

#### TITLE

Admission profile and discharge outcomes for infants aged less than 6 months admitted to inpatient therapeutic care in 10 countries. A secondary data analysis.

#### **AUTHORS AND AFFILIATIONS**

- Carlos S. Grijalva-Eternod. UCL Institute for Global Health, 30 Guilford Street, London, WC1N 1EH, UK. c.eternod@ucl.ac.uk
- Marko Kerac. Department of Population Health, London School of Hygiene & Tropical Medicine, London, Keppel Street, London, WC1E 7HT, UK. Leonard Cheshire Leonard Cheshire Disability and Inclusive Development Centre, Department of Epidemiology & Public Health, 1-19 Torrington Place, London WC1E 6BT, UK. <u>marko.kerac@lshtm.ac.uk</u>

Marie McGrath. ENN, 32 Leopold Street, Oxford, OX4 1TW, UK. marie@ennonline.net

- Caroline Wilkinson. Action Contre la Faim (at the time of data submission) 14/16 Boulevard Douaumont - CS 80060 75854 Paris Cedex 17, France. United Nations High Commissioner for Refugees, Geneva, Switzerland. wilkinso@unhcr.org
- June C. Hirsch. Action Contre la Faim 1416 Boulevard Douaumont CS 80060 75854 Paris Cedex 17, France. jvhirsch@gmail.com
- Pascale Delchevalerie. Medical Department, Médecins Sans Frontières. 46, Rue de l'Arbre Bénit, 1050 Bruxelles. pascale.delchevalerie@brussels.msf.org
- <u>Andrew J. Seal</u>. UCL Institute for Global Health, 30 Guilford Street, London, WC1N 1EH, UK. <u>a.seal@ucl.ac.uk</u>

#### SHORT RUNNING TITLE

Management of acute malnutrition in infants.

#### SOURCE OF FUNDING

The development of this paper was made possible by the generous support of the American people through the United States Agency for International Development (USAID) and through the support of Irish Aid, in grants to ENN. The Global Nutrition Cluster (UNICEF) funded the original review of national guidelines that was updated for this manuscript, undertaken as part of the ENN-led MAMI Project. The contents do not necessarily reflect the views of the funding agencies.

#### **CONFLICT OF INTEREST**

The authors declare that they have no conflict of interest

#### CONTRIBUTIONS

AJS, MK, MM conceived the study. CW, JCH and PD contributed data. CSGE analysed and interpreted the data, and wrote the initial draft of the manuscript. All authors contributed to the manuscript revisions. All authors read and approved the final manuscript.

Word count abstract: 245 words Word count main text: 3,619 words Number of references: 28 references Number of tables: Four tables Number of figures: One figure Supplementary tables & figures: One table and two figures

# Admission profile and discharge outcomes for infants aged less than 6 months admitted to inpatient therapeutic care in 10 countries. A secondary data analysis.

## ABSTRACT

Evidence on the management of acute malnutrition in infants aged less than 6 months (infants <6mo) is scarce. To understand outcomes using current protocols, we analysed a sample of 24,045 children aged 0-60 months from 21 datasets of inpatient therapeutic care programmes in 10 countries. We compared the proportion of admissions, the anthropometric profile at admission, and the discharge outcomes between infants <6mo and children aged 6-60 months (older children).

Infants <6mo accounted for 12% of admissions. The quality of anthropometric data at admission was more problematic in infants <6mo than in older children with a greater proportion of missing data (a 6.9 percentage points difference for length values, 95%CI: 6.0; 7.9, p<0.01), anthropometric measures that could not be converted to indices (a 15.6 percentage points difference for weight-for-length z-score values, 95%CI: 14.3; 16.9, p<0.01), and anthropometric indices that were flagged as outliers (a 2.7 percentage points difference for any anthropometric index being flagged as an outlier, 95%CI: 1.7; 3.8, p<0.01). A high proportion of both infants <6mo and older children were discharged as recovered. Infants <6mo showed a greater risk of death during treatment (risk ratio 1.30, 95%CI: 1.09; 1.56, p<0.01).

Infants <6mo represent an important proportion of admissions to therapeutic feeding programmes and there are crucial challenges associated with their care. Systematic compilation and analysis of routine data for infants <6mo is necessary for monitoring programme performance and should be promoted as a tool to monitor the impact of new guidelines on care.

## **KEYWORDS**

Malnutrition, Infant and Child Nutrition, Management of Acute Malnutrition, Mortality, Anthropometry, Wasting.

- Admission profile and discharge outcomes for infants aged less than 6 months admitted
   to inpatient therapeutic care in 10 countries. A secondary data analysis.
- 3

#### 4 INTRODUCTION

Acute malnutrition is a serious global health concern (Black et al. 2013). Global estimates 5 indicate that wasting, a type of acute malnutrition characterized by acute mass loss (WHO 6 1995), affects 50 million children aged <5 years and accounts for 11.5% of their total deaths 7 8 (UNICEF et al. 2015;Black et al. 2013). Severe wasting affects 16 million children and accounts for 7.8% of their total deaths (UNICEF et al. 2015;Black et al. 2013). Moreover, it is 9 10 also estimated that wasting affects 8.5 million infants aged less than 6 months (henceforth 11 referred to as infants <6mo) (Kerac et al. 2011). Beyond its short-term impact on survival and health, this wasting burden has long-lasting consequences for both individuals and societies 12 (Victora et al. 2008). 13

Despite these high global burdens, infants <6mo were only recently included in the new World Health Organisation (WHO) guidelines for the management of severe acute malnutrition (SAM) (WHO 2013;WHO & UNICEF 2009). Although, inclusion of infants <6mo in these guidelines represents an important development, there is also a recognised need for developing the evidence base in order to improve care in this age group (Angood et al. 2015).

Describing the profile and outcomes associated with the management of acute malnutrition in infants <6mo is central for expanding our understanding about the effectiveness of current care strategies and setting the baseline evidence to help guide improved future care. This study, which preceded the new WHO 2013 guidelines (WHO 2013), aimed at providing evidence on infants <6mo receiving inpatient therapeutic care to determine what is their proportion among children aged 0-60 months, what is their anthropometric profile at admission, and what their outcomes are at discharge.

#### 26 PARTICIPANTS AND METHODS

27 *Ethics* 

This study carried out a secondary analysis of routinely collected and fully anonymised programme data. The analysis of data from programmes in which there is no intervention trial of planned change in procedures is widely classified as audit or service evaluation by research ethics committees. Consequently, no ethical approval was required.

32

#### 33 Field datasets

34 An appeal for datasets containing individual-level programme data on acute malnutrition care of infants <6mo was put out from May to December 2008. We received a total of 30 datasets 35 from Action Contre la Faim (ACF) and one from Médecins Sans Frontières. Of these, only 23 36 37 datasets from ACF contained inpatient therapeutic care programme data from 25,195 children 38 aged 0-60 months from 34 field sites located in 12 countries. Table 1 provides details of the children in these datasets by country and the type of inpatient therapeutic programme care. The 39 40 majority of the individuals in our dataset (81.9%) were admitted into therapeutic feeding centres (TFC). We excluded the data from Afghanistan and Ethiopia (n=1,150) as their 41 programme data included only very young children with no older children for comparison. A 42 final sample of 24,045 children aged 0-60 months was used for analysis. 43

44

#### 45 *Data available*

Age, the presence of bilateral pitting oedema, and anthropometric data, namely weight, length or height, and mid-upper arm circumference (MUAC), were available for most children at admission. For most children, discharge outcomes were also available. Anthropometric data was also available at discharge but there was a large heterogeneity in the type and timing of data collected. Consequently, this analysis focused only on anthropometric and oedema data at
admission and outcomes at discharge.

52

#### 53 Data handling and data analysis

Data was manipulated and analysed in Stata software (Stata Statistical Software: release 14, 54 2015; StataCorp LP). We calculated the anthropometric indices weight-for-age, height/length-55 56 for-age and weight-for-height/length z-scores (WAZ, HAZ, WHZ, respectively) from weight, height or length, age and sex variables at admission, based on the 2006 WHO Growth Standards 57 58 (WHO Multicentre Growth Reference Study Group 2006) using the zscore06 command (Leroy 2011). Extreme z-score values are usually assumed to represent measurement or data entry 59 errors (WHO 1995). We flagged these extreme values as outliers using commonly applied 60 61 cleaning criteria (Crowe et al. 2014) as follows: Flag 1: WAZ <-4 or >4 z-scores from the observed mean; Flag 2: HAZ <-4 or >3 z-scores from the observed mean; and Flag 3: WHZ <-62 4 or >4 z-scores from the observed mean. Acute malnutrition, based on WHZ and/or the 63 64 presence of oedema, was classified as global (GAM; WHZ<-2 and/or oedema), moderate (MAM; WHZ $\leq$ -2 but  $\geq$ -3), and severe (SAM; WHZ $\leq$ -3 and/or oedema). Wasting, based on 65 WHZ among children without reported oedema, was classified as total (WHZ<-2), moderate 66 (WHZ<-2 but  $\geq$ -3), and severe (WHZ<-3). 67

Discharge outcomes were coded differently between and within datasets, differing primarily in
the terminology used and the manner in which they were abbreviated. Discharge codes were
grouped into the four Sphere discharge codes recovered, died, defaulted, and non-recovered
(The Sphere Project 2011); as well as admission error, or missing: if no discharge outcome data
was available (see Table S1).

To describe the burden for programmes providing therapeutic care for acute malnutrition to
infants <6mo, we calculated what proportion of programme admissions were within this age</li>

75 group. To assess the quality of the anthropometric data collected at admission for infants <6mo, we compared the proportion of missing values, the proportion of values that failed to convert 76 into anthropometric indices, and the proportion of anthropometric indices that were categorised 77 78 as outliers against their older counterparts. To assess the nutrition profile at admission for infants <6mo, we compared the proportion of GAM, MAM, SAM, oedema, and of total, 79 moderate and severe wasting against their older counterparts. Lastly, we compared the 80 proportions of discharge outcomes and performed a meta-analysis to assess the risk of death 81 82 during treatment between the different age groups. To test for the equality of means and 83 proportions we used the ztest and prtest commands, respectively. For meta-analysis we used the metan command. 84

#### 85 **RESULTS**

86 *Programme burden - Proportion of children aged <6 months* 

Our sample for analysis included 24,045 children aged 0-60 months who were receiving 87 therapeutic care for acute malnutrition (see Table 1). We observed that a large proportion of 88 these were young; i.e. 17,963 (75%) and 2,939 (12%) were aged 0-24 months and less than 6 89 months, respectively. Figure S1 shows the age frequency distribution of the sample, where one 90 91 can also observe rounding of age to the nearest half-year from 12 months of age onwards. Infants <6mo represented 16% of the sample of children aged 0-24 months. Regarding the type 92 93 of programme therapeutic care, infants <6mo accounted for 6%, 10%, 18% and 13% of the sample for Day Centre (DC), Home Treatment (HT), Stabilisation Centre (SC) and TFC 94 programmes, respectively. The proportion of boys was similar between the two groups, i.e. 95 50.3% and 50.6% for infants <6mo and 6-60 months, respectively. 96

97

#### 98 *Quality of data at admission*

99 Table 2 displays the difference in the quality of anthropometric data at admission for infants 100 <6mo and children aged 6-60 months. At admission, data on infants <6mo contained a 101 significantly greater proportion of missing values for length and for MUAC than their older 102 counterparts; but both age groups had a similar low proportion of missing values for weight 103 and for the presence of bilateral pitting oedema.

Secondly, the WHZ index could not be calculated for a significantly greater proportion of infants <6mo using the anthropometric data collected at admission. The main reason for this difference was that for most infants <6mo, for whom WHZ could not be calculated (467 out of 471), their length was lower than 45cm, the minimum reference value needed for calculating this index. The proportion of WAZ and HAZ indices that could not be calculated was very low for both groups. Lastly, there is a significantly greater proportion of anthropometric indices that were flagged as statistical outliers in infants <6mo compared to children aged 6-60 months. **Figure S2** provides a visual comparison of the difference in the availability of anthropometric data between infants <6mo and their older counterparts. After accounting for missing data, poor quality or out of range anthropometric data, only 74% of the sample of infants <6mo have anthropometric data that would allow for the assessment of wasting, as defined by WHZ, compared with 97% of their older counterparts.

117

### 118 Anthropometric and clinical profile at admission

Table 3 presents the nutrition profile data from the subsample of children aged 0-60 months 119 that had no missing weight or height/length data and their calculated WHZ values were not 120 121 flagged as outliers. Overall, the nutritional profile of infants <6mo was better compared to their older counterparts. Infants <6mo showed a significantly lower GAM proportion than older 122 children, of which a significantly larger and lower proportion were MAM and SAM, 123 respectively. In addition, infants <6mo presented with a significantly lower proportion of 124 bilateral pitting oedema. Similarly, after removing from the sample those reported to have 125 oedema, infants <6mo had a significantly lower proportion of wasting compared to their older 126 counterparts, of which a significantly larger and lower proportion were moderate and severe 127 wasting, respectively. Lastly, mean WHZ values were significantly greater for infants <6mo. 128

129

130 *Discharge outcomes* 

**Table 4** displays the discharge outcomes by age group. Overall, both age groups have a similar high proportion of children being discharged as recovered. However, we observed a significantly lower proportion of infants <6mo discharged as defaulted. Figure 1 presents a forest plot of the pooled risk ratio for death during treatment for infants <6mo against their</p>

- 135 older counterparts. Overall, the risk ratio for death was significantly greater for infants <6mo.
- 136 However, there was a high level of variation in the risk ratio between study sites (86.6%
- 137 variation in risk ratio attributable to heterogeneity; chi-squared = 67.0 p < 0.01).

#### 138 **DISCUSSION**

#### 139 *Main findings*

To our knowledge, this is the first analysis of programme information from a variety of countries and care programmes containing data on infants <6mo receiving therapeutic care for acute malnutrition. One of our main findings is infants <6mo represent an important proportion of the children receiving malnutrition care in the programmes run by international relief agencies.

Our analysis provides insights into some of the main challenges that malnutrition care 145 146 programmes face when assessing infants <6mo. We found that the collection of anthropometric data in infants <6mo is challenging as indicated by the greater proportion of missing data at 147 admission, particularly length. The MUAC data was also missing, far more than in older 148 149 children; however, this was not surprising as MUAC is not recommended as an admission 150 criterion for infants <6mo (Kerac et al. 2012). Furthermore, even when weight and length data were successfully collected, it was not possible to convert a large proportion of them into any 151 useful anthropometric index since WHZ cannot be calculated when length is <45cm. In 152 addition, when this index calculation was possible, a large proportion of the values were 153 observed to be extreme. 154

Furthermore, our study found that infants <6mo who are receiving therapeutic care for acute malnutrition presented a better nutritional profile at admission when compared with their older counterparts. Specifically, infants <6mo presented a lower proportion of oedema and had, on average, greater WHZ values at admission. These differences were manifested in the lower proportion of GAM and total wasting, as well as the proportion of SAM and severe wasting observed in infants <6mo.

Lastly, our analysis showed that infants <6mo have a similar proportion of recovered outcomes</li>
at discharge. However, infants <6mo had a higher risk ratio for death during treatment.</li>

163

#### 164 Programme burden

We have previously showed that despite the lack of focus on assessing the nutritional status in 165 this age group (Lopriore et al. 2007), acute malnutrition among infants <6mo is a public health 166 concern (Kerac et al. 2011); a prevalence that others have characterised as an underestimated 167 public health problem (Patwari et al. 2015). Our analysis contributes to this evidence by 168 169 showing that infants <6mo also account for an important proportion of children receiving inpatient therapeutic care. This burden of care is important given the weak evidence base on 170 171 which care for this age group is based, and their care often falls in the gap between neonatal care and the management of malnutrition for older children (Kerac et al. 2015). 172

It is not possible for us to assess the extent to which the disease burden observed in our sample 173 174 reflects the actual prevalence of acute malnutrition in the catchment areas of the therapeutic programme, as infants <6mo are not routinely included in prevalence surveys of acute 175 malnutrition. Recent evidence has shown that the proportion of infants <6mo suffering from 176 acute malnutrition compared to their older counterparts is greater in hospital settings than in 177 the wider community (Karunaratne et al. 2015). Furthermore, others and we have argued that 178 because it is commonly assumed that this age group is better protected from nutritional stress 179 than their older counterparts, the available estimates are likely to represent an underestimate of 180 its prevalence in both inpatient and community settings that provide malnutrition care. 181 182 However, evidence supporting the assumption of greater protection among infants <6mo exists (Pongou et al. 2006), making it difficult for us to extrapolate our findings to the wider 183 population. 184

185

#### 186 Assessing nutritional status of infants aged <6 months

How acute malnutrition among infants <6mo should be defined is, at present, a top priority 187 research question (Angood et al. 2015). This definition will determine who will receive 188 malnutrition care. The two anthropometric indicators commonly used for assessing SAM in 189 children aged 6-59 months are also being considered for infants <6mo (WHO 2013;WHO & 190 UNICEF 2009), namely low WHZ and low MUAC. Discussions about which of these 191 192 indicators is better suited to assess acute malnutrition in older children have focused almost exclusively on their predictive value for assessing a high risk of death (Walters et al. 2012), in 193 194 spite of the large body of evidence about the long-term consequences of the impaired development associated with acute malnutrition (Victora et al. 2008). 195

Recent evidence, relevant for infants <6mo, has shown that MUAC data, collected at the age 196 197 of routine vaccination, 6-14 weeks of age, predicts child survival at age 12 months better than WHZ data (Mwangome et al. 2012a). Furthermore, collection of MUAC data among infants 198 <6mo has also been shown to be more reliable and accurate than WHZ when collected by 199 trained community health workers, using hanging scales with a precision of 100g (Mwangome 200 et al. 2012b;Mwangome & Berkley 2014). Our study adds to this evidence by showing that for 201 inpatient therapeutic care programmes, obtaining reliable WHZ data in infants <6mo is 202 problematic because of problems arising at different steps, from collection of anthropometric 203 data through calculation of indices and cleaning of data. That a greater number of WHZ were 204 205 flagged for infants <6mo is also relevant from an epidemiological standpoint; and suggests that further work is necessary to better understand if the cleaning criteria originally envisioned to 206 be applied to older children should be applied to this younger age group. It is difficult to draw 207 208 any conclusion regarding the reliability of MUAC data collection in this analysis, as the data 209 was collected during a period when the use of MUAC measures in therapeutic care was not a 210 firmly established practice in older children; and it has never been recommended in infants211 <6mo.</li>

Despite the relative ease of MUAC data collection, compared to WHZ, and its strong 212 association with mortality risk, doubt remains as to how well it indicates acute weight loss in 213 infants  $\leq 6$  months in Ethiopia showed 214 that MUAC values in this very young population are weakly associated with body composition 215 216 (Grijalva-Eternod et al. 2015). MUAC variability among these infants reflects more the variability in length, independently of age and sex, and less the variability of tissue masses. 217 218 Conversely, WHZ variability seems to index nutritional status better as it more closely reflects variability in tissue masses. Given that these two indicators have a different relationship with 219 body composition data and mortality, it has been proposed that MUAC measurements among 220 221 infants <6mo might have a greater capacity to assess growth failure as opposed to an acute loss of tissue mass, for which WHZ might be better-suited (Grijalva-Eternod et al. 2015). Further 222 longitudinal evidence is needed to empirically test this proposal. Nonetheless, even if WHZ is 223 a better indicator of acute tissue mass loss, and MUAC a better indicator of mortality, the 224 challenge remains that collection of anthropometric data, like length, and calculation of indices, 225 like WHZ, among infants <6mo is highly problematic. 226

227

## 228 The nutrition profile of infants aged <6 months at admission to therapeutic care

At admission to therapeutic care, infants <6mo present a better anthropometric profile than their older counterparts do, even after accounting for oedema. To our knowledge, this is the first report of this difference. There is scarce literature to help us understand why oedema was significantly lower among infants <6mo; or why they seem to be admitted to therapeutic care at a less severe stage of malnutrition. In infants <6mo, oedema might be more difficult to diagnose; as in older children of whom most can stand, gravity might influence in narrowing

the location of the oedema to the limbs. Also, infants <6mo compartmentalise body water 235 differently than older children. Studies have shown that total body water, as a percentage of 236 body weight, and extracellular water, as a fraction of total body water, decrease rapidly during 237 the first 130 days of life, with extracellular water decreasing more rapidly (Fomon & Nelson 238 2002). It might be that clinically detectable oedema is more likely only after certain 239 developmental milestones have taken place, such as the decrease in the ratio of extracellular to 240 241 cellular water mentioned above. This idea is supported by the observation that the proportion of oedema among older children with SAM increases with age, peaking at three to five years 242 243 of age (Girma et al. 2013).

244

#### 245 *Care outcomes*

We are not the first to show that a high proportion of infants <6mo admitted to receive care for 246 acute malnutrition recover (Singh et al. 2014; Vygen et al. 2013). However, our findings adds 247 to this evidence. We showed that the proportions discharged as recovered are similar between 248 infants <6mo and older children, as well as the proportions discharged as non-recovered. We 249 also observed a lower proportion of infants <6mo being discharged as defaulted. However, this 250 findings may be because the proportion discharged as dead is higher in this age group 251 (borderline significant). To investigate this borderline fatality, we conducted a meta-analysis 252 of the date from different countries. This revealed that infants <6mo have a higher relative risk 253 254 of death; despite a better nutritional profile at admission.

The higher relative risk of death for infants <6mo observed in our study needs cautious interpretation given the high level of heterogeneity observed between the countries where the data was collected. It is not possible to disentangle whether the observed heterogeneity in our results reflects a different mortality risk among infants <6mo in these different settings, at comparable levels of anthropometrically defined malnutrition; or if this observed heterogeneity may be due to differences in the quality of therapeutic care provided to infants <6mo. Likewise,</li>
it is not possible to assess how much of the higher relative risk of death observed in infants
<6mo may be due to suboptimal care driven by the existence of inadequate care protocols, or</li>
an inadequate provision of care, or both, given the lack of international guidelines for the
management of malnutrition in infants <6mo.</li>

#### 266 *Limitations*

Our study has some limitations. First, most programmes were less likely to have actively sought 267 268 infants <6mo in the community compared to older children aged 6-59 months, and might not have recommended inpatient care for all cases of SAM identified in infants <6mo. This 269 potential bias may have resulted in an under-representation of malnourished infants <6mo, that 270 271 may have varied between contexts, but could not be quantified. Second, the absence of a clear anthropometric criterion for admission to therapeutic feeding of many infants <6mo suggests 272 that alternative criteria were also used. The alternative criteria might include a number of non-273 274 anthropometric criteria, such as clinical signs of infection, disability, feeding difficulties, and maternal factors; an assumption that is supported by a review of admission criteria used for this 275 age group (ENN & CIHD 2010). How much these additional criteria might help explain the 276 differences observed in the nutritional profile of these two age groups where infants seems to 277 be admitted to care at a less severe stage in malnutrition is unknown and could not be quantified 278 279 in our analysis. Lastly, all datasets used for this analysis originate from one international relief agency limiting the study capacity to extrapolate our findings to the other care providers. 280

Our study has also strengths. To our knowledge this is the largest multicentre analysis of inpatient therapeutic care data that includes data on infants <6mo. As such, this dataset allows for a more global understanding of differences in the management of acute malnutrition in these two groups. Likewise, given the paucity of the evidence base for the management of acute

<sup>265</sup> 

malnutrition in infants <6mo (Kerac et al. 2015), even after their inclusion in the WHO</li>
guidelines (WHO 2013), our analysis provides the best available comparisons at admission and
discharge between these two age groups.

288

289 *Conclusions* 

Infants <6mo represent an important proportion of admissions to therapeutic feeding 290 programmes for acute malnutrition. There are numerous challenges associated with their care: 291 292 anthropometric measurement; knowing which measures and signs of illness or poor feeding 293 are best to use for assessment; interpreting current programme outcomes and knowing to what extent the observed mortality is avoidable through better guidelines or better implemented 294 guidelines. Systematic compilation and analysis of routine data of infants <6mo is important 295 296 for monitoring programme performance and should be promoted as a tool to assess the impact of new guidelines on care. 297

#### **KEY MESSAGES**

- Infants aged less than 6 months account for an important proportion of patients that receive inpatient therapeutic care for acute malnutrition.
- Collection of infant's anthropometric data at admission to therapeutic care is problematic compared to that of their older counterparts (children aged 6-60 months). Data on infants had a greater proportion of missing anthropometric data, anthropometric data that could not be used to estimate nutrition indicators, and estimated nutrition indicators that were flagged as extreme and unlikely values.
- At admission to therapeutic care, infants aged less than 6 months presented with a better nutritional profile, including a lower proportion of oedema, global acute malnutrition and severe acute malnutrition compared to their older counterparts.
- The proportion of infants aged less than 6 months and older children discharged as recovered was similar. However, infants aged less than 6 months suffered a higher case fatality rate.
- Systematic compilation and analysis of routine data is an important tool for monitoring programme performance and should be promoted as a tool to monitor the impact of rolling out new guidelines on therapeutic care.

## REFERENCES

Angood, C., McGrath, M., Mehta, S., Mwangome, M., Lung'aho, M., Roberfroid, D. et al. (2015). Research priorities to improve the management of acute malnutrition in infants aged less than six months (MAMI). *PLoS Medicine*, 12, (4) e1001812.

Black, R.E., Victora, C.G., Walker, S.P., Bhutta, Z.A., Christian, P., de Onis, M. et al. (2013). Maternal and child undernutrition and overweight in low-income and middle-income countries. *Lancet*, 382, (9890) 427-451.

Crowe, S., Seal, A., Grijalva-Eternod, C., & Kerac, M. (2014). Effect of nutrition survey 'cleaning criteria' on estimates of malnutrition prevalence and disease burden: secondary data analysis. *PeerJ*, 2, e380.

ENN & CIHD, A. (2010), MAMI Project. Technical Review: Current evidence, policies, practices & programme outcomes. Available at: www.ennonline.net/mamitechnicalreview

Fomon, S.J., Nelson, S.E. (2002). Body composition of the male and female reference infants. *Annual Review of Nutrition*, 22, 1-17.

Girma, T., Kaestel, P., Mølgaard, C., Michaelsen, K.F., Hother A.L., Friis, H. (2013). Predictors of oedema among children hospitalized with severe acute malnutrition in Jimma University Hospital, Ethiopia: a cross sectional study. *BMC Pediatrics*, 13, 204.

Grijalva-Eternod, C.S., Wells, J.C., Girma, T., Kaestel, P., Admassu, B., Friis, H. et al. (2015). Midupper arm circumference and weight-for-length z scores have different associations with body composition: evidence from a cohort of Ethiopian infants. *American Journal of Clinical Nutrition*, 102, (3) 593-599.

Karunaratne, R., Lelijveld, N., Newberry, L., Munthali, C., Kumwenda, E., Lang, H. J. et al. (2015). Management of acute malnutrition in infants aged <6 months (MAMI) in Malawi: Prevalence and risk factors in an observational study. *Archive of Diseases in Childhood* 100[Suppl 3], A103.

Kerac, M., Blencowe, H., Grijalva-Eternod, C., McGrath, M., Shoham, J., Cole, T.J. et al. (2011). Prevalence of wasting among under 6-month-old infants in developing countries and implications of new case definitions using WHO growth standards: a secondary data analysis. *Archives of Diseases in Childhood*, 96, (11) 1008-1013.

Kerac, M., Mwangome, M., McGrath, M., Haider, R., & Berkley, J.A. (2015). Management of acute malnutrition in infants aged under 6 months (MAMI): current issues and future directions in policy and research. *Food and Nutrition Bulletin*, 36, (1 Suppl) S30-S34.

Kerac, M., Trehan, I., Weisz, A., Agapova, S., & Manary, M. (2012), Admission and discharge criteria for the management of severe acute malnutrition in infants aged under 6 months: An AGREE appraisal of national protocols and GRADE review of published literature. Available at:

http://www.who.int/nutrition/publications/guidelines/updates\_management\_SAM\_infantandc hildren\_review8.pdf?ua=1

Leroy, J. F. (2011). zscore06: Stata command for the calculation of anthropometric z-scores using the 2006 WHO child growth standards.

Lopriore, C., Dop, M.C., Solal-Celigny, A., & Lagnado, G. (2007). Excluding infants under 6 months of age from surveys: impact on prevalence of pre-school undernutrition. *Public Health Nutrition*, 10, (1) 79-87.

Mwangome, M.K. & Berkley, J.A. (2014). The reliability of weight-for-length/height Z scores in children. *Maternal and Child Nutrition*, 10, (4) 474-480.

Mwangome, M.K., Fegan, G., Fulford, T., Prentice, A.M., & Berkley, J.A. (2012a). Midupper arm circumference at age of routine infant vaccination to identify infants at elevated risk of death: a retrospective cohort study in the Gambia. *Bulletin of the World Health Organization*, 90, (12) 887-894.

Mwangome, M.K., Fegan, G., Mbunya, R., Prentice, A.M., & Berkley, J.A. (2012b). Reliability and accuracy of anthropometry performed by community health workers among infants under 6 months in rural Kenya. *Tropical Medicine and International Health*, 17, (5) 622-629.

Patwari, A.K., Kumar, S., & Beard, J. (2015). Undernutrition among infants less than 6 months of age: an underestimated public health problem in India. *Maternal and Child Nutrition*, 11, (1) 119-126.

Pongou, R., Ezzati, M., & Salomon, J.A. (2006). Household and community socioeconomic and environmental determinants of child nutritional status in Cameroon. *BMC Public Health*, 6, 98.

Singh, D.K., Rai, R., Mishra, P.C., Maurya, M., & Srivastava, A. (2014). Nutritional rehabilitation of children < 6 mo with severe acute malnutrition. *Indian Journal of Pediatrics*, 81, (8) 805-807.

The Sphere Project (2011). Humanitarian Charter and Minimum Standards in Humanitarian Response, Third Edition. Practical Action Publishing: Rugby.

Victora, C.G., Adair, L., Fall, C., Hallal, P.C., Martorell, R., Richter, L. et al. (2008). Maternal and child undernutrition: consequences for adult health and human capital. *Lancet*, 371, (9609) 340-357.

Vygen, S.B., Roberfroid, D., Captier, V., & Kolsteren, P. (2013). Treatment of severe acute malnutrition in infants aged <6 months in Niger. *Journal of Pediatrics*, 162, (3) 515-521.

Walters, T., Sibson, V., & McGrath, M. (2012). Mid upper arm circumference and weightfor-height z-score as indicators of severe acute malnutrition: A consulation of operational agencies and academic specialist to understand the evidence, identify knowledge gaps and to inform operational guidance. Available at: http://www.cmamforum.org/Pool/Resources/MUAC-WFH-Report-ENN-2013.pdf

UNICEF, WHO & World Bank. (2015). Levels and trends in child malnutrition. UNICEF-WHO-World Bank Group joint child malnutrition estimates, Geneva, New York and Washington DC.

WHO (1995). Physical status: the use and interpretation of anthropometry. Report of a WHO Expert Committee, Geneva.

WHO (2013). Guideline: updates on the management of severe acute malnutrition in infants and children, World Health Organization, Geneva.

WHO & UNICEF (2009), WHO child growth standards and the identification of severe acute malnutrition in infants and children. A Joint Statement.

WHO Multicentre Growth Reference Study Group (2006). WHO Child Growth Standards: Length/height-for-age, weight-for-length, weight-for-height and body mass index-for-age: Methods and development Geneva, World Health Organization.

## **LEGENDS TO FIGURES**

**Figure S1**. Age frequency distribution of a sample of 24,045 children aged 0-60 months from 10 countries (Burundi, Kenya, Liberia, Myanmar, Niger, Democratic Republic of Congo, Somalia, Sudan, Tajikistan and Uganda). The continuous line denotes the cumulative frequency.

**Figure S2**. Proportion of values of the weight-for-height z-score that are available for the assessment of the nutritional status at admission, by age group, from a sample of 24,045 children aged 0-60 months from 10 countries (Burundi, Kenya, Liberia, Myanmar, Niger, Democratic Republic of Congo, Somalia, Sudan, Tajikistan and Uganda).

**Figure 1**. Forest plot of the risk ratio (RR) of death during treatment for infants aged <6 months compared to children aged 6-60 months, by country, from a sample of 24,045 children aged 0-60 months from 10 countries (Burundi, Kenya, Liberia, Myanmar, Niger, Democratic Republic of Congo, Somalia, Sudan, Tajikistan and Uganda). Test for heterogeneity chi-squared = 67.0 (degrees of freedom = 9; p<0.01). I-squared (variation in RR attributable to heterogeneity) = 86.6%. Test of RR=1: z = 2.96 (p<0.01)

## TABLES

		Type of Therapeutic Care		6-60 months	<6 months	<6 months Age (month				
Country	Years	DC	HT	SC	TFC	n (%)	n (%)	mean $\pm$ s.d.	Min	Max
Afghanistan <sup>a</sup>	2002-04	633			460	63 (5.8)	1,030 (94.2)	$3.4 \pm 1.5$	0.5	9
Burundi	2006-07	2,359				2,213 (93.8)	146 (6.2)	$33.5\pm16.7$	0	60
DRC	2005-07				6,229	4,829 (77.5)	1,400 (22.5)	$18.7\pm15.6$	0	60
Ethiopia <sup>a</sup>	2008			57		24 (42.1)	33 (57.9)	$4.3 \pm 1.9$	0	8
Kenya	2005-07				539	502 (93.1)	37 (6.9)	$18.2 \pm 11.4$	1.5	60
Liberia	2006-08				2,436	2,269 (87.1)	167 (6.9)	$16.3\pm9.8$	1	60
Myanmar	2006-08		1,143	248		1,211 (87.1)	180 12.9)	$22.9 \pm 14.6$	0.1	60
Niger	2006-08				1,108	963 (86.9)	145 (13.1)	$14.6\pm9.2$	1	58
Somalia	2006-08				2,997	2,595 (86.6)	402 (13.4)	$17.7\pm13.0$	1	60
Sudan	2005-08			109	5,218	4,967 (93.2)	360 (6.8)	$18.0\pm9.7$	0	60
Tajikistan	2005-06				373	287 (76.9)	86 (23.1)	$10.9\pm6.7$	1	46
Uganda	2005-07				1,286	1,270 (98.8)	16 (1.2)	$21.6\pm10.8$	1	60
Total	2002-08	2,992	1,143	414	20,646	21,193 (84.1)	4,002 (15.9)	$19.0 \pm 14.0$	0	60

 Table 1. Programme datasets by country

DC: Day centre, HT: Home treatment, SC: Stabilisation centre, TFC: Therapeutic feeding centre, DRC: Democratic Republic of Congo <sup>a</sup> This programme data was excluded from analysis as it included only very young children with no older children for comparisons.

Died	Non-recovered	Defaulted	Admission error
Dead	Autres	Abandon	Admission mistake
Death	C.N.R	D	AM
Décédé	CNR	Default	СН
Décès	Critères non-atteints	Defaulter	Cheating
Died	Criteria not reached		Erreur d'admission
Μ	DNG		Error
	Inconnu		Mistake
	Medical transfer		Mistake admission
	Non guéri		
	Non répondant		
	Non respondant		
	Non respondent		
	Non responder		
	Non response		
	Non-respond		
	NR		
	Other		
	Others		
	т		
	TFC		
	To other OTP		
	Transfer		
	Transfer HP		
	Transfer Others		
	Transfer TFC		
	•		
	Unknown		
	Dead Death Décédé Décès Died	DeadAutresDeathC.N.RDécédéCNRDécèsCritères non-atteintsDiedCriteria not reachedMDNGInconnuMedical transferNon guériNon répondantNon respondantNon respondantNon responderNon responderNon responderNon-respondNROtherOthersTTFCTo other OTPTransferTransfer TFCTransfer TFCTransfer TFCTransfer to other OTPTransfer to TFCTransfer to TFCTransfertTransfert Centre de sTransfertTransfert CNTTransfert CNTTransfert HTransfert hopitalTransfert medicalTransfert wers crenas	DeadAutresAbandonDeathC.N.RDDécédéCNRDefaultDécèsCritères non-atteintsDefaulterDiedCriteria not reachedMMDNGInconnuMedical transferNon guériNon répondantNon respondantNon respondentNon responderNon responderNon-respondNROtherOthersTTFCTo other OTPTransferTransfer HPTransfer OthersTransfer to OTPTransfer to OTPTransfer to OTPTransfer to OTPTransfer to TFCTransfer to TFCTransfert CTransfert CCTransfertTransfert CNTTransfertTransfert CNTTransfert CNTTransfert CNTTransfert CSTransfert HTransfert hopitalTransfert hopitalTransfert wers crenas

 Table S1. Coding of original discharge outcomes (Supplementary Appendix)

Table 2. Quality of anthropometric data at	admission by age group.
--	-------------------------

Percentage of anthropome	tric data at admiss	ion that was miss	ing					
		<6 months (n = 2,939)		6 - 60 months		Difference		
				21,106)		1		
	%	95%CI	%	95%CI	%	95%CI	p-value	
Weight	0.51	0.25; 0.77	0.62	0.51; 0.72	-0.11	-0.38; 0.17	0.24	
Height/length	7.55	6.60; 8.50	0.63	0.53; 0.74	6.92	5.96; 7.89	< 0.01	
MUAC	49.4	47.6; 51.2	24.2	23.6; 24.8	25.2	23.3; 27.1	< 0.01	
Oedema data	1.60	1.15; 2.05	1.57	1.41; 1.74	0.03	-0.46; 0.51	0.46	
Percentage of anthropome	tric indices that co	ould not be calculated	ated when mea	surement data wa	s available			
	<6 r	nonths	6 - 60	) months		Difformer		
	(n =	(n = 2,939)		(n = 21, 106)		Difference		
	%	95%CI	%	95%CI	%	95%CI	p-value	
WAZ	0.00		0.01	0.00; 0.02	-0.01	-0.02; 0.00	0.30	
HAZ	0.00		0.00		0.00			
WHZ	16.0	14.7; 17.3	0.40	0.31; 0.48	15.6	14.3; 16.9	< 0.01	
Percentage of anthropome	tric indices flagge	d as outliers						
	<6 r	nonths	6 - 60	) months		Difference		
	(n =	2,939)	(n =	21,106)	Difference			
	%	95%CI	%	95%CI	%	95%CI	p-value	
Flag 1	1.40	0.97; 1.82	0.70	0.58; 0.81	0.70	0.26; 1.14	< 0.01	
Flag 2	6.91	5.99; 7.82	4.26	3.99; 4.53	2.65	1.69; 3.60	< 0.01	
Flag 3	1.91	1.41; 2.40	1.54	1.37; 1.71	0.37	-0.16; 0.89	0.07	
Any flag	8.47	7.47; 9.48	5.71	5.40; 6.02	2.76	1.71; 3.82	< 0.01	

WAZ: Weight-for-age z-score, HAZ: Height-for-age z-score, WHZ: Weight-for-height z-score, MUAC: Mid-upper arm circumference.

Flag 1: WAZ <-4 or >4 z-scores from the observed mean

Flag 2: HAZ <-4 or >3 z-scores from the observed mean Flag 3: WHZ <-4 or >4 z-scores from the observed mean

Table 3. Nutritional profile at admission of	E children aged 0-60 months by age group.	
--	---	--

Proportion of acute malnutrition at	admission						
		months	6-60 months		Difference		
	(n = 2, 190)		(n =	20,556)			
Indicator	mean or %	95%CI	mean or %	95%CI	mean or %	95%CI	p-value
Global (%)	85.4	84.4; 87.3	98.7	98.5; 98.8	-12.8	-14.3; -11.4	< 0.01
Moderate (%)	13.7	12.2; 15.1	4.70	4.41; 4.99	8.95	7.49; 10.4	< 0.01
Severe (%)	72.2	70.3; 74.1	94.0	93.7; 94.3	-21.8	-23.7; -19.9	< 0.01
Oedema (%)	5.53	4.57; 6.48	35.3	34.7; 36.0	-29.8	-31.0; -28.6	< 0.01
Proportion of wasting <sup>a</sup> at admission	on						
	<6 1	months	6 - 6	0 months		Difference	
	(n =	2,069)	(n =	13,295)	Difference		
Indicator	mean or %	95%CI	mean or %	95%CI	mean or %	95%CI	p-value
Total (%)	85.0	83.5; 86.6	98.0	97.7; 98.2	-12.9	-14.5; -11.4	< 0.01
Moderate (%)	14.5	12.9; 16.0	7.27	6.82; 7.71	7.19	5.61; 8.76	< 0.01
Severe (%)	70.6	68.6; 72.5	90.7	90.2; 91.2	-20.1	-22.2; -18.1	< 0.01
WHZ (z-score)	-3.89	-3.93; -3.85	-4.31	-4.32; -4.29	0.42	0.37; 0.46	< 0.01

D. ... c Instaiti . + . 

WHZ: Weight-for-height z-score.

Acute malnutrition: Global (WHZ<-2 and/or oedema), moderate (WHZ<-2 but  $\geq -3$ ) and severe (WHZ<-3 and/or oedema).

Wasting: Total (WHZ $\leq$ -2), moderate (WHZ $\leq$ -2 but  $\geq$ -3) and severe (WHZ $\leq$ -3).

<sup>a</sup> Wasting was measured among children with no reported oedema.

	<6 months (n = 2,939)		6 - 60 months (n = 21,106)		Difference		
Discharge outcome	%	95%CI	%	95%CI	%	95%CI	p-value
Recovered	75.7	74.2; 77.3	74.5	73.9; 75.1	1.23	-0.43; 2.89	0.08
Died	4.60	3.81; 5.31	3.95	3.68; 4.21	0.61	-0.19; 1.41	0.06
Non-recovered	10.2	9.14; 11.3	10.1	9.68; 10.5	0.15	-1.01; 1.32	0.40
Defaulted	6.43	5.54; 7.32	7.75	7.39; 8.11	-1.31	-2.27; -0.36	< 0.01
Admission error	0.37	0.15; 0.60	0.50	0.41; 0.60	-0.13	-0.37; 0.11	0.18
Missing values	2.69	2.10; 3.27	3.24	3.00; 3.48	-0.55	-1.18; 0.08	0.05

 Table 4. Discharge outcomes of children aged 0-60 months

## FIGURES

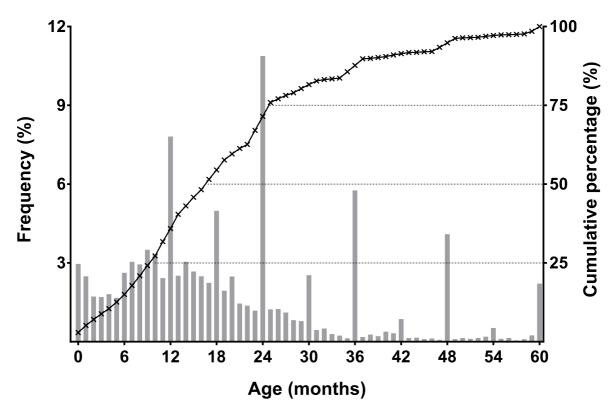


Figure S1 (Supplementary Appendix)

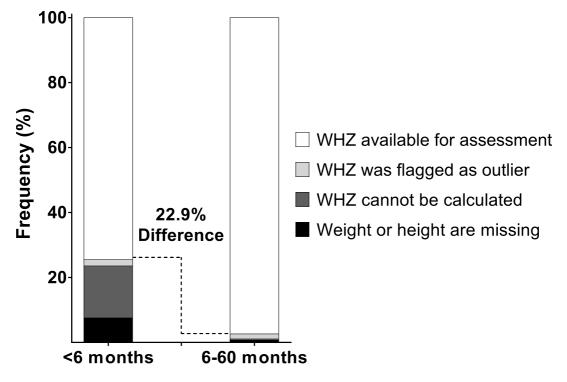


Figure S2 (Supplementary Appendix)

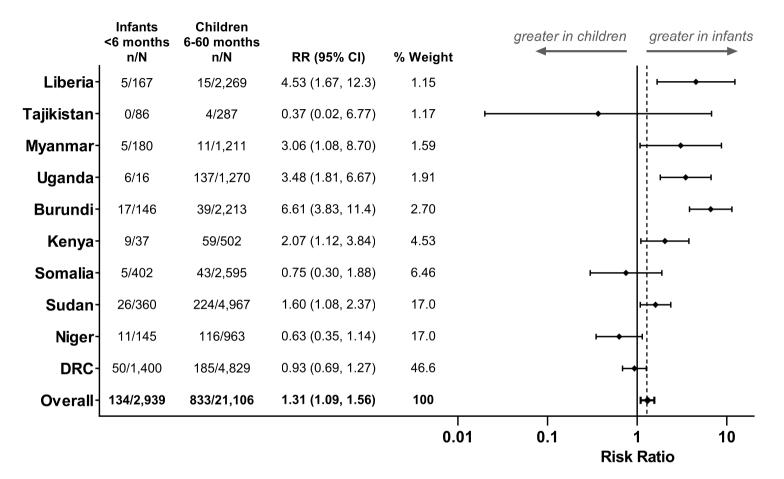


Figure 1