Tackling malnutrition: a systematic review of 15-year research evidence from INDEPTH health and demographic surveillance systems

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Background: Nutrition is the intake of food in relation to the body’s dietary needs. Malnutrition results from the intake of inadequate or excess food. This can lead to reduced immunity, increased susceptibility to disease, impaired physical and mental development, and reduced productivity.

Objective: To perform a systematic review to assess research conducted by the International Network for the Demographic Evaluation of Populations and their Health (INDEPTH) of health and demographic surveillance systems (HDSSs) over a 15-year period on malnutrition, its determinants, the effects of under and over nutrition, and intervention research on malnutrition in low- and middle-income countries (LMICs).

Methods: Relevant publication titles were uploaded onto the Zotero research tool from different databases (60% from PubMed). Using the keywords ‘nutrition’, ‘malnutrition’, ‘over and under nutrition’, we selected publications that were based only on data generated through the longitudinal HDSS platform. All titles and abstracts were screened to determine inclusion eligibility and full articles were independently assessed according to inclusion/exclusion criteria. For inclusion in this study, papers had to cover research on at least one of the following topics: the problem of malnutrition, its determinants, its effects, and intervention research on malnutrition in low- and middle-income countries (LMICs).

Results: The INDEPTH research identified rising levels of overweight and obesity, sometimes in the same settings as under-nutrition. Urbanisation appears to be protective against under-nutrition, but it heightens the risk of obesity. Appropriately timed breastfeeding interventions were protective against malnutrition.

Conclusions: Although INDEPTH has expanded the global knowledge base on nutrition, many questions remain unresolved. There is a need for more investment in nutrition research in LMICs in order to generate evidence to inform policies in these settings.

Keywords: nutrition; malnutrition; under and over nutrition; overweight; low- and middle-income countries; LMICs; health and demographic surveillance system

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Nutrition has been defined as the ‘science of food, the nutrients and other substances therein, their action, interaction and balance in relation to health and disease, and the processes by which the organism ingests, absorbs, transports, utilises and excretes food substances’ (1).

In low- and middle-income countries (LMICs) studies of nutrition generally focus on malnutrition, defined by
including cohort studies, qualitative focus group discussions, key informant interviews, literature reviews, clinical trials, and the testing of diagnostic tools. The centres have also developed and tested measurement and screening tools to facilitate a more accurate diagnosis of nutritional problems and to draw a more robust picture of food intake patterns in low- and middle-income settings.

Methods
In this paper, we review studies of malnutrition conducted by HDSSs published in peer-reviewed English language journals, and we outline the key findings reported by INDEPTH member centres and discuss their implications for future nutrition policy. This study includes 67 published papers on malnutrition between 1998 and 2013, covering nine countries in Africa and four in Asia.

Standard systematic review methods by Higgins and Green (13) and the Centre for Reviews and Dissemination, York, United Kingdom, were used (14). Relevant publications titles (related to malnutrition studies) from the member research centres of INDEPTH were uploaded onto the Zotero research tool from different databases (60% from PubMed). Using the keywords ‘nutrition, malnutrition, and over and under nutrition’, publications were selected that were based only on data generated through research which used the longitudinal HDSS platform. All titles and abstracts were screened to determine inclusion eligibility and full articles were independently assessed according to inclusion/exclusion criteria. For inclusion, papers had to cover research on at least one of the following topics: the problem of malnutrition, its determinants, its effects, and intervention research on malnutrition. One hundred and forty eight papers were identified and independently reviewed by the authors and 67 were selected for this study.

This paper deals with five tropical issues: The problem of malnutrition as manifest in member centres’ demographic surveillance areas, including the prevalence of malnutrition and population groups most affected; the biological and social determinants of malnutrition; the effects of malnutrition; the interventions that have attempted to tackle malnutrition; and concludes with discussion and policy and research recommendations (Fig. 1).

Results

The problem of malnutrition

Under-nutrition

Insufficient or inadequate food intake – be it from lack of quantity or lack of essential nutrients including protein and specific micronutrients – remains a blight across the developing world. Under-nutrition can have effects throughout the life cycle. Physical stunting (defined by the WHO as low height-for-age) can affect cognitive development, increase susceptibility to infection, and impair school attainment and future productivity of survivors as well as of later generations. Deficiencies in iron, iodine, zinc, and vitamins can cause problems ranging from brain damage...
to blindness, anaemia, and stunted growth. Being underweight makes it harder to fight off infection and recover from illness, and harder to study and work.

Inadequate food intake in the early years of life is particularly damaging. In a cohort survey of children at Nouna HDSS in Burkina Faso, the authors found a reduction in underweight prevalence from 40% in June 1999 to 35% in June 2009 and from 43% in December 1999 to 34% in December 2009. The prevalence for wasting and severe wasting prevalence remained high in under-five children (15). In a 2010 study by the same centre found that many women began pregnancy with suboptimal nutritional status and that most did not gain enough weight during pregnancy to avert the risk of adverse health outcomes (20). Malnutrition in pregnant women can pose problems not only for the women themselves but also for their newborns. A study in West Kiang HDSS, Gambia, showed that nutritional rickets, a disease caused largely by vitamin D deficiency, the effects of which can include cardiac failure and hypoglycaemic seizures, was more prevalent in infants whose mothers had poor vitamin D status during pregnancy (21).

A study of 457 people aged 60 years and above in rural Bangladesh found that 50% of older adults had chronic energy deficiency, and 62% were at risk of malnutrition (22). The authors argued that if global targets to reduce world hunger are to be achieved, ‘it is important to recognise that a substantial proportion of the elderly population, particularly in low-income countries, is undernourished’ (22).

**Overweight and obesity**

Under-nutrition is not the only form of malnutrition. Overweight and obesity, caused by an over-consumption of calories relative to energy expended, has long been considered a problem only in high-income countries, but in recent years it has begun to emerge as a threat in poorer countries.

As countries grow wealthier, the intake of refined and fatty foods and foods of animal origin increases. In 2002, the WHO ranked being overweight as the fifth most serious risk factor underlying the disease burden in low-mortality developing countries. INDEPTH member centres have begun to track the rise of obesity in developing countries over the past decade and a half. In the study of adult women by the Purworejo HDSS in Indonesia, researchers found an obesity rate in 1997 of 14% and showed that the rate had increased by almost three percentage points over the previous 12 months (19). Gendered patterns exist in rural South Africa. One study showed that females aged 50 years and older in Agincourt were significantly more obese than males, yet there was no difference in the prevalence of hypertension between men and women (23).

A 2005 study of 3-year-old children in the Limpopo province of South Africa found that 22% were overweight.
and 24% were obese. Nineteen percent of children were found to have both stunted growth and obesity (24). In Agincourt, rural South Africa, Kimani-Murage et al. (25) investigated adolescent weight status and central obesity (measured by abdominal circumference) and found much higher rates of combined overweight and obesity and of central obesity among girls (15% for each) than among boys (4% of whom had combined overweight and obesity, and 2% central obesity). In that study, risk factors for overweight and obesity included high socio-economic status and a household head who had completed secondary education. For central obesity, the risk factors included age, having a mother aged 50 years or above, and having high socio-economic status. A study at Kanchanaburi HDSS in Thailand found that urban residence, affluence, and possession of a television were associated with higher rates of obesity (26). Under-nutrition, on the other hand, was associated with poverty. The studies on over-nutrition point to the impact of modernisation in transforming lifestyles and suggest that as countries develop economically, the risk of obesity may increase.

Like under-nutrition, over-nutrition can also have harmful impacts on health. A cohort study of 1,888 rural women in Bangladesh found that those with the highest body mass index (BMI) had a higher mortality risk than those with intermediate BMI (27). Table 1 displays key findings on under-nutrition and obesity.

The determinants of malnutrition

Biological causes

Under-nutrition does not only come about because of lack of food. Exposure to disease can increase the risks posed to health by under-nutrition.

In a study at the Agincourt HDSS in South Africa, Kimani-Murage et al. (28) found that children infected with HIV had significantly worse nutritional outcomes than their uninfected peers. In Gambia, HIV-positive children had significantly worse scores in terms of weight-for-age, height-for-age, and BMI than HIV-negative children (29). In mothers, too, HIV poses problems. Research at the Africa Centre in KwaZulu-Natal found that breastfeeding mothers in South Africa who were HIV-positive had poorer protein and micronutrient status than HIV-negative women (30). Mean serum concentrations of albumin, pre-albumin, folate, retinol, and haemoglobin were all lower in the group of 92 HIV-positive breastfeeding women than in a control group of 52 HIV-negative women. Filteau et al. (31) found that providing pregnant women who are HIV-positive with vitamin A supplementation may improve the gut function of their HIV-infected infants. They also found that vitamin A supplementation for HIV-infected infants may decrease gastrointestinal morbidity.

Nyakeriga et al. (32) found a significant relationship between malaria and subsequent underweight and stunting in children below the age of 2 years at the Kilifi HDSS in Kenya. However, the authors found no association between malaria and subsequent protein-energy malnutrition. A Kisumu HDSS study in Kenya showed that infants who had had malaria were more likely to have stunting, anaemia, and iron deficiency (33). On the other hand, researchers at Farafenni HDSS in Gambia found no association between malaria and subsequent malnutrition in children below the age of 5 years during the malaria season (34). In Nouna, Burkina Faso, anaemia was significantly associated with malnutrition but not with the frequency of malaria episodes or its prevalence (35).

Table 1. Prevalence of under-nutrition and obesity: findings from INDEPTH studies

| HDSS site                  | Year | Authors               | Main findings                                                                 |
|----------------------------|------|-----------------------|-------------------------------------------------------------------------------|
| Nouna HDSS, Burkina Faso   | 2000 | Beiersmann et al.     | Slight reduction in the underweight prevalence from 40% in June 1999 to 35% in June 2009 and from 43% in December 1999 to 34% in December 2009. |
| Agincourt HDSS, South Africa| 2010 | Kimani-Murage et al. | 20% of children aged 1–4 years had stunted growth.                           |
| Agincourt HDSS, South Africa| 2007 | Saloojee et al.       | 45% of children below the age of 5 had stunted growth.                       |
| Purworejo HDSS, Indonesia  | 2000 | Winkvist et al.       | 15% of adult women had chronic energy deficiency from insufficient food intake; 14% were obese. |
| Matlab HDSS, Bangladesh    | 2006 | Kabir et al.          | 50% of adults over 60 years had chronic energy deficiency; 62% at risk of malnutrition. |
| Agincourt HDSS, South Africa| 2013 | Gomez-Olive et al.    | Women aged 50 years or above were significantly more obese than males.       |
| Dikgale HDSS, South Africa | 2005 | Mamabolo et al.       | 22% of 3-year-old children were overweight or obese; 19% had both stunted growth and obesity. |
| Agincourt HDSS, South Africa| 2011 | Kimani-Murage et al.  | 15% of adolescent girls were overweight or obese, and 4% of adolescent boys. |
A Kilifi HDSS study identified hospital-acquired bacteremia (nosocomial bacteraemia) as a significant risk factor for severe malnutrition. Nosocomial infections, the study authors wrote, ‘are largely unrecognised or undocumented as a health risk in low-income countries, but they are likely to become public health priorities as awareness of their occurrence increases and as other prominent childhood diseases are progressively controlled’ (36). Table 2 displays key findings on biological causes of malnutrition.

Social causes
INDEPTH member centres have extensively investigated the social factors that increase the risk of malnutrition in LMICs. Three major contributors have been identified.

The first is wealth. Families themselves recognise this as important. Focus group discussions among mothers in Gambia revealed that poverty is one of the key constraints preventing mothers from putting into practise their knowledge of child health and nutrition (37). In similar discussions in Kenya, mothers reported that financial constraints were the main cause of severe malnutrition in children, with mothers’ heavy workload and inability to generate income were also reported as key factors. Witchcraft and the violation of sexual taboos were among the other reported causes (38). Household wealth and community socio-economic status were significantly associated with childhood stunting in Nairobi, Kenya (39). The authors of this study posited that the community effect occurs because of the availability of social services, physical environment, and the wealth of individuals. In an earlier paper, the same authors showed that neighbourhood characteristics exert an influence on malnutrition independently of household wealth – ‘when basic socioeconomic and health services are lacking in the poorest communities’, they observed, ‘families therein can hardly take advantage of their increased means, ability and knowledge in caring for their children’ (40). In Agincourt, rural South Africa, cases with severe malnutrition were from households with fewer assets and lower incomes, who were less likely to receive a social support grant or to have anyone employed, compared with control groups in the village (17). In the same area, Kimani-Murage et al. (25) observed four-fold higher odds of overweight for adolescent girls and two-fold higher odds for participants from households with the highest socio-economic status. The causes of childhood malnutrition have been studied more extensively than those of adult malnutrition, but there is evidence that wealth continues to play a part in nutritional status later in life. As a 1998 study at Matlab HDSS in Bangladesh showed, women aged 35 years and older from wealthier households were less likely to have chronic energy deficiency than poorer women (41).

Education is a second key factor. Children born to Nairobi mothers who had been educated to primary level had a 94% lower chance of stunting than those born to mothers with no education (42). In one study in Vietnam, birth weight and growth were statistically significantly

Table 2. Biological causes of malnutrition: findings from INDEPTH studies

| HDSS site          | Year | Authors               | Main findings                                                                 |
|--------------------|------|-----------------------|-------------------------------------------------------------------------------|
| Agincourt HDSS,    | 2011 | Kimani-Murage et al.  | Children infected with HIV had significantly worse nutritional outcomes than their uninfected peers. |
| South Africa       |      |                       |                                                                               |
| West Kiang HDSS,   | 2008 | Nweneka               | HIV-positive children had significantly worse scores in terms of weight-for-age, height-for-age, and BMI than HIV-negative children. |
| Gambia             |      |                       |                                                                               |
| Africa Centre,     | 2007 | Paphathakis et al.    | Breastfeeding mothers in South Africa who were HIV-positive had poorer protein and micronutrient status than HIV-negative women. |
| South Africa       |      |                       | Providing pregnant women who are HIV-positive with vitamin A supplementation may improve gut function of their HIV-infected infants. Vitamin A supplementation for HIV-infected infants may decrease gastrointestinal morbidity. |
| Africa Centre,     | 2001 | Filteau et al.        |                                                                               |
| South Africa       |      |                       |                                                                               |
| Kilifi HDSS, Kenya | 2004 | Nyakeriga et al.      | Significant relationship between malaria and subsequent underweight and stunting in children below the age of 2 years. |
|                    |      |                       | Infants who had had malaria were more likely to have stunting, anaemia, and iron deficiency. |
| Kisumu DSS, Kenya  | 2005 | Mamiro et al.         |                                                                               |
| Farafenni HDSS, Gambia | 2002 | Deen et al.           | No association between malaria and subsequent malnutrition in children below the age of 5 years during the malaria season. |
| Nouna HDSS,        | 2003 | Muller et al.         | Anaemia was significantly associated with malnutrition.                        |
| Burkina Faso       |      |                       |                                                                               |
| Kilifi HDSS, Kenya | 2011 | Aiken et al.          | Hospital-acquired bacteraemia (nosocomial bacteraemia) a significant risk factor for severe malnutrition. |
and positively associated with economic conditions and the education of the mother (43). In another study however, mothers’ education level was associated with their children’s stunting in early childhood (44). As with wealth, education continues to have repercussions for nutritional status later in life – in the above-mentioned study in Bangladesh, women over the age of 35 were half as likely to suffer chronic energy deficiency if they had had one or more years of education, than if they had had no schooling (41).

Linked to both wealth and education is the associated process of urbanisation. Five of the nutrition studies carried out by INDEPTH member centres between 1998 and 2013 demonstrate a link between urban residence and reduced malnutrition. In a study in FilaBavi, Vietnam, Nguyen et al. (45) found an association between household socio-economic status, education of the mother and birth weight. The Nairobi study (39) discussed above confirmed this finding, with urban children significantly less likely to be malnourished than rural children, while two studies in Senegal showed that rural adolescent girls who migrated seasonally to urban areas for work had improved nutritional status, as reflected in BMI, compared with girls who remained in villages (46, 47). Although urbanisation appears to reduce the risk of under-nutrition, it may increase that of obesity. A study at Kanchanaburi HDSS in Thailand found that urban residence was associated with lower rates of under-nutrition in children, but also with higher rates of obesity (26). Household wealth was also associated with obesity.

One additional, yet less extensively studied social factor, which also appears to have an impact on nutrition, is paternal involvement in caring for children. Children in a study in South Africa whose fathers did not provide financial support were found to be at higher risk of malnutrition (48), while in Hanoi, Vietnam, children whose fathers took them to a medical facility to be immunised against childhood diseases were 1.7 times less likely to be underweight and stunted than other children (49). These points to the need for a multi-pronged response to the problem of malnutrition which has many causes (Table 3).

The effects of malnutrition

Obesity is a relatively new threat in LMICs and is, as yet, not generally widespread in these countries. Therefore in researching the effects of malnutrition, HDSS centres have largely focused their attentions on the problem of under-nutrition.

A study by the Kilifi HDSS in Kenya found that among children admitted to a rural district hospital, under-nutrition was a factor in half the in-hospital deaths and half the morbidity from severe diseases such as malaria, gastroenteritis, lower respiratory tract infection, HIV, and invasive bacterial disease (50). At West Kiang in Gambia, young adults born in the ‘hungry season’ – that is, the non-harvest season between July and December when less food is available – had 10 times higher mortality than those born in the harvest season (51). Conversely, two studies at Institut de Recherche pour le Developpement IRD HDSS in Senegal found that although child mortality in the rural Casamance region declined sharply between 1969 and 1992, infants’ nutritional status did not improve over the period, suggesting other factors such as vaccination were more important (52, 53).

As well as increasing mortality, under-nutrition also impairs the development of those who survive it. In a study in Bangladesh, stunting in early childhood, which is in part the result of a combination of disease and under-nutrition, was found to be a significant predictor of stunting in adolescence (54). A four-country study covering Ethiopia, India, Peru, and Vietnam found that stunting in early childhood was associated with lower cognitive achievement in 5-year-old children (44). In infants and young children in Kenya, stunting is also associated with developing severe respiratory syncytial virus-associated pneumonia, and with lower respiratory tract infections from all causes (55).

The effect of under-nutrition on malaria outcomes is disputed in the literature. In a Kilifi HDSS study, Berkley et al. (56) found that malnutrition was associated with severe disease due to falciparum malaria. A cohort study at Farafenni HDSS in Gambia calculated that 51% of children with stunting subsequently experienced malaria during the malaria season, compared with 38% of children who were not stunted. The study authors concluded: ‘Our findings suggest that chronically malnourished children may be at higher risk for developing malaria episodes’ (34). On the other hand, Muller et al. (18), working at Nouna HDSS in Burkina Faso, found no association between protein-energy malnutrition and malaria morbidity, while a Kilifi HDSS study concluded that children with iron deficiency were less likely than other children to develop mild clinical malaria (32). Table 4 displays key findings on effects of malnutrition.

Tackling malnutrition

The role of breastfeeding

Nutrition in infancy, as we have seen, is important to an individual’s prospects of a good life. Breastfeeding has therefore been the subject of much analysis by nutrition researchers, who have assessed its benefits and analysed breastfeeding practices in LMICs.

Working at the Kintampo HDSS in Ghana, Edmond et al. (57) demonstrated for the first time that there is a causal link between early breastfeeding and declines in infection-specific neonatal mortality in infants aged between 2 and 28 days. Delayed initiation of breastfeeding until after the first day of life led to a 2.6 times higher risk of neonatal mortality as a result of infectious disease, while partial breastfeeding led to a 5.7 times higher risk.
Researchers at Bandim HDSS in Guinea-Bissau found that children weaned early for reasons other than ‘being healthy’, including a new pregnancy, had higher mortality (58). Bahl et al. (59), meanwhile, found that non-breastfed infants had a 10-fold higher risk of dying than predominantly breastfed and exclusively breastfed infants, although there was no significant difference in risk between the latter two groups. This finding suggests that in areas where most women already predominantly breastfeed, breastfeeding promotion efforts should be directed at those who chose not to breastfeed or who partially breastfed. A case–control study by Saloojee et al. in rural South Africa (17) also found that risk factors for severe malnutrition included poor weaning practices.

The positive effects of breastfeeding have also been demonstrated in older children. Vohr et al. (60) found that extremely low birth weight babies who ingested breast milk while in the neonatal intensive care unit (NICU) had significantly higher scores in cognitive and behaviour tests at 18 months of age than similar non-breastfed NICU infants.

However, breastfeeding is not universally practised. Researchers at Nanoro HDSS in Burkina Faso who...
conducted a multi-centre study of women infected with HIV, highlighted the size of the task facing those engaged in breastfeeding promotion efforts. In sub-Saharan Africa, fluids other than maternal milk are often introduced within the first six months of life, notwithstanding WHO guidelines that recommend exclusive breastfeeding for the first six months (including where the mother is HIV-positive). Although 53% of women in the Nanoro study saw exclusive breastfeeding as the preferred feeding method, only 11% reported exclusively breastfeeding after previous pregnancies (61). Conversely, some mothers choose to prolong breastfeeding beyond the recommended duration, which has been found to be associated with reduced nutritional status and impaired growth. However, as Simondon and Simondon (62) and Simondon et al. (63) found in two studies conducted in Senegal, these consequences are a result not of prolonged breastfeeding itself, but of the fact that women prolong breastfeeding beyond 12 months for children who are undernourished and tend to wean well-nourished children. The authors surmised that this is a result of mothers’ awareness of the mortality risks to children following weaning.

Prevention efforts
A number of interventions to prevent or cure malnutrition have been tested by HDSS sites. The vast majority of the trials have focused on infants, children and women of childbearing age.

Efforts to tackle malnutrition in infants and children have met with mixed success. Studies by the Bandim HDSS in Guinea-Bissau of malnutrition campaigns during that country’s 1998 civil war found that vitamin A supplementation for children (64) and a supplementary feeding programme for children (65) helped to reduce mortality during the conflict. Compared with the 3-year period before the war, the authors reported that children offered vitamin A at home during the war had a 12% reduction in mortality, whereas the overall impact of the conflict was an 89% increase in mortality.

In a randomised controlled trial in Vietnam, fortification with multiple micronutrients of biscuits provided in schools was found to reduce the risk of anaemia and of iron and zinc deficiencies by 40% (66). At Nouna HDSS in Burkina Faso, on the other hand, a randomised controlled trial to assess the effects of zinc supplementation on the growth of young children in an area with high rates of malnutrition found no significant impact on height-for-age, weight-for-age, and weight-for-height scores. The study authors concluded that zinc supplementation does not have an effect or public health importance on growth in West African populations of young children with a high prevalence of malnutrition, and that multi-nutrient interventions are likely to be more effective (67).

The effectiveness of vitamin A supplementation has also been questioned. Although it appeared to be beneficial in

### Table 4. Effects of malnutrition: findings from INDEPTH studies

| HDSS site                  | Year  | Authors       | Main findings                                                                                                                                 |
|----------------------------|-------|---------------|-----------------------------------------------------------------------------------------------------------------------------------------------|
| Kilifi HDSS, Kenya         | 2008  | Bejon et al.  | Under-nutrition a factor in half of in-hospital deaths of children and half of morbidity from severe diseases.                                |
| West Kiang HDSS, Gambia   | 2011  | Ngom et al.   | Ten times higher mortality among young adults born in the hunger season than in harvest season.                                             |
| Niakhar HDSS, Senegal      | 2006  | Enel et al.   | Decline in child mortality despite no improvement in nutritional status among children.                                                        |
| Niakhar HDSS, Senegal      | 2004  | Pinchinat et al. | Decline in child mortality despite no improvement in nutritional status among children.                                                     |
| Matlab HDSS, Bangladesh    | 2008  | Bosch et al.  | Stunting in early childhood a significant predictor of stunting in adolescence.                                                               |
| Multiple sites in Ethiopia, India, Peru, Vietnam | 2013  | Barnett et al. | Stunting in early childhood associates with lower cognitive achievement among 5-year-olds.                                               |
| Kilifi HDSS, Kenya         | 2009  | Berkley et al. | Malnutrition associated with severe falciparum malaria.                                                                                     |
| Farafenni HDSS, Gambia     | 2002  | Deen et al.   | 51% of children with stunting subsequently experienced malaria during the malaria season, compared with 38% of children who were not stunted.|
| Nouna HDSS, Burkina Faso   | 2003  | Muller et al. | No association between protein-energy malnutrition and malaria morbidity.                                                                     |
| Kilifi HDSS, Kenya         | 2004  | Nyakeriga et al. | Children with iron deficiency were less likely than other children to develop mild clinical malaria.                                       |
the above-mentioned trial in Guinea-Bissau, researchers at Kintampo HDSS in Ghana found no positive effect of vitamin A supplements in combination with vaccines during the first few months of life (68). This finding echoes the findings of similar trials in Peru and India. Fish oil supplementation, too, appears to have limited effects for infants. A study in Gambia found that fish oil had no effect on growth, intestinal integrity, morbidity, or cognitive development (69).

Only two studies investigated methods to treat children suffering from severe malnutrition. Current guidelines for treating children who develop shock as a result of severe malnutrition recommend using low-dose hypotonic fluid resuscitation. Working at the Kilifi HDSS in Kenya, Akech et al. (70) tested the safety and efficacy of this recommendation and found ‘universally poor’ outcomes in terms of persistent shock, oliguria, and high case fatality. The authors concluded that the low recommended dose was insufficient to correct shock and suggested that for future guidelines to be more effective, clinical investigation is needed to determine the appropriate volume and rate of dosage of isotonic fluids. Researchers at Agincourt HDSS, meanwhile, found that the introduction of WHO guidelines in two hospitals strengthened the management of malnutrition. The authors concluded that implementation of WHO guidelines on severe malnutrition was feasible, affordable, and sustainable (71).

Studies of malnutrition prevention campaigns for women of childbearing age also had mixed results. In Ghana, vitamin A supplementation was found to increase liver reserves of mothers during the postpartum period, with the effects persisting for at least five months (72). A study at Niakhar HDSS in Senegal found that providing high energy, nutrient-dense food supplements to infants aged between four and seven months contributed to decreased postpartum weight loss in their mothers (73). The supplements reduced the intake of breast milk and therefore the energy costs of lactation on mothers. It also increased the interval between births, thereby potentially having effects on fertility. As with supplements for children, a number of studies – all conducted at the West Kiang HDSS in Gambia – found that supplements of individual nutrients for women were ineffective. Hawkesworth et al. (74) found that nutrient supplementation during pregnancy had little effect on infants’ risk of cardiovascular disease. Another study found that protein-energy supplementation during pregnancy had no effect on the blood pressure of their children in adolescence (75), while a third study found that calcium supplementation for pregnant mothers had no effect on foetal and infant growth (21). A fourth West Kiang study found that calcium supplements could be harmful – women given calcium supplements while pregnant were found to have lower bone mineral content, bone area, and bone mineral density at the hip during the 12-month lactation period than other women. The authors surmised that such supplements may disrupt metabolic adaptation (76).

Two studies conducted by INDEPTH members in South Africa and Bangladesh addressed the problem of implementing nutrition programmes. In the first, at Dikgale HDSS, the majority of people with type 2 diabetes had poor glycaemic control and were obese or had high blood pressure. Quantitative and qualitative research showed that these patients were often given incorrect and inappropriate dietary advice by health educators (77). In the second study, researchers showed that agencies working to tackle a flood disaster in Bangladesh in 1998 had mixed results in terms of meeting international disaster response standards. Compliance with standards was variable. Preliminary nutritional analysis was one of the areas in which agencies performed poorly in this regard (78).

Discussion

The state of research

Over the past 15 years, INDEPTH member centres have expanded the global knowledge base on nutrition. They have shown how under-nutrition continues to plague children and adults of all ages in LMICs and they have highlighted the emergence of obesity as a growing threat. In some areas, combined under-nutrition early in life with overweight and obesity in adolescence and adulthood, warns of rapidly escalating risk for cardiometabolic disease. The HDSS centres have demonstrated how malnutrition early in life continues to have effects on physical and mental development throughout the life-cycle. They have examined links between disease and malnutrition, and shown how wealth, education, and urbanisation all reduce the risks of under-nutrition but at the same time can increase the risks of obesity. The double burden of malnutrition in LMICs is of major concern to the World Bank, the WHO, and other international bodies.

In terms of tackling malnutrition, INDEPTH members have demonstrated the importance of breastfeeding for infant nutrition and health, and for the health of the infants as they grow older, and they have assessed the reasons why some women choose not to breastfeed or to wean infants too early or too late. Interventions such as micronutrient supplementation for children and their mothers can have beneficial impacts on child health and survival, with some studies suggesting that multiple micronutrient supplementation may be more effective than supplementation with single micronutrients (Table 5).

Nutrition is something of an under-researched field; however, over the period 1999–2013, INDEPTH member centres produced almost four times as many published papers on HIV/AIDS as they did on nutrition. Many issues remain unresolved – for example, the impact of malaria on malnutrition and of malnutrition on susceptibility
to malaria. There was little attention paid to nutrient supplements or to education and information campaigns to help families improve their diet, while activism campaigns to encourage companies to label foods accurately or to reduce sugar or fat content received no attention from HDSS centres. Likewise, the economic benefits of nutrition interventions also received little attention from HDSSs. One study by the Africa Centre calculated the costs of adding nutritional supplementation to the management of HIV-infected children (79), but studies on the cost-effectiveness of nutrition programmes were lacking.

**Policy implications**

In this review, we noted the importance of adopting a multi-pronged approach to tackling malnutrition. Countries including Vietnam and South Africa have had some success in reducing malnutrition in recent years. Researchers at Kaya HDSS in Burkina Faso carried out a nine-site study to assess the impact of United Nations Millennium Project guidelines to reduce child stunting. This holistic approach incorporated nutrition-specific, health-based interventions with food system and livelihood-based approaches, and its adoption reduced stunting in children by up to

| HDSS site                  | Year | Authors             | Main findings                                                                 |
|----------------------------|------|---------------------|-------------------------------------------------------------------------------|
| Bandim HDSS, Guinea-Bissau | 2004 | Nielsen et al.      | During civil war, vitamin A supplementation for children helped reduce mortality. |
| Bandim HDSS, Guinea-Bissau | 2005 | Nielsen et al.      | During civil war, supplementary feeding programme helped reduce mortality in children. |
| FilaBavi HDSS, Vietnam      | 2009 | Nga et al.          | Fortification with multiple micronutrients of biscuits provided in schools reduced risk of anaemia and of iron and zinc deficiencies by 40%. |
| Nouna HDSS, Burkina Faso   | 2003 | Müller et al.       | No significant impact of zinc supplementation for children on height-for-age, weight-for-age, and weight-for-height scores. |
| Kintampo HDSS, Ghana       | 2007 | Newton et al.       | No positive effect of vitamin A supplements in combination with vaccines during the first few months of life. |
| Farafenni HDSS, Gambia     | 2013 | Van der Merwe et al.| Fish oil supplementation had no effect on growth, intestinal integrity, morbidity, or cognitive development. |
| Kilifi HDSS, Kenya         | 2010 | Akech et al.        | Use of low-dose hypotonic fluid resuscitation for children who develop shock as a result of severe malnutrition has poor outcomes in terms of persistent shock, oliguria, and high case fatality. |
| Agincourt HDSS, South Africa | 2003 | Deen et al.        | Introduction of WHO guidelines in two hospitals strengthened the management of malnutrition. |
| Kintampo HDSS, Ghana       | 2006 | Tchum et al.       | Vitamin A supplementation increased liver reserves of mothers during the postpartum period, with effects persisting for at least five months. |
| Niakhar HDSS, Senegal      | 2006 | Ly et al.          | Provision of high energy, nutrient-dense food supplements to infants aged between four and seven months contributed to decreased postpartum weight loss in their mothers. |
| West Kiang HDSS, Gambia    | 2011 | Hawkesworth et al. | Nutrient supplementation during pregnancy had little effect on infants’ risk of cardiovascular disease. |
| West Kiang HDSS, Gambia    | 2009 | Hawkesworth et al. | Protein-energy supplementation of mothers during pregnancy had no effect on the blood pressure of their children in adolescence. |
| West Kiang HDSS, Gambia    | 2009 | Prentice et al.    | Calcium supplementation for pregnant mothers had no effect on foetal and infant growth. |
| West Kiang HDSS, Gambia    | 2010 | Jarjou et al.      | Women given calcium supplements while pregnant had lower bone mineral content, bone area and bone mineral density at the hip during the 12-month lactation period than other women. |
| Dikgale HDSS, South Africa | 2002 | Nthangeni et al.   | The majority of people with type 2 diabetes had poor glycaemic control and were obese or had high blood pressure and were given poor advice by health educators. |
| Africa Centre HDSS, South Africa | 2002 | O’Donnell et al. | Agencies working to tackle a flood disaster in Bangladesh in 1998 achieved mixed results in terms of meeting international disaster response standards. Some agencies complied with most standards, some with few. |
43% over 3 years (80). Programmes in Vietnam to increase nutrition, particularly via animal-source foods, have also been successful in reducing under-nutrition in children and chronic energy deficiency in women (81), but otherwise investigations of nationwide or multi-pronged nutrition programmes have not been extensively studied by HDSS centres. Lessons from other countries on how to reduce malnutrition may be a useful guide for policy-makers who are attempting to tackle the problem in their own countries.

Although a small number of studies have examined the success of efforts to implement nutrition programmes, there is a need for further investigation in this area. How to effectively change dietary habits, how to educate individuals in LMICs about the availability of healthy natural foods, how to improve compliance with food supplementation programmes, and how to train health workers to provide useful nutrition advice and effective treatment for nutrition problems, are just a few of the many implementation challenges that remain.

The INDEPTH HDSSs have unearthed important findings over the past decade and a half – for example, the vital role of breastfeeding in promoting infant and child health, the benefits of multiple micronutrient supplementation, the under-appreciated prevalence of malnutrition in adults and the elderly, and the rise of obesity in lower-income settings. Although much knowledge has already been achieved, there is broad scope for future research to inform policy-makers and researchers on ways to tackle the double burden of malnutrition in order to have a major impact on policy and on the lives of people living in LMICs.

Authors’ contributions
All the authors contributed to drafting, analysing, reviewing, and finalising this manuscript.

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