

## Letter to the Editor

# Reconsidering milrinone as a primary treatment for neonatal pulmonary hypertension: an alternative to inhaled nitric oxide in resource-limited Asian settings

Sir,

Neonatal pulmonary hypertension is a commonly encountered condition with underlying low systemic blood pressure and reduced cardiac output secondary to right ventricular overload and myocardial dysfunction.<sup>1</sup> Research indicates a high incidence of this condition, occurring in 0.5 to 7 neonates per 1000 live births, with a mortality rate of 4-33%, underscoring the need for early diagnosis and effective management.

The standard treatment protocols for persistent pulmonary hypertension of the newborn (PPHN) involve the use of inhaled nitric oxide (iNO) as the first-line therapy. However, the unavailability of iNO in developing countries necessitates the exploration of alternative therapies. A randomized clinical trial conducted in Iran demonstrated that milrinone is an equally effective treatment for PPHN, with both inhaled and infused milrinone showing comparable benefits and minimal adverse effects.<sup>2</sup>

A case report by Tzialla et al highlights improved outcomes in neonatal pulmonary hypertension when intravenous milrinone was used as an adjunct to iNO in cases resistant to iNO alone.<sup>3</sup> Milrinone inhibits phosphodiesterase-3, thereby improving pulmonary hemodynamics through its inotropic and lusitropic effects. Its use in neonates with pulmonary hypertension has been associated with significant elevation in PaO<sub>2</sub>, reduction of oxygen index, and decreased reliance on iNO. Echocardiographic findings further reveal lower pulmonary artery pressure, improved right and left ventricular output, and reduced right-to-left shunting.<sup>4</sup>

Implementing milrinone as the primary treatment modality could serve as a cost-effective strategy, reducing iNO demand and hospital stay durations.<sup>5</sup> Resource-limited countries could significantly benefit from using milrinone alone or as an adjunct to iNO, offering both short- and long-term advantages for patients and the healthcare system.<sup>6</sup>

Considering the high cost and prolonged hospital stay associated with iNO, I urge higher authorities to promote the use of milrinone in PPHN treatment and conduct further studies to establish its role as a primary therapy,

particularly in regions with limited iNO access. This shift could improve patient outcomes while alleviating the financial burden on families and healthcare institutions.

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