Therapeutic Intranasal Drug Delivery

Needleless treatment options for medical problems

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Concepts:

Featured new IN medication articles

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Hypoglycemia

Hospice

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Education:

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Teaching

Peer Reviewed Articles

Bibliography



Informational Links Related to Intranasal medication delivery

CONTACT INFORMATION for specific questions not addressed on this web site:

E-mail address - info@intranasal.net

(Be aware that I am usually busy caring for patients, spending time with my kids or climbing mountains and only check this e-mail during the times I update the site - about every 4 to 6 weeks. So if you are in a hurry you may be disappointed)

History behind this web site:

The primary author of this site, Tim Wolfe MD, is a practicing emergency physician with a background in EMS. Some of his physician and paramedic colleagues also assist with the writing of this site and many contribute clinical insights.



He trained in a medically underserved county hospital during the onset of the HIV epidemic in the 1980's where he noticed that many of his patients were able to "safely" self medicate using elicit drugs without needles - they used a form of nasal drug delivery called snorting. This concept led him to test the efficacy of having his patients "snort" inexpensive generic medicines (by dripping it in their noses) in hopes of finding both a safe and a painless method of drug delivery for paramedics and ER nurses (who at that time were seroconverting to HCV and sometime HIV simply for doing their jobs). Unfortunately his clinical experience was mixed -

sometimes nasally delivered drugs were very effective, other times not so effective. His epiphany occurred in the middle of the night while dripping a measured dose of a benzodiazepine into a child's nose - the child spit it back in his face. At that moment he realized that he was giving ORAL drugs via the nose which probably explained the variable efficacy he had noticed all along. He needed to change the mode of delivery. If there was only a way to rapidly

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Links

Educational links regarding nasal medication delivery

aerosolize an exact dose of the drug into a fine mist that could be delivered directly to the nasal mucosal with no loss into the environment (nebulization does not achieve these needs since most is not delivered to the nose or to the lung, rather is it lost to the environment, takes time to deliver and usually requires a compliant patient). Through trial and error using compressed air, dual syringes, etc the idea morphed into a very simple syringe driven atomizer. He and a friend started a company selling that device (the MAD - mucosal atomization device).

A few years ago the company was sold and Dr. Wolfe currently receives no royalty nor any money from the MAD, allowing him to write this web site and pursue other interests. Dr. Wolfe consults for interested device and pharmaceutical companies and occasionally is paid to educate their sales people specifically about the MAD device, but he earns no money for any sales of any topical drug delivery products or drugs nor does he have shares in these companies nor do they have any control of this web site. Never the less an author can never completely remove his or her bias from any writings and you will find that Dr. Wolfe feels strongly that nasal drug delivery is a valuable drug delivery adjunct to providing efficient and effective patient care. Furthermore due to his training in a poor, medically underserved community you will find a bias towards using inexpensive generic drugs that are proven by the literature to be clinically effective (rather than focusing on drug levels and bioavailability which do not necessarily translate into clinical effect when applied in the real world - case in point the tiny 0.1 ml volumes that supposedly work via the nose of compliant patients but are extremely difficult to administer in the ER setting). Since the MAD is commonly used in these research studies there is also a great deal of literature that involves that product so a bias could be suggested here as well.

Because he gets questions from clinicians and patients all over the world and because he enjoys teaching about the topic he and some colleagues created this site to help provide a place for the curious clinician seeking detailed information. He apologizes for the somewhat disorganized look of this site - he is just a doctor, not a web master. Someday when he has the time he may rewrite the site to make it less cluttered. If you personally want to communicate with him feel free to do so via the email address above providing your name and affiliations and he will eventually respond.

Note from a Fellow EMS director, emergency physician:

January 14, 2010

Dear Tim,

I am offering this little note to share with you some thoughts regarding the importance of mucosal administration in the care of our patients. It is clear to us in the fields of emergency medicine and EMS medicine that we have had a long neglected opportunity for care. You have brought this advance to the forefront. You saw clearly that through a novel yet simple idea, we could garner a new advantage that had long been missing in clinical care. Because of your insight, we can now approach many of the long-dealt-with dilemmas in emergency care that before often eluded us, indeed caused us to offer in many cases less than optimal care.

One important example is in the treatment of seizures. Those of us who have dealt with pediatric patients suffering from convulsions for example, in the absence of intravenous access, have been left with virtually no other alternative than the per rectum administration of diazepam to halt the seizure. You gathered the budding information in the literature and quickly adapted an invention that could simply, safely treat this difficult problem in a much more proper, much less toxic manner. This has been a great advance in the history of medicine.

Another critical area, the dilemma of "excited delirium," presents a hazard to patient and responder alike. These patients must often be sedated to provide safety in therapy and management. The intranasal route of administration presents a fast alternative that is much

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less hazardous for the responding emergency provider as regards the potential for infectious disease contamination from the patient, especially in the difficult problem of the spread of viruses such as hepatitis C and HIV through needle-stick exposure.

Finally, we all understand how important the prompt treatment of severe pain is to our patients. We now have the capability of prompt administration of fentanyl via the intranasal route and thousands of patients will no longer suffer as a result of this important medical advance.

I would close by thanking you for your contributions to medicine, Tim. You have seen clearly where others did not. Because of your persistence and your great loving heart, patients everywhere – who will never know you – are being treated promptly and humanely, for the benefit of all concerned.

Thank you for your service and dedication.

With kind regards, I am

Respectfully,

Raymond L. Fowler, MD, FACEP

Professor of Emergency Medicine

Chief of EMS Operations

Southwestern Medical Center at Dallas

Attending Emergency Medicine Faculty

Parkland Memorial Hospital

INTERNET DISCUSSIONS regarding IN therapy

Reviews

Synopsis of a review article regarding intranasal medication use in children published in the journal Pediatrics Sept 2010 (click here)

Abstract of this article

Pain discussions

- Intranasal fentanyl relieves breakthrough cancer pain
- Industry sponsored discussion of intranasal pain medications
- <u>Intranasal fentanyl or intravenous morphine for analgesia in adults Best</u> <u>Bets evidence based topic review.</u>
- Intranasal Fentanyl versus parenteral opiates for acute pain in children -Best Bets evidence based topic review.
- Prommer et al, Intranasal fentanyl review article 2011
- <u>Stay on top of kids pain in the ED (use IN fentanyl) ACEP news</u>
 <u>December 2010</u>

EMS links of interest

- The Nose Knows cost effectiveness of transnasal therapy.
- Australian Lay public education regarding treating seizures, calling 112
 and using intranasal midazolam (PDF 0.2 MB)
- <u>New South Wales EMS protocol for pediatric seizures</u> (PDF 0.17 MB)
- Audio interview with Debra Kerr IN naloxone researcher
- <u>Contra Costa EMS update 2010 (Intranasal naloxone pp 38-50)</u>
- Intranasal fentanyl use in ski resort

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Educational links regarding nasal medication delivery

- Inhaled methoxyflurane vs intranasal fentanyl for adult visceral pain in an ambulance
- <u>Dyer, Snuffing out the overdose: The Boston BLS nasal naloxone program</u> <u>slide presentation (11 MB)</u>
- Intranasal Versed usage Paramedic perception of Utility in urban EMS system (Columbus OH)
- Use of IN naloxone by ALS and BLS providers in Boston MA
- <u>National public radio Overdose rescue kits save lives</u>
- <u>Chris Colwell, MD The Nose Knows: Intranasal Medications options are</u> <u>growing (Gathering of Eagles EMS conference feb 2011)</u>
- <u>Excited Delirium in EMS treatment with IN midazolam, Emergency</u> <u>Medicine News October 2010</u>
- <u>Prepackaged syringes of drug ready for use with an atomizer cool idea</u>
- Intranasal naloxone as a layperson administered procedure the Massachusetts experience
- Intranasal fentanyl in EMS discussion (Editorial comment the dose is too low - 1.5 to 2 mcg/kg should be used as initial dose. If you only give an adult 75 mcg you can expect it to fail so don't make this common error. Since this publication, the author has convinced his medical control of the right dose and now uses that shown to be effective on this web site.)
- <u>Callopy and Snyder, Intranasal drug delivery in EMS. EMS today May</u>
 <u>2011</u>

Hospice links of interest

- Palliative info web site by Mike Harlos, MD
- Lecture slides by Dr. Portenoy (Beth Isreal) regarding breakthrough Pain in hospice patients
- <u>www.palliativedrugs.com</u>

Sedation discussions

- <u>Another very detailed link discussing IN Precedex for pediatric sedation in</u> <u>multiple settings in the hospital</u>
- <u>Excited Delirium in EMS treatment with IN midazolam, Emergency</u> <u>Medicine News October 2010</u>

POSTED INTERNET ARTICLES (to go to peer reviewed articles for more PDF and web linked articles - click here)

Nose Brain Pathway articles

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Pain Articles

- Industry sponsored articles in nasal pain medications
- Borland, IN fentanyl vs IV morphine for fractures in children randomized controlled trial, Ann Emerg Med 2007
- Sitte, IN fentanyl for episodic breathlessness, J pain sym manage 2008
- <u>Rickard, IN fentanyl vs IV morphine for fractures in the prehospital</u> setting, 2008
- Heshmati, IN sufentanil for postoperative pain, IJPT 2006
- Wilson, IN diamorphine for pediatric analgesia, J Accid EM 1998

Sedation articles

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- Bkakta, IN midazolam for preoperative sedation, Ind J Anaesth 2007
- <u>Mathai, Preanesthetic sedation of preschool children: Comparison of</u> intranasal midazolam versus oral promethazine. Anesth Essays Res 2011.

Seizure articles

- Lahat, IN midazolam vs IV diazepam for pediatric seizures, BMJ 2000
- <u>Ma, Clinical Guidelines on Management of Prolonged Seizures, HK J</u> <u>Paediatrics 2010</u>
- <u>Klein-Kremer, IN midazolam for treating seizures in children in the</u> <u>emergency setting. Is J Med 2007</u>
- French JA.. Benzo vs Benzo: And the winner is.. Epilepsy Curr 2011;11(5):143-144.

Intranasal Naloxone articles

- <u>Barton, IN naloxone as a needleless alternative to treatment of opioid</u> overdose in the prehospital setting, J Emerg Med 2005
- <u>MMWR Feb 17 2012 Community based opioid overdose prevention</u>
 <u>programs providing naloxone United States 2010</u>
- <u>Time Magazine Lifesaving Overdose antidote should be made more</u> widely available 2012 article
- <u>Time Magazine Lifesaving Overdose antidote should be made more</u> widely available 2012 editorial
- McDermott, C. and N. C. Collins (2012). "Prehospital medication administration: a randomised study comparing intranasal and intravenous routes." Emerg Med Int 2012: 476161.
- <u>RehabCenter.net Naloxone rescue kit downloadable protocol</u>

The PDF file downloaded from their site (click here).

Topical anesthetic articles

- <u>A guidance for the use of topical anesthetics for nasal/oropharyngeal and laryngotracheal procedures</u>
- <u>Chan, Lidocaine spray prior to NGT placement, Hong Kong Medical</u> Journal 2010
- Whit Fisher, MD Procedurettes how to create a NGT frozen curve to enhance passage through nose - link to video

Pediatric reviews

INTRANASAL DRUG DELIVERY REVIEWS AND MISC

- <u>References for incident pain and incident dyspnea protocol</u>
- Pires, Intranasal drug delivery How, why, what for -extensive review, J Pharm Pharmaceut Sci 2009
- <u>Mercus, Influence of head position and anatomy on nasal drug</u> <u>deposition, Rhinology 2006.pdf</u>

INTERNET PROTOCOLS

- Intranasal diamorphine dosing table
- <u>Palliative care incident pain and dyspnea protocol</u>
- Sunshine Hospital emergency department pediatric IN fentanyl protocol

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