

Evaluation of 'TRY' an algorithm for neonatal CPAP in low-income settings.

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ABSTRACT:

Background: Non-invasive respiratory support using continuous positive airway pressure (bCPAP) is useful in treating babies with respiratory distress syndrome. Despite its proven clinical and cost effectiveness, implementation is hampered by the inappropriate administration of bCPAP in low-resource settings. A clinical algorithm - 'TRY' (based on **T**one: good, **R**espiratory distress: **Y**es: heart rate above 100b/min) has been developed to correctly identify which newborns would benefit most from bCPAP in a teaching hospital in Malawi.

Objective: To evaluate the reliability, sensitivity and specificity of TRY when employed by nurses in a Malawian district hospital.

Methods: Nursing staff in a Malawian district hospital baby unit were asked, over a 2-month period, to complete TRY assessments for every newly admitted baby with the inclusion criteria: clinical evidence of respiratory distress and/or birth weight less than 1.3kg. A visiting paediatrician, blinded to the nurses' assessments, concurrently assessed each baby providing both a TRY assessment and a clinical decision regarding the need for CPAP administration. Interrater reliability was calculated comparing nursing and paediatrician TRY assessment outcomes. Sensitivity and specificity were estimated comparing nurse TRY assessments against the paediatrician's clinical decision.

Results: 287 infants were admitted during the study period; 145 (51%) of these met the inclusion criteria and of these, 57 (39%) received joint assessments. The inter-rater reliability was high, (kappa(κ) 0.822). Sensitivity and specificity were 92% and 96% respectively.

Conclusions: District hospital nurses, using the TRY-CPAP algorithm, reliably identify babies that might benefit from bCPAP and thus improve its effective implementation.

INTRODUCTION:

Globally 2.7 million newborns die each year in the first 28 days of life comprising 45% of under-5 mortality (U5MR) (1). More than a third of these deaths (36%, representing 1 million babies) are attributable to complications of prematurity (2) and one of the most important causes is respiratory distress syndrome (RDS), resulting from surfactant deficiency (3). In well-resourced settings RDS is managed with ventilatory support – either in the non-invasive form of continuous positive airway pressure (CPAP), or as invasive mechanical ventilation with surfactant replacement therapy. These are costly interventions requiring a high level of newborn clinical skill to implement.

In response to this, bubble CPAP (bCPAP) has been developed as an affordable, practical, non-invasive mode of delivering CPAP. bCPAP provides respiratory support through a closed system, via nasal prongs to a spontaneously breathing infant. By delivering blended oxygen and air, at variable pressures and flow rates, it can deliver adjustable levels of oxygen (Fraction of inspired oxygen (F_{iO_2}) typically starting at 70%) (4) keeping the lungs expanded and reducing the work of breathing. The *Pumani* CPAP system (fig 1.) (*Pumani* meaning breathe in Chichewa) was developed for poor-resourced settings by clinicians and bioengineers at the University of Malawi, College of Medicine, and Rice University, Texas (5). It costs much less than any currently available commercial CPAP machine, uses a similar patient interface and delivers airflows and pressures similar to bCPAP devices at Texas Children's Hospital (6). (fig.1).

Two non-randomised studies in South Africa showed bCPAP to be associated with improved survival (7,8). In Malawi, studies found *Pumani* CPAP to be efficacious (9) and cost-effective (5). However bCPAP is challenging to implement correctly; health care workers (HCWs) are inexperienced and untrained in when and how to use it (10). Previously, the standard of care for initiating CPAP was simply to apply it when the health care workers felt that the baby had signs of significant respiratory

distress. However, during the roll-out of bCPAP to over 2000 patients in 28 facilities in Malawi it emerged that bCPAP was being given to floppy babies with severe birth asphyxia - hypoxic ischaemic encephalopathy (HIE), who may have respiratory distress caused by neurological damage rather than prematurity and surfactant deficiency. In such settings these babies are unlikely to benefit from bCPAP therapy, having a 75% combined probability of severe handicap or death (11) and often die in the first few days of life. A robust implementation strategy is therefore important to minimise opportunity costs, maximize resource allocation cost-effectiveness, and avoid the misconception that deaths could be attributed to bCPAP.

Other clinical-decision support tools for guiding bCPAP implementation in low resource settings are available, namely the Silverman Anderson Respiratory Severity Score (RSS) (12) and the Downes RDS score (13). These are both 5-part scoring systems that only consider one factor (respiratory effort). Short clinical scoring systems and algorithms have been shown to be effective in resource-poor settings, for example the highly successful three part Blantyre Coma Score (a simplified version of the paediatric Glasgow coma score) (14). Furthermore, neither the RSS nor the Downes RDS algorithms address the concern around bCPAP being applied inappropriately to newborns with HIE. The TRY-CPAP algorithm (T: Is Tone good, R: Is Respiratory distress present and Y: Yes, Heart rate is above 100b/min) (fig 2) is designed to address these deficits. It has three main steps and includes a measure of potential HIE. (11). It aims to assist nursing staff in deciding on the appropriate application of bCPAP, rationalising the use of a limited number of bCPAP machines towards infants most likely to benefit, such as premature babies with RDS and away from babies with HIE.

The word TRY is a useful aid memoire to recall the main elements of the algorithm if the chart is not available, however the steps in the flowchart (fig 2) do not follow the order T, R, Y and weight cut-offs are also included. The algorithm is for use on any baby with signs of respiratory distress and/or a birth weight of less than 1.3 kg. This

is an estimate for gestation of 30 weeks and early bCPAP is recommended for gestations < 30 weeks because at this gestation the lungs are sufficiently immature to need CPAP regardless of the signs of respiratory distress (11).

The first step of the algorithm (fig 2) is weight over 1kg and is designed to select patients likely to survive. Survival at < 1kg, in the absence of surfactant and intensive care, is estimated to be low at around 10%(15,16). Hence, babies weighing less than 1kg are given supportive care including intra-nasal oxygen. The next step selects babies that do not need active resuscitation, i.e. those with a heart rate (HR) > 100b/min. If the HR is < 100b/min the baby may need resuscitation and bCPAP would not be appropriate until successfully resuscitated. If the baby has good tone, the TRY algorithm then helps the user decide between early bCPAP for babies <1.3kg and delayed bCPAP for larger infants. For larger babies; if respiratory rate (RR) is >60 and/or oxygen saturations (O₂ sats) are <90%, intra-nasal oxygen (IN-O₂) is commenced first. If O₂ sats are persistently <90% after 2 hours, they are initiated on bCPAP.

This algorithm has shown promising results. In a prospective validation study conducted in Queen Elizabeth Central Hospital, Blantyre, Malawi (a tertiary referral centre), TRY-CPAP was shown to have a high interrater reliability, $k=0.9$ 95% CI (0.86-0.95), when comparing TRY assessments between doctor and nurses/nursing students (11). However, this study did not represent the more typical Malawian health care setting of a district-hospital, where newborn care is delivered by non-physicians and predominantly by nursing staff.

AIM:

To evaluate the reliability, sensitivity and specificity of the TRY-CPAP algorithm in a nurse-led district hospital environment in Malawi.

METHODS:

Setting: The study was conducted in a 40-bed Special Care Baby Unit (SCBU) in Zomba district hospital, Malawi.

Inclusion criteria: All newly admitted babies with respiratory distress *and* babies with an admission weight of less than 1.3kg with/without respiratory distress. Weight was measured using weighing scales. Respiratory distress was defined as any of the following signs - nasal flare, tracheal tug, chest in-drawings, recessions, head bobbing or grunting.

Exclusion criteria: Any baby with a suspected congenital abnormality requiring surgical intervention such as a congenital abnormality of the respiratory system and any baby weighing more than 1.3kg without respiratory distress.

Data collection:

Demographic, admission and outcome information were collected and recorded prospectively in the notes of all medical admissions (eligible and non-eligible for TRY-CPAP). Any newly admitted babies meeting the inclusion criteria would undergo a triple assessment on admission or during the ward round immediately following admission, consisting of a) TRY-CPAP assessment by a nurse (TRY-HCW) b) TRY-CPAP assessment by the paediatrician blinded to the nurse's assessment (TRY-Paed) and c) clinical assessment by the same paediatrician without using the TRY-CPAP algorithm according to her overall subjective impression. (Fig.2 depicts the TRY-CPAP algorithm where RR and O₂ sats are key-components) The TRY-HCW assessment was recorded on a proforma at the back of the patient's notes. TRY-Paed was done immediately following the nurse's assessment and documented in the patient's medical record. Then after a full assessment of the baby, the paediatrician would document her final clinical decision regarding CPAP.

Diagnoses were made according to the paediatrician's previous neonatal experience, local guidelines (Care of the Infant Newborn (COIN)(17)) and diagnostics available

(full blood count and gram stain but no blood culture, C reactive protein or x-ray). Please see appendix for summary of diagnostic definitions. Ballard scores were conducted as often as possible on any baby weighing less than 2500g or babies who appeared clinically premature irrespective of weight, but was done after the TRY triple assessment so as not to affect the initial decision on whether to instigate CPAP.

Training:

New nurses and students had not been trained to use TRY-CPAP and so weekly interactive teaching sessions were conducted by the visiting paediatrician prior to and during the study period. These sessions included practicing use of TRY in four theoretical case vignettes and real-time supervision on one real baby by the paediatrician. The TRY-algorithm (fig 2) was attached to the bCPAP machines and copies were printed for the clinical notes.

Time-scale: Data were collected from Monday to Friday between the hours of 09.00 and 17.00, from 27 April 2015 to 21 June 2015.

Sample: Convenience sampling was used to achieve as many TRY assessments as possible within the available time-period. Patients admitted overnight, on the weekend, or who died before the next weekday ward round were not included.

Ethical considerations: Ethical approval was obtained from both the Malawi College of Medicine and University College of London ethics committees (UK). Verbal consent was obtained from mothers to start CPAP where it was necessary but not for carrying out the TRY assessment as this did not affect the final clinical decision whether to instigate CPAP.

Analysis:

The number of eligible infants was compared to the total medical admissions (eligible and non-eligible) using univariate analyses. A score of 1-4 was obtained for each TRY-CPAP assessment according to its outcome (Room Air (RA) = 1, Intra-Nasal oxygen (IN-O₂) = 2, CPAP = 3, Early CPAP (ECPAP) = 4). The kappa statistic was calculated to measure interrater reliability between the TRY-CPAP scores of the

HCWs and the paediatrician. To facilitate sensitivity and specificity calculations, using the paediatrician's final decision as the gold standard, TRY-CPAP scores were made dichotomous, and recorded as 'CPAP' or 'no CPAP' with a score of 3 or 4 indicating the 'CPAP' group and 1 or 2 the 'no CPAP' group.

RESULTS:

There were 301 medical, neonatal admissions during the study period. 14 patient records were missing leaving 287 patient records for analysis: of these 145 (51%) patients were eligible for the TRY algorithm: 60 (41%) eligible babies had a TRY proforma completed by a HCW, of whom 57 (39%) also had a proforma completed and clinical assessment made by the paediatrician. (Fig 3). Cadres of HCW participating included matron, registered nurse mid-wife, nurse-midwife technician (NMT), student nurses and student clinical officers.

Table 1 shows characteristics of the whole sample (287 patients) and the sub-sample of 57 patients who had triple assessments using the TRY algorithm. In accordance with the eligibility criteria, babies who had the TRY algorithm were of lower weight and earlier estimated gestational age, had lower oxygen saturations and more prematurity with RDS. The TRY subgroup had slightly lower admission temperatures (a difference in medians of 0.6 degrees centigrade ($p=0.024$)) in accordance with lower weights and earlier gestations. Otherwise, there were no significant differences. Of these 57, mortality was 42% (24/57) and the primary diagnoses (presumed causes of death) were prematurity and RDA (13/24 = 54%), Severe birth asphyxia (8/24 = 33%), prematurity (2/24 = 8%) and meconium aspiration (1/24 = 4%). No significant complications of CPAP, such as pneumothorax, were suspected clinically when the visiting paediatrician was present. Chest x-ray was not available.

Of the 57 patients who had a TRY algorithm completed by both a HCW and a

paediatrician, 51 received the same outcome score. (Table 2) Of the six who differed the HCW overestimated the need for respiratory support for five and underestimated it in one. A Kappa statistic of 0.82 ($p < 0.001$), 95% confidence interval (CI) (0.69-0.95) was calculated as a measure of interrater reliability between the four possible outcomes of TRY in HCWs and the paediatrician.

With the paediatrician's final decision as the reference standard and using dichotomous outcomes of 'CPAP' or 'no CPAP', the sensitivity of TRY was 92.6% (95% CI 76.6-97.9%) and the specificity was 96.7% (CI 76-97%). The area under the receiver operator curve (AUC) or c-statistic was 0.946. The length of bCPAP treatment in the study ranged from 1-9 days, (mean 3.8, median 3.0 days).

DISCUSSION:

This is the only study in a resource-poor, district level hospital where newborn care is delivered predominantly by nursing staff, to assess the interrater reliability and validity of a clinical decision tool to aid the appropriate administration of bCPAP. We showed high interrater reliability, sensitivity and specificity of the TRY-bCPAP algorithm supporting the results of the previous study (Hundalani et al 2015). Key differences here are the lower resource district-level setting, only nursing staff of different cadres are included and babies of all weights are included to truly assess the whole algorithm.

A high kappa statistic (> 0.8) suggests that the TRY algorithm gives reproducible results with an 'outstanding measure of agreement' (18). A slightly higher kappa value in the previous QECH Blantyre study, at 0.9 for nurses and 0.97 for paediatricians in training (11) is expected as nursing expertise in a district level facility may not be equivalent to that in a tertiary centre. Nurses with less training and supervision may differ in their assessment, especially of tone and respiratory distress

and here a wide range of HCW cadres filled in the algorithm. A kappa of 0.82 implies that TRY can yield reproducible results even for different cadres of HCW. Total numbers were too small to disaggregate by cadre.

In some cases, parts of the algorithm were not adhered to, for example HR of <100b/min was disregarded by both HCW and paediatrician in an older baby from the community due to extreme work of breathing and both used clinical judgment to continue down the algorithm. Hence the 'HR above 100bpm' step of the algorithm is primarily to identify newborns that might need resuscitation straight after delivery, rather than older babies. The paediatrician's final decision for this patient was 'early CPAP' despite a weight of 4.0kg (well above the 1.3kg threshold in TRY). These 'over-rulings' illustrate how the algorithm may be more useful for newborns rather than those admitted from the community.

In this study, a HCW allocated bCPAP to a baby with severe birth asphyxia in only one case evidencing its success in directing bCPAP away from floppy babies with HIE. It is possible the HCW did not appreciate how floppy the baby was and instead focused on the low oxygen saturations of 80%, and signs of respiratory distress. There may have been more than one diagnosis in this case, such as mild birth asphyxia and meconium aspiration. This baby was kept comfortable on oxygen and died after three days due to hypoxic brain injury. Giving this baby bCPAP, may have supported the breathing temporarily but would not have improved his brain injury, or survival.

The sensitivity of TRY-CPAP of 92% was higher than in QECH, Blantyre (83%), while its specificity in Zomba (97%) was slightly lower than in Blantyre (100%) (11). A higher sensitivity (ie less false negatives) in a district hospital setting is encouraging and suggests the algorithm is safe, 'missing' less babies in lower resourced facilities.

Indeed, if the algorithm is slightly less specific i.e. more likely to allocate CPAP unnecessarily, this is preferable to missing babies who need CPAP. A greater sensitivity in this study may also be related to study design and reflect a learning effect in HCWs over time, as they observed the paediatrician using the algorithm. The paediatrician's TRY assessment was over-ruled by clinical judgement in 8 of 57 cases demonstrating that no algorithm can completely compensate for clinical knowledge and experience.

Comparisons of mortality in the TRY group versus the group who did not receive TRY are not reported because these comparisons are significantly confounded by sicker babies being prioritized for TRY due to low resources (as shown in table 1) and the numbers are not high enough to stratify for severity of illness. Also the final decision to use CPAP or not was made clinically by the paediatrician independently of TRY. A future bigger trial comparing TRY-CPAP with no TRY-CPAP, used only by HCWs without a visiting paediatrician, significantly powered to detect differences in mortality is recommended.

If the babies assessed in this study were representative of newborn admissions to other similar sized and resourced hospital we can estimate the need for bCPAP in similar facilities. Twenty-four (42%) of the 57 TRY algorithms completed by the paediatrician deemed babies appropriate for bCPAP. If TRY is completed in all babies with respiratory distress (51% of admissions), a facility admitting 150 infants per month, will need CPAP for 32 of these infants. If CPAP is on average continued for 3.8 days (as in this study), and given it is unlikely CPAP admissions will be spread evenly throughout the month, at least 5 machines may be needed per facility.

Other algorithms have been evaluated for their effect on mortality in low & middle-income countries. For example the Integrated Management of Neonatal and Childhood Illness (IMNCI) from the World Health Organisation (19) is a large scale

algorithm-based strategy, the scale up of which was thought to be a key factor in Malawi reaching MDG4 (Millennium Development Goal 4) (20). A recent Cochrane review (21) however, presented only moderate evidence that IMNCI 'may reduce infant mortality' and the strategy was difficult to evaluate because it employs multiple interventions across facility, primary care and community settings. The results of this TRY-CPAP evaluation (and the previous QECH study) add to the evidence that clinical algorithms are useful in facility settings. This is important for Malawi because 89% of women now give birth in a health care facility (22).

LIMITATIONS:

The main limitation of the study was that it was impossible for the paediatrician to be blinded to her own TRY assessment. Although she was blinded to the nurses TRY outcome, completion of the algorithm herself in the second part of the triple assessment may have biased her final decision whether to give CPAP. Unfortunately, there were no local paediatricians present to overcome this limitation. The convenience sampling method was heavily influenced by time pressures and staff availability (See Fig 3 Box: 'Reasons for TRY not done'), hence sicker babies were prioritised for TRY. The paediatrician was present throughout the study hence a follow-up study would be needed to assess how the algorithm is used in the absence of a clinician and could in addition examine intra-rater reliability and compare different cadres of HCWs to guide implementation.

The TRY algorithm itself, has limitations, for example it relies on weight as an estimate for gestation. It assumes that all babies with HIE are floppy, whereas they may be hypertonic. Some premature infants without HIE and weighing more than 1.3kg may be hypotonic due to causes such as sepsis, hypoglycaemia or prematurity itself. However, in this setting there are many more markedly hypotonic newborns with severe HIE who do not survive. Respiratory support in these cases would be futile, and when CPAP equipment and trained staff are limited, the support given to

such an infant may be at the expense of another infant who would benefit from CPAP support. One solution would be to adjust this step in the algorithm to say 'marked hypotonia due to asphyxia' rather than simply 'hypotonia'. A footnote could also be added to the algorithm to state that 'an experienced clinician may decide to use CPAP in circumstances beyond the scope of the TRY algorithm'.

NEXT STEPS:

Low-cost CPAP has been shown to be cost-effective and efficacious in improving outcomes for neonates in Malawi. Our study has shown the TRY-CPAP algorithm can help HCWs decide who would benefit from CPAP in a district hospital setting. However new technologies should only be introduced in a unit after careful assessment of the facility, equipment, work load and staffing numbers. A simple check list of minimum requirements in terms of staff, power supply, space and other equipment, should be drawn up, so that CPAP and TRY are used appropriately and in the right setting. CPAP should only be added when there is good evidence that basic neonatal nursing needs are already being provided and CPAP will not overstretch the unit's capabilities to deliver good care. Maynard et al have provided a list of technologies that are essential to newborn care and advise how to deliver them(23).

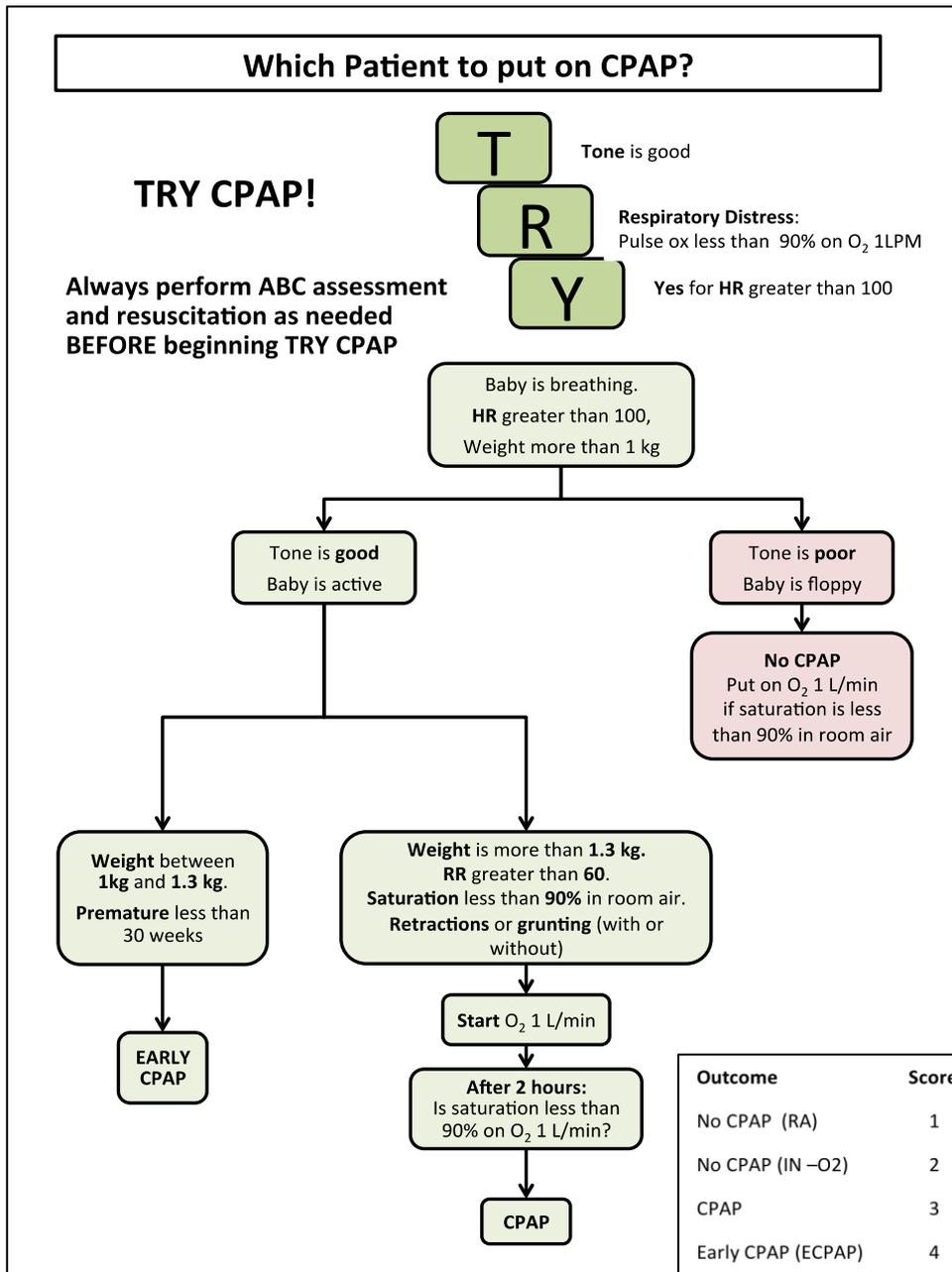
CONCLUSION:

This study demonstrates that the TRY-CPAP algorithm has a good interrater reliability, sensitivity and specificity between HCW and paediatrician. The TRY-CPAP algorithm was helpful, in guiding HCWs in the safe and appropriate application of low-cost bubble CPAP in a district hospital setting where usually physicians are absent and care is nurse-led. This evaluation supports the use of clinical algorithms for improving the quality of newborn care in low-resource settings and suggests that more staff are needed to support implementation of new technologies.

Figure 1. Pumani CPAP (Image from 3rd Stone Design, 2016)(24)

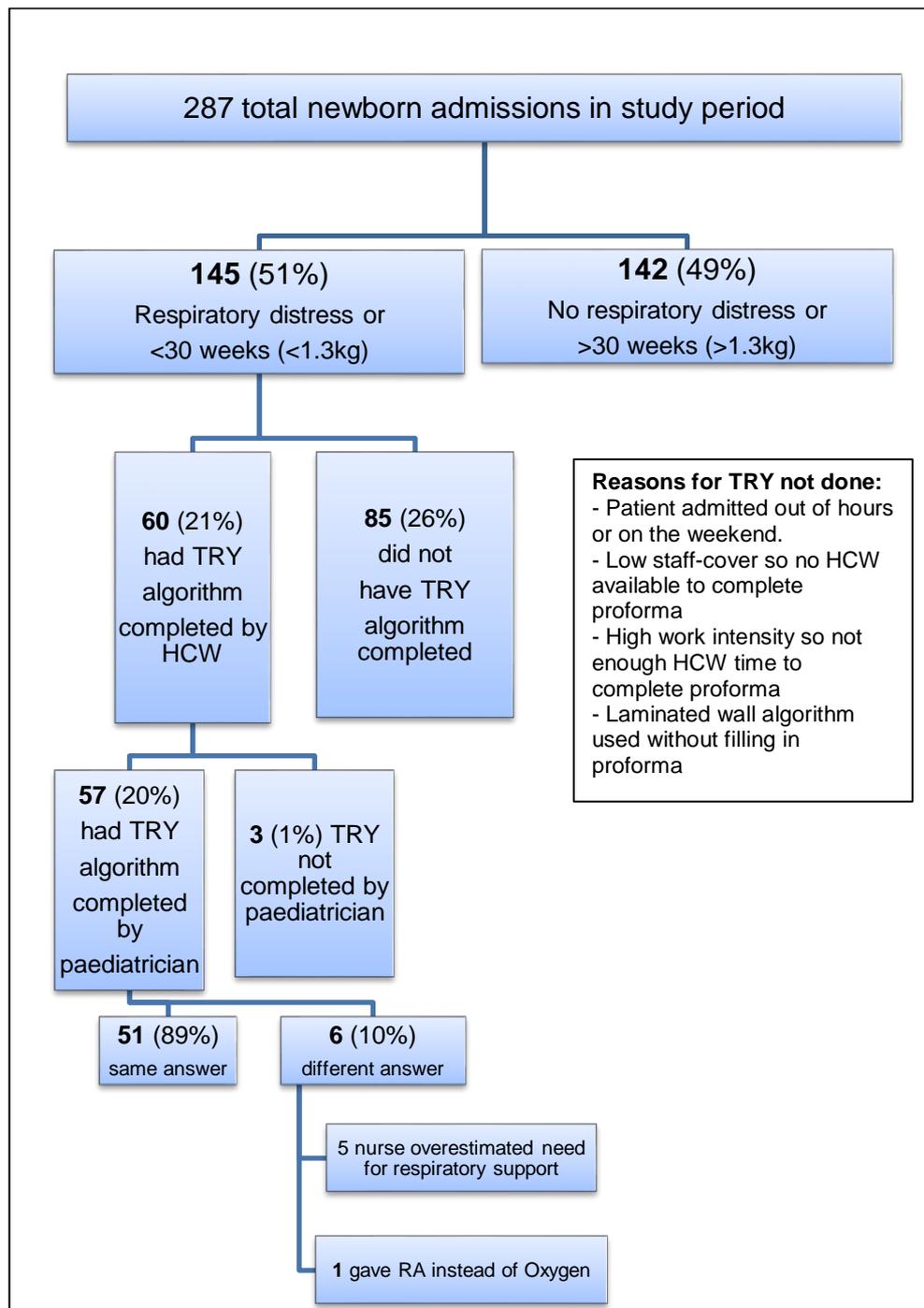


Figure 2. TRY-CPAP algorithm



CPAP = continuous positive airways pressure, HR = Heart Rate, RR = Respiratory rate O₂ = oxygen, RA = Room air, IN-O₂ = Intra-Nasal oxygen

Figure 3: Overview of participants



HCW = Health Care Worker, RA = Room Air

Table 1. Population characteristics					
		Whole Sample	Had TRY algorithm	p-value	
Number of study participants		287	57		
Sex (n(%))	Male	140 (48)	32 (56)	0.307	
	Female	146 (51)	24 (42)		
	Unrecorded	1 (1)	1 (2)		
Mean Birth weight (g)		2418	2127	0.047	
Birth weight (n(%))	<1300g	19 (7)	13 (23)	<0.001	
	>=1300g	253 (88)	40 (70)		
	<2500g (LBW)	127 (45)	33 (58)	0.038	
	>2500g	145 (50)	20 (35)		
Unrecorded		15 (5)	4 (7)		
Mean Gestation at birth (weeks)		38	36	0.006	
Gestation at birth (n(%))	<=30 weeks	8 (3)	5 (9)	0.030	
	>30 weeks	279 (97)	52 (91)		
Age (n(%))	Newborns <=48hrs	227 (7)	46 (81)	0.859	
	Older babies	60 (21)	11 (19)		
Weight for gestation category (n(%))	SGA	76 (27)	18 (32)	0.733	
	AGA	161 (56)	28 (49)		
	LGA	13 (4)	2 (3.5)		
	Unknown	37 (13)	9 (16)		
Admitted from (n(%))	Zomba District Hospital	167 (58)	31 (54)	0.3	
	Outside health unit	101 (35)	22 (38)		
	Unknown	19 (6)	4 (7)		
Singleton/twin/triplet (n(%))	Singletons	237 (83)	49 (86)	0.662	
	Twins	47 (16)	8 (14)		
	Triplets	3 (1)	0 (0)		
HIV status (n(%))	Exposed	32 (11)	8 (14)	0.612	
	Unexposed	146 (51)	31 (54)		
	Unknown	109 (38)	18 (32)		
Vital signs	(Mean(SD))	Admission HR (beats/minute)	132 (28)	138 (33)	0.238
		Admission RR (breaths/minute)	57 (16)	56 (18.6)	0.746
		Admission oxygen saturations (%)	89 (13)	79 (16)	<0.001
	(Median(IQR))	Admission Temperature (deg C)	36.1 (1.9)	35.5 (2.65)	0.024
Diagnosis (n(%))	Neonatal Sepsis	53 (18)	5 (9)	0.111	
	Birth Asphyxia (mild/moderate/severe)	46 (16)	11 (20)	0.73	
	LBW	44 (15)	3 (5)	0.07	
	Prematurity with RDS	40 (14)	19 (33)	<0.001	
	Pneumonia/Bronchiolitis	32 (12)	9 (16)	0.445	
	Prematurity only	29 (10)	4 (7)	0.614	
	Meconium aspiration	13 (5)	4 (7)	0.648	
	Transient Tachypnoea Newborn	5 (2)	0 (0)		
	Skin infections	5 (2)	0 (0)		
	Other	10 (3)	2 (3)	0.7	
	Well baby	9 (3)	0 (0)		
Outcome (n(%))	Absconded	8 (3)	1 (2)	<0.001	
	Discharged alive	233 (81)	32 (56)		
	Neonatal death	46 (16)	24 (42)		

ELBW = Extremely low birth weight, VLBW = Very low birth weight, LBW = Low birth weight, SGA = small for gestational age, AGA = appropriate for gestational age, LGA = Large for gestational age, HIV = Human immunodeficiency virus. HR = heart rate, RR = respiratory rate, Sats = Oxygen saturations, RDS = respiratory distress syndrome. For diagnosis Chi² tests were done for each separate diagnosis because there were many categories, Chi² could not be done where there was 0 in one group. Statistically significant p-values are highlighted in bold

Table 2: Cross-tabulation of Paediatrician & HCW TRY-CPAP Scores						
		HCW TRY-CPAP score				Total
		1: RA	2: IN-O ₂	3: CPAP	4: ECPAP	
Paediatrician TRY-CPAP score	1: RA	0	1	0	0	0
	2: IN-O ₂	1	29	2	1	33
	3: CPAP	0	0	15	1	17
	4: ECPAP	0	0	0	7	7
Total		2	28	18	9	57

HCW = Health Care Worker, TRY-CPAP = Tone, Respiratory distress, Yes HR > 100 – Continuous Positive Airways Pressure, RA = Room Air, IN-O₂ = intranasal oxygen, ECPAP = Early CPAP

REFERENCES

1. Liu L, Oza S, Hogan D, Perin J, Rudan I, Lawn JE, et al. Global , regional , and national causes of child mortality in 2000 – 13 , with projections to inform post-2015 priorities : an updated systematic analysis. *Lancet*. 2015;385(9966):430–40.
2. Lawn JE, Blencowe H, Oza S, You D, Lee ACC, Waiswa P, et al. Every newborn: Progress, priorities, and potential beyond survival. Vol. 384, *The Lancet*. 2014. p. 189–205.
3. Ho JJ, Subramaniam P, Henderson-Smart DJ, Davis PG. Continuous distending airway pressure for respiratory distress syndrome in preterm infants. *Cochrane Database Syst Rev*. 2000;(7):CD002271.
4. Hadley Health Technologies. Pumani bCPAP User Manual and Repair Manual [Internet]. [cited 2017 Nov 15]. p. 5 (Appendix A). Available from: <https://www.viaglobalhealth.com/wp-content/uploads/2016/01/3-378-Pumani-CPAP-User-Manual-H.pdf>
5. Chen A, Deshmukh AA, Richards-Kortum R, Molyneux E, Kawaza K, Cantor SB. Cost-effectiveness analysis of a low-cost bubble CPAP device in providing ventilatory support for neonates in Malawi, a preliminary report. *BMC Pediatr* [Internet]. 2014;14(1):288. Available from: <http://www.pubmedcentral.nih.gov/articlerender.fcgi?artid=4247223&tool=pmcentrez&rendertype=abstract>
6. Brown J, Machen H, Kawaza K, Mwanza Z, Iniguez S, Lang H, et al. A High-Value, Low-Cost Bubble Continuous Positive Airway Pressure System for Low-Resource Settings: Technical Assessment and Initial Case Reports. *PLoS One*. 2013;8(1).
7. Ballot DE, Chirwa TF, Cooper PA. Determinants of survival in very low birth weight neonates in a public sector hospital in Johannesburg. *BMC Pediatr* [Internet]. 2010;10:30. Available from: <http://www.biomedcentral.com/1471-2431/10/30>
8. Pieper C et al. Is nCPAP of value in extreme preterms with no access to Neonatal intensive care. *J Trop Pediatr*. 2003;49:148–52.
9. Kawaza K, Machen HE, Brown J, Mwanza Z, Iniguez S, Gest A, et al. Efficacy of a low-cost bubble CPAP system in treatment of respiratory distress in a neonatal ward in Malawi. *PLoS One*. 2014;9(1).
10. van den Heuvel M, Blencowe H, Mittermayer K, Rylance S, Couperus A, Heikens GT, et al. Introduction of bubble CPAP in a teaching hospital in Malawi. *Ann Trop Paediatr* [Internet]. 2011;31(1):59–65. Available from: <http://www.ncbi.nlm.nih.gov/pubmed/21262111>
11. Hundalani SG, Richards-Kortum R, Oden M, Kawaza K, Gest A, Molyneux E. Development and validation of a simple algorithm for initiation of CPAP in neonates with respiratory distress in Malawi. *Arch Dis Child - Fetal Neonatal Ed* [Internet]. 2015;100(4):F332–6. Available from: <http://fn.bmj.com/content/100/4/F332.abstract>
12. McAdams RM, Hedstrom AB, DiBlasi RM, Mant JE, Nyonyintono J, Otai CD, et al. Implementation of Bubble CPAP in a Rural Ugandan Neonatal ICU. *Respir Care*. 2015;60(3):437–45.
13. Rusmawati A, Haksari EL, Naning R. Downes score as a clinical assessment for hypoxemia in neonates with respiratory distress [Internet]. Vol. 48, 342±*Paediatr Indones*. 2008. Available from: <http://paediatricaindonesiana.org/pdf/48-6-5.pdf>
14. Taylor TE. Caring for children with cerebral malaria: insights gleaned from 20 years on a research ward in Malawi. *Trans R Soc Trop Med Hyg*. 2009;103(1 SUPPL.).
15. Rylance S, Ward J. Early mortality of very low-birthweight infants at Queen Elizabeth Central Hospital, Malawi. *Paediatr Int Child Health*. 2013;
16. Ahlsén AK, Spong E, Kafumba N, Kamwendo F, Wolff K. Born too small: who survives in the public hospitals in Lilongwe, Malawi? *Arch Dis Child - Fetal Neonatal Ed*. 2015;
17. COIN Course T. Care of the infant and newborn in Malawi The COIN Course Participants Manual. [cited 2017 Dec 3]; Available from: http://cms.medcol.mw/cms_uploaded_resources/41905_12.pdf
18. Landis J, Koch G. The measurement of observer agreement for categorical data. *Biometrics*. 1977;33:159–74.
19. WHO. Integrated management of childhood illness, Maternal, Newborn, Child and

- adolescent health. [Internet]. 2016. Available from:
http://www.who.int/maternal_child_adolescent/topics/child/nutrition/breastfeeding/en/
20. Kanyuka M, Ndawala J, Mleme T, Chisesa L, Makwemba M, Amouzou A, et al. Malawi and Millennium Development Goal 4: A Countdown to 2015 country case study. *Lancet Glob Heal*. 2016;4(3):e201–14.
 21. Gera T, Shah D, Garner P, Richardson M, HS S. Integrated management of childhood illness (IMCI) strategy for children under five. *Cochrane Database Syst Rev*. 2016;(6):1–61.
 22. Malawi Ministry of Health. Malawi Every Newborn Action Plan [Internet]. Malawi MOH; 2015. Available from: www.who.int/pmnch/media/events/2015/malawi_enap.pdf
 23. Maynard KR, Causey L, Kawaza K, Dube Q, Lufesi N, Maria Oden Z, et al. New technologies for essential newborn care in under-resourced areas: what is needed and how to deliver it. *Paediatr Int Child Health* [Internet]. 2015;35(3):192–205. Available from: <http://www.tandfonline.com/doi/full/10.1179/2046905515Y.0000000034>
 24. 3rd Stone Design. Pumani CPAP 3rd stone design [Internet]. 2016. Available from: <http://www.3rdstonedesign.com/work/pumanibcpap>

What is already known on this topic?

- 1) The TRY-CPAP algorithm has previously been shown to be reliable, sensitive and specific when used by nurses in a teaching hospital, using a neonatologist's decision as the standard.
- 2) This was a single-centre study conducted in Malawi's teaching hospital, so does not represent a typical Malawian health facility.
- 3) Other algorithms, previously evaluated elsewhere, only consider respiratory effort, and do not tackle the problem of bCPAP being applied inappropriately to newborns with HIE.

What does this study add?

- 1) TRY was developed to support the use of bCPAP in district hospitals and smaller facilities, where there may not be any qualified doctors.
- 2) This study assessed TRY in a district Hospital where there are few qualified doctors and no paediatricians. Results showed the algorithm to have good reliability, sensitivity and specificity.
- 3) This study therefore confirms the value of TRY in helping nurses to correctly implement CPAP in the absence of specialist support.