Approved by P&T Committee: 2/2/21

Date Reviewed/Updated: 8/3/21 by H. Taekman, MD; R. Sterling, MD

Date Approved by P&T Committee: 8/3/21

Date Reviewed/No Updates: 2/1/22 by H. Taekman, MD; R. Sterling, MD

Date Approved by P&T Committee: 2/1/22

Date Reviewed/No Updates: 1/31/23 by H. Taekman, MD; R. Sterling, MD

Date Approved by P&T Committee: 1/31/23

Date Reviewed/No Updates: 2/13/24 by H. Taekman, MD; R. Sterling, MD

Date Approved by P&T Committee: 2/13/24

Date Reviewed/Updated: 2/18/25 by H. Taekman, MD; R. Sterling, MD

Date Approved by P&T Committee: 2/18/25

Revision Date	Content Revised (Yes/No)	Contributors	Review/Revision Notes
1/24/17	No	Catherine Sanders, MD; Robert Sterling, MD	Annual review
1/23/18	No	Catherine Sanders, MD; Robert Sterling, MD	Annual review
1/22/19	No	Catherine Sanders, MD; Robert Sterling, MD	Annual review
2/18/20	No	Howard Taekman, MD; Robert Sterling, MD	Annual review
2/2/21	No	Howard Taekman, MD; Robert Sterling, MD	Annual review
8/3/21	Yes	Howard Taekman, MD; Robert Sterling, MD	Minor changes; References added
2/1/22	No	Howard Taekman, MD; Robert Sterling, MD	Annual review
1/31/23	No	Howard Taekman, MD; Robert Sterling, MD	Annual review
2/13/24	No	Howard Taekman, MD; Robert Sterling, MD	Annual review
2/18/25	Yes	Howard Taekman, MD; Robert Sterling, MD	Updated dosing information and references