

Policy #: 100

Original policy date: 6/1/09

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Title

Inhaled Nitric Oxide as a Treatment of Hypoxic Respiratory Failure in Neonates

Description

Inhaled nitric oxide (iNO) is an effective treatment in the near-term to full-term neonate diagnosed with persistent pulmonary hypertension (PPHN) as an isolated condition or as an associated condition resulting from any of the following: respiratory distress syndrome (hyaline membrane disease), meconium aspiration syndrome, pneumonia, sepsis, congenital diaphragmatic hernia and lung hypoplasia.

Nitric oxide in its natural form is a product of cell metabolism that acts as a blood vessel relaxant (vasodilator) which increases blood flow to the tissues; it also regulates the binding and releasing of oxygen to hemoglobin. When used as an inhalant to treat PPHN nitric oxide produces selective pulmonary vasodilatation and redistributes pulmonary blood flow from areas of the lung with low gas exchange capability (decrease ventilation capacity) to the healthier lung tissue with better gas exchange capability, thus improving oxygenation.

The effect of iNO and ventilator support on improved oxygenation also reduces the need for the use of a more surgically invasive lung oxygenation treatment known as extracorporeal membrane oxygenation (ECMO).

When services are covered for all Products (including Medicare HMOB, Medicare PPO Blue and Blue Medicare PFFS PlusRx Products)

We cover **inhaled nitric oxide** when it is administered as a component of treatment of hypoxic respiratory failure in neonates born at 34 or more weeks of gestation.

When services are not covered for all Products (including Medicare HMOB, Medicare PPO Blue and Blue Medicare PFFS PlusRx Products)

We do not cover indications for **inhaled nitric oxide**, including, but not limited to the following since it is considered investigational¹ and does not meet the BCBSMA Medical Technology Assessment Guidelines, #350:

- For adults with respiratory distress (ARDS) *or*
- For premature neonates.

Individual consideration

All our medical policies are written for the majority of people with a given condition. Each policy is based on medical science. For many of our medical policies, each individual's unique clinical circumstances may be considered in light of current scientific literature. For consideration of an individual patient, physicians may send relevant clinical information to:

For services already billed

Blue Cross Blue Shield of Massachusetts
Provider Appeals
P. O. Box 986065
Boston, MA 02298

Prior to performance of service

Blue Cross Blue Shield of Massachusetts,
Case Creation/Medical Policy
One Enterprise Drive
Quincy, MA 02171
Tel: 1-800-327-6716
Fax: 1-888-641-5330

Managed care guidelines

- Medicare HMO Blue members: referrals are not required.
- Any specialist visit requires a referral for Medicare HMO Blue.
- For all other Managed Care plans, any specialist visit requires a referral.
- Authorization for inpatient admission is required.

Indemnity and PPO guidelines

All authorization requirements are determined by the individual's subscriber certificate, however:

- Authorizations are required for all inpatient services.
- Authorizations are not required for most outpatient services as determined by the individual's subscriber certificate.
- Referrals to a specialist are not required.

Coding information

Procedure codes are from current CPT, HCPCS Level II, Revenue Code, and/or ICD-9-CM manuals, as recommended by the American Medical Association, Centers for Medicare and Medicaid Services and American Hospital Associations. Blue Cross Blue Shield Association national codes may be developed when appropriate.

The following codes are included below for informational purposes. Inclusion or exclusion of a code does not constitute or imply member coverage or provider reimbursement. Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage as it applies to an individual member.

Facility

ICD-9-CM procedure code 00.12, administration of inhaled nitric oxide

Other information

- For our Medical Technology Assessment Guidelines, see document # 350.

Policy update history

New Policy, effective 6/2009.

Footnotes

¹ Based upon BCBSA national policy, *Inhaled Nitric Oxide as a Treatment of Hypoxic Respiratory Failure*, 8.01.37.

References**References for Footnote 1**

Neonatal Inhaled Nitric Oxide Study Group. Inhaled nitric oxide in full-term and nearly full-term infants with hypoxic respiratory failure. *N Engl J Med* 1997; 336(9):597-604.

1. Clark RH, Kueser TJ, Walker MW et al. Low-dose nitric oxide therapy for persistent pulmonary hypertension of the newborn. Clinical Inhaled Nitric Oxide Research Group. *N Engl J Med* 2000; 342(7):469-74.
2. Finer NN, Sun JW, Rich W et al. Randomized, prospective study of low-dose versus high-dose inhaled nitric oxide in the neonate with hypoxic respiratory failure. *Pediatrics* 2001; 108(4):949-55.
3. American Academy of Pediatrics. Use of inhaled nitric oxide. *Pediatrics* 2000; 106(2 pt 1):344-5.
4. Schreiber MD, Gin-Mestan K, Marks JD et al. Inhaled nitric oxide in premature infants with the respiratory distress syndrome. *N Engl J Med* 2003; 349(22):2099-107.
5. Martin RJ. Nitric oxide for preemies – not so fast. *N Engl J Med* 2003; 349(22):2157-9.
6. Van Meurs KP, Wright LL, Ehrenkranz RA et al. Inhaled nitric oxide for premature infants with severe respiratory failure. *N Engl J Med* 2005; 353(1):13-22.
7. Field D, Elbourne D, Truesdale A et al. Neonatal ventilation with inhaled nitric oxide versus ventilatory support without inhaled nitric oxide for preterm infants with severe respiratory failure. *Pediatrics* 2005; 115(4):926-36.
8. Martin RJ, Walsh MC. Inhaled nitric oxide for preterm infants - who benefits? *N Engl J Med* 2005; 353(1):82-4.
9. Dellinger RP, Zimmerman JL, Taylor RW et al. Effects of inhaled nitric oxide in patients with acute respiratory distress syndrome: results of a randomized phase II trial. Inhaled Nitric Oxide in ARDS Study Group. *Crit Care Med* 1998; 26(1):15-23.
10. Lundin S, Mang H, Smithies M et al. Inhalation of nitric oxide in acute lung injury: results of a European multicentre study. The European Study Group of Inhaled Nitric Oxide. *Intensive Care Med* 1999; 25(9):911-9.
11. Taylor RW, Zimmerman JL, Dellinger RP et al. Low-dose inhaled nitric oxide in patients with acute lung injury: a randomized controlled trial. *JAMA* 2004; 291(13):1603-9.
12. Ballard RA, Truog WE, Cnaan A et al. Inhaled nitric oxide in preterm infants undergoing mechanical ventilation. *N Engl J Med* 2006; 355(4):343-53.
13. Kinsella JP, Cutter GR, Walsh WF et al. Early inhaled nitric oxide therapy in premature newborns with respiratory failure. *N Engl J Med* 2006; 355(4):354-64.
14. Hintz SR, Van Meurs KP, Perritt R et al. Neurodevelopmental outcomes of premature infants with severe respiratory failure enrolled in a randomized controlled trial of inhaled nitric oxide. *J Pediatr* 2007; 151(1):16-22.
15. Stark AR. Inhaled NO for preterm infants – getting to yes? *N Engl J Med* 2006; 355(4):404-6.
16. Steinhorn RH, Porta NF. Use of inhaled nitric oxide in the preterm infant. *Curr Opin Pediatr* 2007; 19(2):137-41.
17. Angus DC, Clermont G, Linde-Zwirble WT et al. Healthcare costs and long-term outcomes after acute respiratory distress syndrome: a phase III trial of inhaled nitric oxide. *Crit Care Med* 2006; 34(12):2883-90.
18. Di Fiore JM, Hibbs AM, Zadell AE et al. The effect of inhaled nitric oxide on pulmonary function in preterm infants. *J Perinatol* 2007; 27(12):766-71.
19. Su PH, Chen JY. Inhaled nitric oxide in the management of preterm infants with severe respiratory failure. *J Perinatol* 2008; 28(2):112-6.
20. Hibbs AM, Walsh MC, Martin RJ et al. One-year respiratory outcomes of preterm infants enrolled in the Nitric Oxide (to Prevent) Chronic Lung Disease trial. *J Pediatr* 2008; June 3 [epub ahead of print].
21. Hintz SR, Van Meurs KP, Perritt R et al. Neurodevelopmental outcomes of premature infants with severe respiratory failure enrolled in a randomized controlled trial of inhaled nitric oxide. *J Pediatr* 2007; 151(1):16-22.
22. Barrington KJ, Finer NN. Inhaled nitric oxide for preterm infants: a systematic review. *Pediatrics* 2007; 120(5):1088-99.

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