1	Title: Pulse oximetry in paediatric primary care: Catalyzing implementation in			
2	low-income and middle-income countries			
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Each year pneumonia kills more children before their fifth birthday than any
other infectious disease.¹ To end preventable deaths of newborns and
children under five by 2030 (United Nations Sustainable Development Goal
3.2), effective primary care interventions for child pneumonia are needed.

Hypoxaemia, a low blood oxygen level, is a key risk factor for child pneumonia mortality.² In high-income settings, pulse oximeters, non-invasive portable devices that measure the peripheral arterial oxyhaemoglobin saturation (SpO₂), have been used in routine paediatric clinical practice for over 30 years.³ In contrast, most paediatric primary care settings in low-income and middle-income countries (LMICs) do not routinely use pulse oximeters at all.³

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While barriers to implementation have included cost and weak maintenance and supervision structures, a lack of policy recommendations has meant pulse oximeter roll-out has lacked prioritization and investment. We argue, that two knowledge gaps underpin this – device selection and high quality evidence – that if addressed can further catalyze both policy and demand for pulse oximetry in paediatric primary care in LMICs.

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First, what do clinicians and nurses need from a pulse oximeter in LMIC paediatric primary care? They need accurate devices designed to work on small, distressed children even when they are moving or have compromised perfusion. If pulse oximeters are to be used as screening tools among children with suspected pneumonia (i.e., children with observed or reported cough or difficult breathing) – our recommendation – then devices must work

74 quickly in overburdened facilities reading >90% of SpO₂ measurements within 75 120 seconds. Devices must be robust, incorporate reusable probes, disinfect 76 easily, work despite electricity outages and with rechargeable batteries, and 77 employ a simple and intuitive interface. To date, surprisingly few pulse oximeters meet these requirements and potential purchasers in LMICs have 78 79 no access to independent device evaluation. Instead, when LMICs procure 80 pulse oximeters they are purchased solely on price and manufacturer 81 specifications. Evidence comparing performance of different models is scarce. 82 especially regarding LMIC performance in children. Recent studies have confirmed that not all inexpensive pulse oximeters are accurate,⁴ which 83 84 makes them unsuitable for paediatric use, and that even expensive devices 85 perform differently under certain conditions common to children in LMICs, such as motion and low perfusion.^{5,6} 86

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88 Although formal cost effectiveness analyses are missing, basic cost 89 projections suggest oximeters may be a best buy for LMICs. Specifically, if 90 LMICs consider pulse oximeter costs on a per patient basis a \$345 USD 91 investment in one quality device (\$250 USD/unit) with three additional 92 paediatric probes (\$25/probe) and one spare battery (\$20/battery) would cost 93 less than \$0.07 USD per patient over five years in a clinic serving three to four 94 children daily. LMICs must have the ability to transparently determine the 95 most appropriate device for use with children in their setting, considering cost, 96 performance, durability and usability.

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98 Next, quality pulse oximetry and hypoxaemia data on children accessing

primary care services in LMICs is scarce. While there is evidence hypoxaemia
is common and that pulse oximeters effectively identify children with
hypoxaemia in hospitals,^{7,8} similar data at the primary care level is lacking,
especially outcome data, prevalence data, and healthcare worker device use
and decision-making data.

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105 A recently published World Health Organization report indicates that outcome data may soon be available from both Malawi and Bangladesh.⁹ Although one 106 107 large study in Malawi showed hypoxaemia was prevalent and government-108 sector healthcare providers effectively used pulse oximeters during paediatric primary care,¹⁰ similar data is needed from other countries and regions. Local 109 data is also required to understand the optimal SpO₂ threshold for hospital 110 111 referral. In addition to mortality risk, the optimal threshold is likely to be driven 112 by two factors, altitude and health system capacity. Although our understanding is limited, children adapted to living at higher altitudes are likely 113 to be more tolerant of a lower SpO₂ than children at lower altitudes.^{11,12} Health 114 system capacity will also differ in LMICs such that a one-size-fits-all SpO₂ 115 116 threshold may no longer be appropriate. Areas with greater health system 117 capacity may be able to accommodate higher SpO₂ referral thresholds while 118 areas with more limited capacity may not. In addition to driving policy, such 119 data will inform LMICs where to locally prioritize distribution of pulse 120 oximeters, training and supervision. 121

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123 Health Research (RESPIRE), Save the Children (United

Kingdom)/GlaxoSmithKline (INSPIRING) aim to pilot pulse oximeters during
primary care in LMICs and may address some of these evidence gaps. We
hope these projects and others will expand our understanding of how
oximeters may improve LMIC paediatric primary care and help end
preventable child deaths from pneumonia.

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130 Citations

131 1. Liu L, Oza S, Hogan D, Chu Y, Perin J, Zhu J, Lawn JE, Cousens S, 132 Mathers C, Black RE. Global, regional, and national causes of under-5 133 mortality in 2000-15: an updated systematic analysis with implications 134 for the Sustainable Development Goals. Lancet 2016; 388: 3027–3035. 135 2. Lazzerini M, Sonego M, Pellegrin MC. Hypoxaemia as a Mortality Risk Factor in Acute Lower Respiratory Infections in Children in Low and 136 137 Middle-Income Countries: Systematic Review and Meta- Analysis. 138 PLos ONE 2015; 10: e0136166. 139 King C, Graham H, McCollum ED. Point-of-Care Pulse Oximetry for 3. 140 Children in Low-Resource Settings. In: Atkinson K, Mabey D, eds. 141 Revolutionizing Tropical Medicine: Point of Care Tests, New Imaging 142 Technologies, and Digital Health. Hoboken, New Jersey, USA: John Wiley and Sons Inc; 2019: 327-43. 143 144 4. Lipnick MS, Feiner JR, Au P, Bernstein M, Bickler PE. The Accuracy of 145 6 Inexpensive Pulse Oximeters Not Cleared by the Food and Drug Administration: The Possible Global Public Health Implications. 146

147 Anesth Analg 2016; **123**: 338–45.

148	5.	Louie A, Feiner JR, Bickler PE, Rhodes L, Bernstein M, Lucero J. Four
149		Types of Pulse Oximeters Accurately Detect Hypoxia during Low
150		Perfusion and Motion. Anesthesiology 2018; 128 : 520–30.
151	6.	Baker K. Can improved paediatric pneumonia diagnostic aids support
152		frontline health workers in low resource settings? Large scale
153		evaluation of four respiratory rate timers and five pulse oximeters in
154		Cambodia, Ethiopia, South Sudan and Uganda. 2019.
155		https://openarchive.ki.se/xmlui/handle/10616/46833 (Accessed
156		September 26, 2019).
157	7.	Subhi R, Adamson M, Campbell H, Weber M, Smith K, Duke T. The
158		prevalence of hypoxaemia among ill children in developing countries: a
159		systematic review. Lancet Infect Dis 2009; 9: 219–27.
160	8.	Duke T, Wandi F, Jonathan M, Matai S, Kaupa M, Saavu M, Subhi R,
161		Peel D. Improved oxygen systems for childhood pneumonia: a
162		multihospital effectiveness study in Papua New Guinea. Lancet 2008;
163		372 : 1328–33.
164	9.	Exploratory meeting to review new evidence for Integrated
165		Management of Childhood Illness (IMCI) danger signs. Geneva: World
166		Health Organization, 2019.
167		https://apps.who.int/iris/bitstream/handle/10665/326100/WHO-MCA-
168		19.02-eng.pdf?ua=1 (Accessed September 26, 2019)
169	10.	McCollum ED, King C, Deula R, Zadutsa B, Mankhambo L, Nambiar B,
170		Makwenda C, Masache G, Lufesi N, Mwansambo C, Costello A,
171		Colbourn T. Pulse oximetry for children with pneumonia treated as

- 172 outpatients in rural Malawi. Bull World Health Organ 2016; 94: 893–
- 173 **902**.
- 174 11. Subhi R, Smith K, Duke T. When should oxygen be given to children at
- 175 high altitude? A systematic review to define altitude-specific
- 176 hypoxaemia. *Arch Dis Child* 2009; **94**: 6-10.
- 177 12. Tasker RC. Oxygen and living at altitude. *Arch Dis Child* 2009; **94**: 1-2.