References and Literature Grading

Is There a Role for Intravenous Sub-Dissociative-Dose Ketamine Administered as an Adjunct to Opioids or as a Single Agent for Acute Pain Management in the ED? (9/6/2015)

- 1. Johansson P, Kongstad P, Johansson A. *The effect of combined treatment with morphine sulphate and low-dose ketamine in a prehospital setting.* Scand J Trauma Resusc Emerg Med. 2009 Nov 27;17:61.
- 2. Jennings PA, Cameron P, et al. *Morphine and ketamine is superior to morphine alone for out-ofhospital trauma analgesia: a randomized controlled trial.* Ann Emerg Med. 2012 Jun;59(6):497-503.
- 3. Jennings PA, Cameron P, Bernard S. *Ketamine as an analgesic in the pre-hospital setting: a systematic review.* Acta Anaesthesiol Scand. 2011 Jul;55(6):638-43
- 4. Galinski M, Dolveck F, et al. *Management of severe acute pain in emergency* settings: ketamine reduces morphine consumption. Am J Emerg Med. 2007 May;25(4):385-90
- Ahern TL, Herring AA, Stone MB, Frazee BW. Effective analgesia with low-dose ketamine and reduced dose hydromorphone in ED patients with severe pain. Am J Emerg Med. 2013 May;31(5):847-51
- 6. Beaudoin FL, Lin C, Guan W, Merchant RC. *Low-dose ketamine improves pain relief in patients receiving intravenous opioids for acute pain in the emergency department: results of a randomized, double-blind, clinical trial.* Acad Emerg Med. 2014 Nov;21(11):1193-202
- Miller JP, Schauer SG, Ganem VJ, Bebarta VS. Low-dose ketamine vs morphine for acute pain in the ED: a randomized controlled trial. Am J Emerg Med. 2015 Jan 7. pii: S0735-6757 (14)00979-6. [Epub ahead of print]
- Motov S, Rockoff B, et al. Intravenous Subdissociative-Dose Ketamine Versus Morphine for Analgesia in the Emergency Department: A Randomized Controlled Trial. Ann Emerg Med. 2015 Mar 26. pii: S0196-0644(15)00191-2. [Epub ahead of print]

Search: 2010-2015

- Tier 1: 1 systematic review
- Tier 2: 4 randomized controlled trials
- Tier 3: 2 prospective observational studies

Reference and Associated Literature Grade	Grade	Quality	Comments

1. Johansson P, <u>Kongstad P</u> , <u>Johansson A</u> . The effect of combined treatment with morphine sulphate and low-dose ketamine in a prehospital setting. <u>Scand J Trauma</u> <u>Resusc Emerg Med.</u> 2009 Nov 27;17:61.	С	Good	Prospective Clinical Cohort study- Supportive
2. Jennings PA, <u>Cameron P</u> , et al. Morphine and ketamine is superior to morphine alone for out-of-hospital trauma analgesia: a randomized controlled trial. <u>Ann Emerg</u> <u>Med.</u> 2012 Jun;59(6):497-503.	A	Outstanding	Randomized Controlled Trial-Supportive
3. Jennings PA, <u>Cameron P</u> , <u>Bernard S</u> . <i>Ketamine as an analgesic in the pre-</i> <i>hospital setting: a systematic review</i> . <u>Acta</u> <u>Anaesthesiol Scand</u> . 2011 Jul;55(6):638-43	D	Good	Systematic review- Supportive (Literature review with large exclusion criteria.) 6 articles included in review. No standardized approach for the 6 of them. Some had very small "N" (4 in one which limits its value.)
4. Galinski <u>M</u> , <u>Dolveck F</u> , et al. Management of severe acute pain in emergency settings: ketamine reduces morphine consumption. <u>Am J Emerg</u> <u>Med.</u> 2007 May;25(4):385-90	A	Good	Prospective, Multicenter, Randomized, Double- blind, Clinical trial. Supportive.
5. Ahern TL, <u>Herring AA</u> , <u>Stone MB</u> , <u>Frazee</u> <u>BW</u> . Effective analgesia with low-dose ketamine and reduced dose hydromorphone in ED patients with severe pain. <u>Am J Emerg Med.</u> 2013 May;31(5):847-51	С	Good	Prospective Observational study Supportive

6. <u>Beaudoin FL</u> , <u>Lin C</u> , <u>Guan W</u> , <u>Merchant</u> <u>RC</u> . Low-dose ketamine improves pain relief in patients receiving intravenous opioids for acute pain in the emergency department: results of a randomized, double-blind, clinical trial. <u>Acad Emerg</u> <u>Med.</u> 2014 Nov;21(11):1193-202	A	Adequate	Randomized, Double- blind, Clinical trial- overall supportive Small sample size and authors' admitted "possible confounding" issues with regard to the statement.
7. Miller JP, Schauer SG, Ganem VJ, Bebarta VS. <i>Low-dose ketamine vs</i> <i>morphine for acute pain in the ED: a</i> <i>randomized controlled trial.</i> Am J Emerg Med. 2015 Jan 7. pii: S0735-6757 (14)00979-6. [Epub ahead of print]	A	Good	Prospective, Randomized, Double- blind, Clinical trial- Supportive small sample size Authors pointed out there were limitations.
8. Motov S, <u>Rockoff B</u> , et al. <i>Intravenous</i> <i>Subdissociative-Dose Ketamine Versus</i> <i>Morphine for Analgesia in the Emergency</i> <i>Department: A Randomized Controlled</i> <i>Trial.</i> <u>Ann Emerg Med.</u> 2015 Mar 26. pii: S0196-0644(15)00191-2. [Epub ahead of print]	A	Good	Prospective, , Randomized, Double- blind, Clinical trial – Supportive small sample size and "the potential for unblinding"