

Case-Series of Nurse-Administered Nitrous Oxide for Urinary Catheterization in Children

Judith L. Zier, MD, FAAP*

Gloria J. Drake, CRNA†

Paul C. McCormick, MD‡

Katherine M. Clinch, MD‡

David N. Cornfield, MD‡

BACKGROUND: Children undergoing urologic imaging studies requiring urethral catheterization experience considerable discomfort and psychological distress. Nitrous oxide sedation may mitigate these detriments but the requirement for physician administration has limited the applicability of this technique.

METHODS: Registered nurses underwent the nitrous oxide training requirements prescribed for state licensure of dentists and dental hygienists, with special emphasis on pediatric sedation principles. To evaluate the safety of nurse-administered nitrous oxide, we consecutively enrolled all children (ASA PS I-II) sedated for urethral catheterization for urologic imaging in an observational trial designed to identify sedation-related adverse events.

RESULTS: Nitrous oxide was administered on 1018 occasions. There were no major adverse events (apnea, oxygen saturation <92%). Minor adverse events (diaphoresis, nausea, vomiting) occurred in 4% of patients. Eight patients (1%) were described as over-sedated. In 11 (1%) patients, nitrous oxide provided insufficient sedation for completion of urologic imaging.

CONCLUSIONS: Nitrous oxide sedation can be provided by a nurse-administered program in pediatric radiology. Administration of nitrous oxide for pediatric procedures by adequately trained nursing staff with appropriate multidisciplinary oversight may increase children's access to this sedative/analgesic drug.

(Anesth Analg 2007;104:876-9)

Considerable and well-warranted attention has been focused recently on the issue of pediatric pain management (1,2) and procedural sedation (3). Pain and distress experienced by children during medical procedures increases distress and anxiety during subsequent procedures (4). Urethral catheterization, although not particularly traumatic from an adult perspective, can be especially troublesome and painful in children who lack the emotional or cognitive maturity to cooperate or to understand the reasons for the procedure (5,6). Urethral catheterization is required for urologic imaging with voiding cystourethrography (VCUG) and radionuclide cystography (RNC). As many children with urologic abnormalities will require testing at regular intervals, it is particularly important to use strategies that minimize distress and discomfort. Oral midazolam effectively mitigates the anxiety and distress associated with these procedures, but has a half-life which significantly exceeds the time required for imaging, and often results in

unwanted behavioral side effects (7). Recent data suggest that nitrous oxide (N₂O) is as effective a sedative drug as oral midazolam for VCUG (8). Given the cost and workforce requirements of anesthesiologist or intensivist-administered N₂O sedation, Children's Hospitals and Clinics of Minnesota developed a nurse-administered N₂O program to facilitate procurement of VCUG and RNC studies. If both safe and effective, nurse-administered N₂O procedural sedation is likely to be less costly and more readily available than the traditional anesthesiologist-led procedure. The present study was undertaken to evaluate the safety and efficacy of the program by determining the incidence of adverse events and the ability to successfully complete the imaging using N₂O sedation administered by registered nurses.

METHODS

This study was approved by the IRB of Children's Hospitals and Clinics of MN. Given the observational study design, written informed consent was not required.

Nurse-administered N₂O Program

Registered nurses trained and experienced in monitoring deeply sedated pediatric patients underwent the training requirements prescribed for state licensure of dentists and dental hygienists for N₂O administration. Accreditation entailed attendance at an 8-h course designed to address the pharmacology, toxicity, and environmental safety of N₂O as well as the

From the Divisions of *Pediatric Critical Care; and †Anesthesiology, Children's Hospitals and Clinics of Minnesota, Minneapolis, Minnesota; and ‡Department of Pediatrics, Stanford School of Medicine, Stanford, California.

Address correspondence to Judith L. Zier, MD, Children's Respiratory and Critical Care Specialists, 2545 Chicago Ave. S, Suite 517, Minneapolis, MN 55404. Address e-mail to judy.zier@childrensmn.org.

Reprints will not be available from the author.

Copyright © 2007 International Anesthesia Research Society
DOI: 10.1213/01.ane.0000258763.17768.ce

equipment used for its delivery. After successful completion of this course, the clinical competency of each qualified nurse was assessed during four additional hours of observation and mentored administration of N₂O. The program met the guidelines of the American Nurses' Association for registered nurses charged with the management of patients receiving IV medication for short-term diagnostic procedures (9).

N₂O Administration

A standard dental flowmeter and rubber goods were used for N₂O administration and scavenging. Inhaled N₂O was administered via a continuous flow device (Porter Instrument Company, Hatfield, PA) which allows titration of N₂O concentration from zero to a maximum of 70%, with oxygen as the remaining gas. Unlike the commercially available fixed 50:50 N₂O:O₂ mixture, there was no need for the patient to overcome a demand valve to maintain N₂O delivery. The equipment incorporates built-in safety features, including a non-rebreathing valve, emergency air intake valve, and fail-safe device that automatically terminates the flow of N₂O in the event of an interruption in oxygen flow. The equipment includes an apparatus for exhaled gas scavenging and evacuation. An adequate seal could be comfortably maintained using the nasal hood over the nose of the older child or over the nose and mouth of a toddler. Before clinical use, the equipment was assembled and tested for N₂O leakage by the hospital's biomedical department. Badge dosimetry monitoring was performed periodically to ensure compliance with National Institute of Occupational Safety and Health N₂O occupational exposure limit of less than 25 ppm time-weighted average.

N₂O Sedation for Urologic Imaging

All patients underwent a pre-sedation assessment before sedation administration to identify potential contraindications to sedation (e.g., gastroesophageal reflux, craniofacial abnormalities) and inhaled N₂O (e.g., pneumothorax, bowel obstruction). A set of vital signs, including temperature, respiratory rate, heart rate, arterial blood pressure, and baseline pulse oximetry reading, were obtained during the pre-sedation assessment. By hospital policy, all patients with ASA classification >II were ineligible to receive nurse-administered N₂O and did not participate in this study. During the initial 4 mo of the study, patients were kept fasting for a minimum of 4 h before sedation. On the basis of further literature review (10,11) and interim analysis of the study data, subsequent patients were instructed to restrict intake to at most a light meal for 4 h before the procedure. N₂O was administered at 70% N₂O/30% O₂ until completion of urethral catheterization. After catheterization, 100% oxygen was administered for 2–5 min. Throughout the N₂O administration, and until the child returned to the pre-sedation level of alertness, the patient was monitored with continuous

Table 1. Adverse Effects of Nitrous Oxide for Urologic Imaging in 1018 Children

	N	%
Apnea (>15 s)	0	0
O ₂ saturation <92% (>1 s)	0	0
Diaphoresis	7	1
Nausea	9	1
Vomiting	21	2
Other (crying, pallor, agitation)	8	1
Total unique cases with any side effect	36	4

pulse oximetry and direct nursing observation. No additional arterial blood pressure recordings were obtained. VCUG or RNC was then performed as determined by radiology protocol. Venous access was not obtained in any child, either for the imaging study or, per policy, for N₂O sedation.

Data Collection

From September, 2004 through April, 2006, all children receiving N₂O sedation for urethral catheterization for VCUG or RNC in the radiology department of the St. Paul campus of Children's Hospitals and Clinics of MN were enrolled consecutively in the study. Data collection sheets were attached to each N₂O sedation order to ensure compliance with data collection for each patient. Data collected for this study included presence or absence of the side effects listed in the Table 1, duration of N₂O administration (<15, 15–30, >30 min), and whether the level of sedation was sufficient to allow successful completion of the procedure. Level of sedation was assessed as presence or absence of over-sedation, defined as sedation deeper than a drug-induced state during which patients respond normally to verbal commands.

RESULTS

N₂O was administered on 1018 occasions for urethral catheterization for either VCUG or RNC during the 20-mo study period. Review of departmental scheduling records revealed that there were 3398 VCUGs and RNCs scheduled during that time. One thousand ninety-three procedures were scheduled with sedation, representing data collection on a minimum of 93% of possible sedation encounters. The actual percentage may be higher, as last minute cancellations were not removed from the records. Patients ranged in age from 11 mo to 17 yr, with a mean age of 5.4 yr and median of 4.8 yr. Almost all (94%, *n* = 952) received N₂O for <15 min. Sixty-three patients (6%) received N₂O between 15 and 30 min and three patients (0.3%) received N₂O for longer than 30 min. No patient developed apnea (>15 s) or oxygen saturation below 92% (>1 s) at any time during N₂O administration or recovery. Thirty-six patients (4%) had minor adverse effects, including nausea, diaphoresis, and/or vomiting (Table 1). Eleven procedures (1%) were unsuccessful due to sedation failure.

Eight patients (1%) were described as over-sedated. Charts of all eight patients were reviewed. One patient was described as “snoring,” with rapid response to discontinuation of N₂O and initiation of 100% oxygen. None required airway intervention.

DISCUSSION

Dentists have been administering N₂O alone or combined with other sedatives and analgesics since the 1800s (12). Eighty-five percent of pediatric dentists use N₂O for patient sedation (13). Although an article published in the *Journal of the American Medical Association* more than 20 yr ago described the use of N₂O for more than 3000 patients in a general pediatric office in UT, (14) the use of N₂O for pediatric procedural sedation in the United States likely remains sporadic. Use of N₂O outside of the operating room or dental clinic has been reported primarily in the pediatric emergency department for laceration repair (15,16) or fracture reduction (17,18).

N₂O administration by non-physician providers is routine. Either dentists or dental hygienists can deliver N₂O in much of the United States. Advanced practice nurses deliver N₂O to pediatric patients for minor surgical procedures in the United States (19). Registered nurses deliver N₂O in the emergency department and outpatient setting in Australia and England (20,21). The safety of N₂O administration to children by pre-hospital providers, including lay responders, has been documented (22).

A recent article (23) addressed the efficacy of a tiered approach to pediatric sedation including nurse-administered protocols. The authors stressed the importance of accurately matching the pharmacologic approach to appropriately trained personnel. The present report provides data that supports the notion that inhaled N₂O can be used by registered nurses for specific urologic procedures. While several articles have addressed the safety of N₂O sedation for a variety of pediatric procedures (24–31), the current study adds to this body of knowledge by reporting the largest series of patients sedated with N₂O using a nurse-administered protocol. In the present series, the use of N₂O to expedite the performance of urologic imaging studies in more than 1000 children did not result in a single major adverse event (apnea or arterial oxygen saturation <92%). Despite using N₂O at a concentration of 70%, the incidence of minor adverse effects of N₂O (nausea, vomiting, diaphoresis) in the present study was less than previously reported (20,24). The difference may derive from the relatively short duration of administration of N₂O required to expedite urethral catheterization.

Restriction of N₂O administration privileges to physicians or nurse anesthetists may not only considerably limit the use of N₂O as a sedative/analgesic drug due to workforce requirements, but also increase the cost of the procedure. An editorial (32) critical of

N₂O use for laceration repair in a pediatric emergency department cited the labor-intensive need for a physician to administer the N₂O in addition to the physician performing the procedure and asked the question, “can the substantial logistical hurdle of a separate sedating physician be overcome through special nurse training in this technique?” The current study seems to answer that question in the affirmative.

The present study does not address the quality of sedation for urethral catheterization. Rather the study end-point was the successful completion of the imaging study. Similarly, the study does not address discomfort that may have been encountered during the remainder of the urologic imaging (bladder filling, voiding). Nevertheless, our experience mirrors the report of Keidan et al. (8) wherein N₂O provided a reduction in anxiety and distress associated with urologic imaging comparable to oral midazolam, but with a shorter recovery time.

N₂O administered at <50% concentration in oxygen with no other sedative or analgesic medications is recognized as minimal sedation (33). For this study, patients characterized as “over-sedated” correspond to a level of sedation deeper than minimal sedation, as outlined in the ASA Continuum of Depth of Sedation (33). Even though N₂O was administered at a concentration of 70%, more than 99% of the study patients remained at the level of minimal sedation, as judged by responsiveness to verbal stimulation. The fact that eight patients reached a level of moderate sedation nevertheless reinforces the importance of preparation to appropriately manage patients at the level of moderate sedation when N₂O is used at this concentration. Measurement of patient sedation level during N₂O administration using a validated sedation scale is important information for future study. Rates of dysphoria should also be measured in future studies with larger sample sizes to better estimate rates of adverse events.

In conclusion, the present data support the notion that N₂O sedation can be safely and effectively provided using a nurse-administered program in a hospital-based radiology department. Administration of N₂O for pediatric procedures by adequately trained nursing staff with appropriate multidisciplinary oversight may increase children’s access to this sedative/analgesic drug.

ACKNOWLEDGMENTS

Thank you to Marsha Finkelstein and Song Chen, Center for Care Innovation and Research of Children’s Hospitals and Clinics of MN, for their contributions to this research.

REFERENCES

1. Brislin RP, Rose JB. Pediatric acute pain management. *Anesthesiol Clin North Am* 2005;23:789–814.
2. Young KD. Pediatric procedural pain. *Ann Emerg Med* 2005;45:160–71.
3. Cravero JP, Bilke GT. Review of pediatric sedation. *Anesth Analg* 2004;99:1355–64.
4. Chen E, Zeltzer LK, Craske MG, Katz ER. Alteration of memory in the reduction of children’s distress during repeated aversive medical procedures. *J Consult Clin Psychol* 1999;67:481–90.

5. Phillips D, Watson AR, Collier J. Distress and radiological investigations of the urinary tract in children. *Eur J Pediatr* 1996;155:684-7.
6. Stashinko EE, Goldberger J. Test or trauma? The voiding cystourethrogram experience of young children. *Issues Compr Pediatr Nurs* 1998;21:85-96.
7. Elder JS, Longenecker R. Premedication with oral midazolam for voiding cystourethrography in children: safety and efficacy. *AJR* 1995;164:1229-32.
8. Keidan I, Zaslansky R, Weinberg M, et al. Sedation during voiding cystourethrography: comparison of the efficacy and safety of using oral midazolam and continuous flow nitrous oxide. *J Urol* 2005;174:1598-601.
9. Odom-Forren J. The evolution of nurse-monitored sedation. *J Perianesth Nurs* 2005;20:385-98.
10. Babl FE, Puspitadewi A, Barnett P, et al. Preprocedural fasting state and adverse events in children receiving nitrous oxide for procedural sedation and analgesia. *Pediatr Emerg Care* 2005;21:736-43.
11. American Academy of Pediatric Dentistry. Guideline on appropriate use of nitrous oxide for pediatric dental patients. *Pediatr Dent* 2005;27:107-9.
12. Yagiela JA. Office-based anesthesia in dentistry. *Dent Clin North Am* 1999;43:201-15.
13. Dock M, Creedon RL. Pharmacologic management of patient behavior. In: McDonald RE, Avery DR, Dean JA, eds. *Dentistry for the child and adolescent*. St. Louis: Mosby, 2004:285-311.
14. Griffin GC, Campbell VD, Jones R. Nitrous oxide-oxygen sedation for minor surgery. Experience in a pediatric setting. *JAMA* 1981;245:2411-3.
15. Luhmann JD, Kennedy RM, Porter FL, et al. A randomized clinical trial of continuous-flow nitrous oxide and midazolam for sedation of young children during laceration repair. *Ann Emerg Med* 2001;37:20-7.
16. Gamis AS, Knapp JF, Glenski JA. Nitrous oxide analgesia in a pediatric emergency department. *Ann Emerg Med* 1990;19:843-4.
17. Gregory PR, Sullivan JA. Nitrous oxide compared with intravenous regional anesthesia in pediatric forearm fracture manipulation. *J Pediatr Orthop* 1996;12:187-91.
18. Hennrikus WL, Shin AY, Klingelberger CE. Self-administered nitrous oxide and a hematoma block for analgesia in the outpatient reduction of fractures in children. *J Bone Joint Surg Am* 1995;77:335-9.
19. Burnweit C, Diana-Zerpa JA, Nahmad MH, et al. Nitrous oxide analgesia for minor pediatric surgical procedures: an effective alternative to conscious sedation? *J Pediatr Surg* 2004;39:495-9.
20. Frampton A, Browne GJ, Lam LT, et al. Nurse administered relative analgesia using high concentration nitrous oxide to facilitate minor procedures in children in an emergency department. *Emerg Med J* 2003;20:410-3.
21. Cleary AG, Ramanan AV, Baildam E, et al. Nitrous oxide analgesia during intra-articular injection for juvenile idiopathic arthritis. *Arch Dis Child* 2002;86:416-8.
22. Faddy SC, Garlick SR. A systematic review of the safety of analgesia with 50% nitrous oxide: can lay responders use analgesic gases in the prehospital setting? *Emerg Med J* 2005;22:901-8.
23. Dalal PG, Murray D, Cox T, et al. Sedation and anesthesia protocols used for magnetic resonance imaging studies in infants: provider and pharmacologic considerations. *Anesth Analg* 2006;103:863-8.
24. Kanagasundaram SA, Lane LJ, Cavalletto BP, et al. Efficacy and safety of nitrous oxide in alleviating pain and anxiety during painful procedures. *Arch Dis Child* 2001;84:492-5.
25. Annequin D, Carbajal R, Chauvin P, et al. Fixed 50% nitrous oxide oxygen mixture for painful procedures: a French survey. *Pediatrics* 2000;105:e47.
26. Michaud L, Gottrand F, Ganga-Zandzou PS, et al. Nitrous oxide sedation in pediatric patients undergoing gastrointestinal endoscopy. *J Pediatr Gastroenterol Nutr* 1999;28:310-4.
27. Bar-Meir E, Zaslansky R, Regev E, et al. Nitrous oxide administered by the plastic surgeon for repair of facial lacerations in children in the emergency room. *Plast Reconstr Surg* 2006;117:1571-5.
28. Fishman G, Botzer E, Marouani N, DeRowe A. Nitrous oxide-oxygen inhalation for outpatient otologic examination and minor procedures performed on the uncooperative child. *Int J Pediatr Otorhinolaryngol* 2005;69:501-4.
29. Ekbom K, Jakobsson J, Marcus C. Nitrous oxide is a safe and effective way to facilitate procedures in paediatric outpatient departments. *Arch Dis Child* 2005;90:1073-6.
30. Faroux B, Onody P, Gall O, et al. The efficacy of premixed nitrous oxide and oxygen for fiberoptic bronchoscopy in pediatric patients. *Chest* 2004;125:315-21.
31. Gall O, Annequin D, Benoit G, et al. Adverse events of premixed nitrous oxide and oxygen for procedural sedation in children. *Lancet* 2001;358:1514-5.
32. Krauss B. Continuous-flow nitrous oxide: searching for the ideal procedural anxiolytic for toddlers. *Ann Emerg Med* 2001;37:61-2.
33. American Society of Anesthesiologists Task Force on Sedation and Analgesia by Non-Anesthesiologists. Practice guidelines for sedation and analgesia by non-anesthesiologists. *Anesthesiology* 2002;96:1004-17.