Advancing Early Childhood Development: from Science to Scale 2

Nurturing care: promoting early childhood development


The UN Sustainable Development Goals provide a historic opportunity to implement interventions, at scale, to promote early childhood development. Although the evidence base for the importance of early childhood development has grown, the research is distributed across sectors, populations, and settings, with diversity noted in both scope and focus. We provide a comprehensive updated analysis of early childhood development interventions across the five sectors of health, nutrition, education, child protection, and social protection. Our review concludes that to make interventions successful, smart, and sustainable, they need to be implemented as multi-sectoral intervention packages anchored in nurturing care. The recommendations emphasise that intervention packages should be applied at developmentally appropriate times during the life course, target multiple risks, and build on existing delivery platforms for feasibility of scale-up. While interventions will continue to improve with the growth of developmental science, the evidence now strongly suggests that parents, caregivers, and families need to be supported in providing nurturing care and protection in order for young children to achieve their developmental potential.

Introduction

Although global attention to early childhood development has been established through its inclusion in the UN Sustainable Development Goals, 250 million children (43%) younger than 5 years in low-income and middle-income countries are at risk of not achieving their developmental potential, as discussed in Paper 1 of this Series.1 We suggest that this gap in human potential is partly due to two reasons: the failure to apply emerging scientific knowledge on nurturing care to shape young children’s development; and the failure to take action at scale, using a multi-sector approach across key stages in the early life course.

We define nurturing care as a stable environment that is sensitive to children’s health and nutritional needs, with protection from threats, opportunities for early learning, and interactions that are responsive, emotionally supportive, and developmentally stimulating. As an overarching concept, nurturing care is supported by a large array of social contexts—from home to parental work, child care, schooling, the wider community, and policy influences.1 Nurturing care consists of a core set of inter-related components, including: behaviours, attitudes, and knowledge regarding caregiving (eg, health, hygiene care, and feeding care); stimulation (eg, talking, singing, and playing); responsiveness (eg, early bonding, secure attachment, trust, and sensitive communication); and safety (eg, routines and protection from harm).4 The single most powerful context for nurturing care is the immediate home and care settings of young children often provided by mothers, but also by fathers and other family members, as well as by child-care services.

The brain has evolved to adapt in response to a wide range of early experiences, which supports the rapid acquisition of language, cognitive skills, and socio-emotional competencies. Nurturing care mediates the development of key brain regions and promotes developmental adaptations. These developments have lifelong benefits for children, including an increased ability to learn, greater achievement in school and later life, citizenship, involvement in community activities, and overall quality of life.6,7 The period of early development is key stages in the early life course.

Key messages

- Advances in basic and intervention science indicate that early childhood is a period of special sensitivity to experiences that promote development, and that critical time windows exist when the benefits of early childhood development interventions are amplified.
- The most fundamental promotive experiences in the early years of life come from nurturing care and protection received from parents, family, and community, which have lifelong benefits including improved health and wellbeing, and increased ability to learn and earn.
- Nurturing care and protection are supported by a range of interventions delivered pre-pregnancy and throughout birth and the newborn period, infancy, and early childhood. Many of these interventions have shown benefits for child development, nutrition, and growth, and reductions in morbidity, mortality, disability, and injury.
- Interventions that integrate nurturing care and protection can target multiple risks to developmental potential at appropriate times, and can be integrated within existing preventive and promotive packages.
- Preventive and promotive packages can build on existing platforms, such as community-based strategies and social safety nets, for delivering parental and child services at scale to vulnerable and difficult-to-reach populations, enhancing their effectiveness and sustainability.
one of enormous change and is characterised by a high degree of plasticity in brain organisation.\textsuperscript{7,8} Advances in developmental science have also provided an understanding of the multiple and overlapping critical windows of time when development of specific capacities and abilities is most powerfully enhanced.\textsuperscript{9,10} Nurturing, caring, enriching, and protective interactions provide the early environments needed for developmental progression to occur, and protect infants and children from the negative effect of stress and adversity (panel 1). Studies from across the globe, including from Jamaica,\textsuperscript{11-13} Pakistan,\textsuperscript{14} and Turkey,\textsuperscript{15,16} have demonstrated that including elements of nurturing care in interventions significantly improves childhood development and even later adult outcomes (appendix pp 22–25). The interplay between the elements of nurturing care, the timing of experiences, and complexity of risks requires action beyond single sector interventions.

**Selection of interventions for review**

This paper provides a comprehensive update of early childhood development interventions across key sectors. Although progress has been made with early childhood development-related interventions, existing research is at different levels of maturity across sectors and distributed across numerous populations and settings. Experts from research communities in reproductive, maternal, newborn, and child health (RMNCH), nutrition, parenting, early childhood education, maltreatment prevention, and social protection worked in teams using standard methods to critically appraise the available evidence that addressed child outcomes, including: mortality; malformations, disability, and injury; nutrition and growth; and severe morbidity (panel 2). The primary focus, however, was direct measures of child development outcomes (eg, language, cognition, motor, social and emotional development, and psychosocial wellbeing). Most papers in each sector were published after the last *Lancet* Series on early childhood development—ie, from October, 2011, to April, 2015. Search strategies in each group were tailored to the existing evidence in each sector. The RMNCH and nutrition group relied on the most recent overviews of systematic reviews featuring good quality methods for all sectors, and found 15 types of interventions that show benefit on multiple outcomes including child development, based on high-quality systematic reviews (table). Many of those with effects on childhood development encompass aspects of nurturing care including parenting support and social protection, care for the caregiver, and early learning opportunities provided in or out of the home environment.

**Maternal health**

While nurturing care interventions usually begin at birth, established RMNCH interventions can reduce adverse growth and health outcomes—including stunting, low birthweight, and iron deficiency anaemia—that are strongly related to early childhood development. In our review of low-income and middle-income countries (LMICs), we identified five such RMNCH interventions during the period from conception to birth and labour that have significant effects on child development, in addition to growth, mortality, morbidity, or disability (appendix pp 2–9). These interventions include: iodine supplementation before or during pregnancy,\textsuperscript{6} antenatal corticosteroids for women at risk of preterm birth,\textsuperscript{57} magnesium sulphate for women at risk of preterm birth,\textsuperscript{18} antiplatelet drugs for women at risk of pre-eclampsia,\textsuperscript{19} and therapeutic hypothermia\textsuperscript{6} for hypoxic ischaemic encephalopathy. One review\textsuperscript{20} found mixed effects of delayed cord clamping on measures of neurodevelopment at 4 months, based on the results of one study.\textsuperscript{21} Tobacco and alcohol use are viewed as serious threats to the health of pregnant women and their children. A review of 86 randomised controlled trials showed that psychosocial programmes have been successful during pregnancy for smoking cessation, reducing low birthweight and preterm births, but evidence is limited on such interventions in LMICs.\textsuperscript{22}

**Maternal nutrition, micronutrients, and iodine supplementation**

The ability of a mother to support the health and development of her children is critically dependent on her own health and wellbeing before, during, and after pregnancy. Intrauterine growth restriction influences multiple aspects of child development and has been linked to poorer neurodevelopmental outcomes, risks of prematurity, reduced school performance, and heightened behavioural problems in children.\textsuperscript{23} Evidence suggests that linear growth is correlated across generations and short maternal stature is associated with low birthweight, stunting, childbirth complications, and increased child mortality.\textsuperscript{24} The provision of a balanced energy and protein...
during pregnancy can disrupt maternal programming, having a significant effect on children's cognitive development. Intervention during pregnancy with evidence of a severely iodine-deficient area is the only nutrition-related outcomes. Iodine supplementation in moderate-to-high-income countries prevents neural tube defects and reduces the risk of adverse birth outcomes, while folic acid fortification is associated with improved mother-infant interaction, improved cognitive development, and increased immunisation rates. Anti-depressants for treatment of antenatal depression have

diet, as well as multiple micronutrients, for women of childbearing age and expectant mothers at risk of deficiencies shows potential benefits in reducing the risk of intrauterine growth restriction, small-for-gestational-age births, and stillbirths. 

Maternal stress, depression, and mental disorders

The onset of caregiving in humans is triggered by hormonal signals beginning in pregnancy (eg, oxytocin and lactogens). Mental disorders and the timing of stress during pregnancy can disrupt maternal programming, which prepares women to respond to their infants, and can have negative effects on the fetus. Disruption to maternal programming might account for associations between maternal mental disorders, insecure mother-infant attachment, and exposure to maltreatment. Mental disorders in women, including depression and anxiety, are among the most common conditions to coexist with pregnancy and are associated with a range of negative child outcomes, including poor infant growth, children’s emotional and behavioural difficulties, and insecure attachment with caregivers. Recent evidence is emerging that paternal mental health during pregnancy can also influence the socioemotional and behavioural development of children. A systematic review of 13 trials of psychological interventions, delivered by local community health workers, for women with antenatal depression in LMICs showed positive effects on reducing maternal depression. Benefits to children included improved mother–infant interaction, improved cognitive development and growth, reduced frequency of diarrhoea episodes, and increased immunisation rates. Anti-depressants for treatment of antenatal depression have

Panel 1: Co-occurrences among bio-ecological or contextual risk factors in low-income and middle-income countries

Although there are parallels in the types of risk and promotive factors encountered by children in high-income, middle-income, and low-income countries, the limited evidence indicates that children from low-income and middle-income countries are more likely to encounter a greater number and range of risk factors and fewer promotive influences for development than poor children in high-income countries (HICs). Toxins, chronic severe malnutrition, direct exposures to armed conflict and displacement, and refugee status are risk factors that occur in LMICs, but are rarely seen in HICs. Exposure to environmental factors that reduce blood–brain barrier integrity will decrease protection of the developing brain. Poor sanitation, severe childhood diarrhoea, iron deficiency anaemia, orphan status, substandard housing, domestic violence, harsh physical punishment, and maternal depression are risk factors that occur at a higher rate in LMICs than in HICs and can be frequently amplified by exposure to conflict and population displacement. Some evidence indicates that there might be a reduced availability of promotive factors in LMICs, such as routine neonatal screening for iodine deficiency, childbirth attended by skilled health personnel, and fewer learning resources in the home. In addition to a greater range and prevalence, there are higher levels of co-occurrence among risk factors in LMICs compared with HICs. Based on analysis of UNICEF Multiple Indicator Cluster Survey data, multiple risk factors co-occur. For example, 85% of children aged 3–4 years in west and central Africa and 56% in east Asia and Pacific experience multiple risks. Data estimating risks for children living in conflict, crises, and insecure conditions are scant; however, we estimate increased levels of co-occurrence of risk factors in such situations. The findings support the application of coordination or combining of interventions, within packages, to reduce exposure to multiple risk factors. The following are specific examples of co-occurrence.

Nutritional deficiencies in infancy and early childhood are likely to occur with:

- Being born small for gestational age, or preterm, or both
- Parents who are less involved, sensitive, or responsive to the needs of the child
- Extreme poverty and food insecurity
- Suboptimal infant and young child feeding practices
- High exposure to pathogens and corresponding burden of infectious disease in infancy and childhood
- Home environments characterised as less stimulating than others
- Exposure to domestic violence

Maternal depression and anxiety are likely to coexist with:

- Preterm birth
- Low birthweight
- Poor infant growth and reduced cognitive development
- Less adequate prenatal care
- Less adequate caregiving including:
  - Suboptimal infant and child feeding practices (including not exclusive breastfeeding)
  - Insufficient communication and play to stimulate learning
  - Delayed and inappropriate care-seeking
  - Increased child morbidity
  - Increased use of harsh discipline
  - Increased family stress

Exposure to societal violence is likely to occur with:

- Child abuse and neglectful parenting
- Disruption of family or community support systems
- Disrupted and dysfunctional health systems

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Development of early childhood programmes (2014);32 and a Cochrane review of centre-based day care for children under 5 (2014).33 A meta-analysis of education programmes was conducted to determine non-cognitive developmental benefits of parenting and early childhood education programmes, as this information was not available in existing systematic reviews. The review of maltreatment prevention publications updated the Lancet article on prevention of child maltreatment (2009)34 and a systematic review of child maltreatment prevention reviews,5 by including recent reviews of maltreatment prevention interventions,33–34 such as home visiting,36 parenting training programmes,40,41 sexual abuse prevention programmes,42 universal campaigns to prevent physical abuse,43 and three narrative reviews on prevention of child maltreatment.44 The social protection literature review examined five systematic reviews that focused on the effects of social programmes, including conditional and unconditional cash transfers and microcredit schemes.26–31 After examining the systematic reviews, the literature was searched for papers that had been published since the systematic reviews. 24 new studies were included that investigated the effects of conditional cash transfers or unconditional cash transfers on measures of health, nutrition, or developmental outcomes. The search focused on research conducted in low-income and middle-income countries (LMICs), but systematic reviews based on evidence from high-income countries were included for maltreatment prevention where evidence from LMICs was either unavailable or limited. Data were double-extracted using a standardised form.

The education review was based on four recent reviews, including: the Lancet Child Development in Developing Countries Series (2011);31 a systematic review of parenting interventions published by the Annual Review of Psychology (2015);39 a literature review of parenting and early childhood programmes (2014);32 and a Cochrane review of centre-based day care for children under 5 (2014).33 A meta-analysis of education programmes was conducted to determine non-cognitive developmental benefits of parenting and early childhood education programmes, as this information was not available in existing systematic reviews. The review of maltreatment prevention publications updated the Lancet article on prevention of child maltreatment (2009)34 and a systematic review of child maltreatment prevention reviews,5 by including recent reviews of maltreatment prevention interventions,33–34 such as home visiting,36 parenting training programmes,40,41 sexual abuse prevention programmes,42 universal campaigns to prevent physical abuse,43 and three narrative reviews on prevention of child maltreatment.44 The social protection literature review examined five systematic reviews that focused on the effects of social programmes, including conditional and unconditional cash transfers and microcredit schemes.26–31 After examining the systematic reviews, the literature was searched for papers that had been published since the systematic reviews. 24 new studies were included that investigated the effects of conditional cash transfers or unconditional cash transfers on measures of health, nutrition, or developmental outcomes. The search focused on research conducted in low-income and middle-income countries (LMICs), but systematic reviews based on evidence from high-income countries were included for maltreatment prevention where evidence from LMICs was either unavailable or limited. Data were double-extracted using a standardised form. Methodological quality of systematic reviews was assessed using the AMSTAR criteria, where appropriate. More detailed information on the search strategies for each review topic can be found in the supplementary appendix.

Panel 2: Criteria for identifying relevant research

We identified peer-reviewed overviews, systematic reviews, and individual studies that focused primarily on child development outcomes, published between January, 2009, and April, 2015. We used established guidelines to search, evaluate, and synthesise the results of relevant research.6 The reproductive maternal, newborn, and child health and nutrition reviews relied primarily on six recent overviews of reviews, including: the Lancet Breastfeeding Series (2016);34 the Lancet Every Newborn Series (2014);35 the Lancet Maternal and Child Nutrition Series (2013);36 the Lancet Childhood Pneumonia and Diarrhoea Series (2013);37 the Reproductive Health 2014 supplement on essential maternal, newborn, and child health interventions;38 and the Essential Interventions for Reproductive, Maternal, Newborn, and Child Health report by the Partnership for Maternal, Newborn, and Child Health (2011).39 The education review was based on four recent reviews, including: the Lancet Child Development in Developing Countries Series (2011);31 a systematic review of parenting interventions published by the Annual Review of Psychology (2015);39 a literature review of parenting and early childhood programmes (2014);32 and a Cochrane review of centre-based day care for children under 5 (2014).33 A meta-analysis of education programmes was conducted to determine non-cognitive developmental benefits of parenting and early childhood education programmes, as this information was not available in existing systematic reviews. The review of maltreatment prevention publications updated the Lancet article on prevention of child maltreatment (2009)34 and a systematic review of child maltreatment prevention reviews,5 by including recent reviews of maltreatment prevention interventions,33–34 such as home visiting,36 parenting training programmes,40,41 sexual abuse prevention programmes,42 universal campaigns to prevent physical abuse,43 and three narrative reviews on prevention of child maltreatment.44 The social protection literature review examined five systematic reviews that focused on the effects of social programmes, including conditional and unconditional cash transfers and microcredit schemes.26–31 After examining the systematic reviews, the literature was searched for papers that had been published since the systematic reviews. 24 new studies were included that investigated the effects of conditional cash transfers or unconditional cash transfers on measures of health, nutrition, or developmental outcomes. The search focused on research conducted in low-income and middle-income countries (LMICs), but systematic reviews based on evidence from high-income countries were included for maltreatment prevention where evidence from LMICs was either unavailable or limited. Data were double-extracted using a standardised form. Methodological quality of systematic reviews was assessed using the AMSTAR criteria, where appropriate. More detailed information on the search strategies for each review topic can be found in the supplementary appendix.

For more on the Care for Child Development Package see http://www.unicef.org/earychildhood/index_68195.html

For more on Reach Up and Learn see http://www.reachupandlearn.com

Interventions from birth to 5 years of age

Parenting support

Opportunities for stimulation, responsive parent–child interactions, directed focused enrichment, early learning, and positive parenting are crucial for children’s development.40 Parenting programmes are operationally defined as interventions or services aimed at improving parenting interactions, behaviours, knowledge, beliefs, attitudes, and practices. Three recent reviews32,33,36 of parenting programmes in LMICs found positive effects on direct measures of children’s cognitive and language development across diverse policy, service delivery, and social contexts. We updated and expanded on the previous reviews by conducting a meta-analysis of non-cognitive outcomes and concluded that parenting programmes increased scores on measures of psychosocial development (standardised mean difference [SMD] 0·35, 95% CI 0·14–0·56, 13 studies) and motor development (0·13, 0·07–0·19, nine studies), in addition to child cognitive development (0·36, 0·22–0·49, 19 studies) (appendix pp 10–15). The effect of parenting programmes on child growth was not significant.

Parenting programme implementation varied in relation to dose of intervention, setting, and curriculum. The total amount of contact with parents, which ranged from less than 10 h to 120 h, did not have a clear association with the size of effect.37 Some programme models have used only home visits—eg, Roving Caregivers in Jamaica77—and others, such as Pastoral del Niño in Paraguay, have used group sessions.46 Combined group sessions and home visits in Bangladesh and Brazil produced better outcomes than did home visits alone. The most effective parenting programmes used several behaviour-change techniques, including media such as posters and cards that illustrate enrichment practices, opportunities for parent practice of play and responsive talk with their child, guidance and support for changing practices, and problem-solving strategies.43 Examples include the Care for Child Development package developed by UNICEF and WHO, and Reach Up and Learn, which provide opportunities to use multiple strategies to strengthen nurturing care by parents.48 A notable gap in published reviews is the role of fathers in promoting nurturing care and protection.49 Parenting programmes that combine nutrition and stimulation have been effective in improving child cognitive and language development outcomes.40 Taken together, the
results suggest that parenting support programmes that promote nurturing care and protection can substantially augment the positive effects of basic health and nutrition, education, and protection interventions on early child development outcomes.

### Attachment and bonding

Different brain systems enhance nurturing by supporting infant–mother attachment, as well as emotional wellbeing, learning and memory, attention, and executive functions.

Secure attachment forms with a caregiver who provides security, safety, affection, and comfort. Aspects of nurturing care during birth and labour include early initiation of breastfeeding and interventions such as Kangaroo Mother Care, which promotes thermal sufficiency in preterm infants, and early bonding. Kangaroo Mother Care has been associated with an increase in bonding indicators such as infant-mother attachment at 3 months (mean difference [MD]=6.24, 95% CI 5.57–6.91), infant growth, and rates of early exclusive breastfeeding (at 1–3 months) (risk ratio [RR]=1.20, 95% CI 1.01–1.43). Most of these evaluations were undertaken in health facilities; there is a need for research focusing on effectiveness of Kangaroo Mother Care or variants thereof when delivered at scale in community settings.

### Breastfeeding

Breastfeeding has clear short-term benefits for child health, reducing mortality and morbidity from infectious diseases, encouraging healthy food preferences, and promoting the establishment of a healthy gut microbiome. A recent review of 17 observational studies of breastfeeding presents evidence that optimal breastfeeding supports improved performance in intelligence tests in childhood and adolescence, demonstrating an intelligence quotient (IQ) increase of 3.44 points (95% CI 2.30–4.58). Findings from a 2015 analysis of the Pelotas birth cohort in Brazil also showed a dose-response association between breastfeeding duration and increased child intelligence, educational attainment, and income at the age of 30 years. The positive effect of breastfeeding was observed in one randomised trial in Belarus, in which duration of total and exclusive breastfeeding was higher in the intervention group that received the Baby-Friendly Hospital Initiative than in a control group that was not exposed to the breastfeeding counselling intervention; performance in intelligence tests at 6–5 years was also higher in the intervention group. A cohort analysis from South Africa found that exclusive breastfeeding was associated with fewer than average conduct disorders.

### Micronutrients and child feeding

Malnutrition remains a serious challenge in developing countries, undermining the survival, growth, and development of young children. Stunting and severe acute malnutrition (wasting) are often associated with concomitant micronutrient deficiencies—among these,

<table>
<thead>
<tr>
<th></th>
<th>Childhood development</th>
<th>Nutrition and growth</th>
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<th>Disability, injury, and malformations</th>
<th>Severe morbidity</th>
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<td>Periconceptional folic acid fortification or supplementation</td>
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<td>Birth interval at least 36–60 months</td>
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<td>Preconceptional diabetes care</td>
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<td>Multiple micronutrient supplementation during pregnancy</td>
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<td>Balanced protein-energy supplementation during pregnancy</td>
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<td>Intermittent preventive therapy and use of bednets for malaria prevention in mothers and children</td>
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<td>✓</td>
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<tr>
<td>Antibiotics for premature rupture of membranes</td>
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<td>-</td>
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<tr>
<td>Antibiotics for asymptomatic bacteriuria in children</td>
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</table>

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vaccine campaigns of children within high- and low-income countries.

**Table: Summary of effective interventions related to early childhood development**

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<tr>
<th>Intervention</th>
<th>Childhood development</th>
<th>Nutrition and growth</th>
<th>Mortality</th>
<th>Disability, injury, and malformations</th>
<th>Severe morbidity</th>
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<td>Detection and treatment of syphilis in pregnant mothers</td>
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<tr>
<td>Smoking cessation interventions in parents</td>
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</tr>
<tr>
<td>Continuous support during childbirth</td>
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<td>–</td>
<td>–</td>
<td>–</td>
<td>✓</td>
</tr>
<tr>
<td>Kangaroo Mother Care, skin-to-skin, cap and wrap (thermal care)</td>
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<td>✓</td>
<td>✓</td>
<td>–</td>
<td>✓</td>
</tr>
<tr>
<td>Topical emollient therapy for preterm neonates</td>
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<td>✓</td>
<td>–</td>
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<td>–</td>
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<tr>
<td>Intramuscular vitamin K for neonates</td>
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<td>–</td>
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<td>✓</td>
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<td>Handwashing behaviour and water quality improvement eg, water, sanitation, and hygiene (WASH)</td>
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<tr>
<td>Rotavirus, Hib, and pneumococcal vaccinations in children</td>
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<td>Vitamin A supplementation in children</td>
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<td>Zinc supplementation and treatment for acute diarrhoea in children</td>
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<td>✓</td>
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<td>Deworming drug treatment in children</td>
<td>–</td>
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<tr>
<td>Complementary feeding education and provision</td>
<td>–</td>
<td>✓</td>
<td>–</td>
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<td>✓</td>
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<tr>
<td>Treatment of moderate and severe acute malnutrition in children</td>
<td>–</td>
<td>✓</td>
<td>–</td>
<td>–</td>
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<tr>
<td>Interventions to prevent child maltreatment (eg, specific home-visiting and parenting programmes)</td>
<td>–</td>
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</table>

Interventions were for improving child development, nutrition and growth, mortality, disability, and morbidity in low-income and middle-income countries (LMICs), based on high-quality systematic review evidence discussed in text. Checkmarks pertain to significant pooled effect sizes. Hib=Haemophilus influenzae type B. *Most rigorous trials of interventions to prevent child maltreatment have been conducted in high-income countries, with far fewer in LMICs, and are not uniformly effective in reducing injuries, physical abuse, and neglect.

**Prevention of child maltreatment**

Family violence is increasingly recognised as a key public health problem in LMICs. Maltreatment during childhood is associated with reduced volume of both the mid-sagittal area and hippocampus, which are brain regions involved in learning and memory.7 Children who receive inadequate care, especially in the first 24 months of life, are more sensitive to the effects of stress and display more behavioural problems than do children who receive nurturing care.8 There is increasing evidence that one of the most powerful predictors of caregiving behaviour is how caregivers, especially mothers, were cared for themselves.9 Children who grow up neglected or abused by their parents, or under conditions of extreme distress within their families, are at risk of developing a host of unhealthy behaviours that affect their own lives. When these children grow up, they tend to be less equipped to take on a parenting role and are more likely to perpetuate a cycle of adverse caregiving across generations. The maltreatment prevention interventions with the best evidence that shows positive results following the intervention are selective programmes (eg, Nurse Family Partnership) characterised by intensive visits by professional home visitors and beginning prenatally, but these programmes have not been evaluated in LMICs (appendix pp 16–18). The extent to which these findings are generalised beyond the specific HICs where they have been evaluated is unknown. A systematic review of 12 parenting interventions for reducing harsh or abusive parenting in LMICs found potentially positive results on a range of parenting measures, but the quality of included trials was generally low.10 Early intervention that occurs before the onset of abusive and neglectful parenting is crucial to preventing maltreatment. One specific parenting programme, Triple P, has shown some promise in one HIC randomised trial.101 There is an urgent need for more rigorously evaluated maltreatment prevention interventions in LMICs, focusing on parenting and child outcomes, and adapted for low resource contexts. More recent reviews of early childhood development

average age 10 years; MD –3·00, –5·96 to –0·04);12 a second review on iron supplementation found an improvement in mental development (SMD 0·30, 0·15 to 0·46) and IQ (SMD 0·41, 0·20 to 0·62).13 One other review, which focused on the effect of supplementary food given to socioeconomically disadvantaged children aged from 3 months to 5 years, found that food supplements improved psychomotor development (SMD 0·41, 0·10 to 0·72), but found mixed effects on measures of cognitive development in different trials (SMD –0·40, –0·79 to 0 for Bayley II: Mental Development; and SMD 0·58, 0·17 to 0·98 for cognitive development test battery).14 Results from individual studies in Bangladesh15 and India16 suggest that responsive feeding can be effective in promoting child growth and developmental outcomes.
interventions in LMICs are suggesting associations with violence reduction and peace promotion (appendix pp 26–28).

**Out-of-home interventions**

Effects of early learning programmes, including high-quality child care, and formal and informal preschools, are well established in LMICs. On the basis of an update of an earlier published review, we found that formal and non-formal or community-based preschools in LMICs improved scores on direct measures of children's cognitive development (SMD=0.67, 95% CI 0.43–0.91, 26 studies) and psychosocial development (0.23, 0.06–0.4, five studies; appendix pp 10–15). The effects of early learning programmes on child growth were not significant and one study measuring motor effects of early learning programmes on child growth (0.23, 0.06–0.4, five studies; appendix pp 10–15). The effects of early learning programmes on child growth showed non-significant effects. The earlier review[1] found that the effects of non-formal preschools on child outcomes were typically weaker than those of formal preschools; yet some low-cost and innovative programmes, such as home-based preschool[101] and a child-to-child approach,[102] improve developmental outcomes in participants compared with non-participants. Regardless of type, programme quality is a key predictor of effectiveness; important factors of preschool quality include greater number, variety and challenging play materials, interactive or dialogic reading, classroom organisation, and instructional support. Nurturing environments, in the form of care and positive interactions and individualised attention, appear to be important in early learning programmes. A positive emotional climate at child-care centres in Chile[103] and Ecuador, [104] including individualised attention, positive affect or positive moods, and reinforcement of children's behaviours, has shown positive associations with children's early childhood cognitive and socioemotional skills.

**Social safety net interventions**

Our analysis of the systematic reviews[50–54] and the new literature (appendix pp 19–21) on social safety net interventions suggests positive effects of conditional cash transfer programmes on some child outcomes, including birthweight, illness, or morbidity. Outcomes with mixed-group or subgroup effects included height-for-age or stature, weight-for-age or underweight, and cognitive and language development. Conditional cash transfer programme participation consistently had no effects on haemoglobin concentration or prevalence of anaemia in children. In terms of indirect effects of these programmes, results were significant for effects of participation on prenatal care, growth monitoring, micronutrient supplementation, and household food consumption. It is difficult, however, to compare results across countries and contexts because programmes differ greatly. The effect of cash transfers on child development might be improved by combining social protection and early childhood development interventions. Cash transfer programmes try to address many issues at multiple levels that influence child development, such as parental and community levels, but these programmes do not directly change the factors that are linked with improving development outcomes. For example, programmes providing parental support for child development within the context of larger social protection efforts in Latin America have shown substantial benefits for child development, over and above the benefits of conditional cash transfer programmes.[105,106] Bringing these two interventions together can address both economic and nurturing care factors that impact developmental outcomes.

**Intervention packages that integrate nurturing care with sector-specific programmes**

Building on the earlier Lancet child development Series, the subsequent literature on early childhood interventions has expanded to include new longitudinal data and cohort data from LMICs. Most interventions during the period from preconception to birth focus on the physical and mental health of the mother to support a healthy pregnancy and improve birth outcomes. Interventions focusing on nurturing care and protection are usually introduced at birth; however, maternal programming for nurturing care begins during pregnancy and even earlier, with the caregiver's own childhood experiences. Evidence-based interventions during infancy that combine basic sectoral elements in health, nutrition, child and social protection, and child care and learning, with nurturing care and protection can synergistically improve child developmental outcomes. For example, including stimulation in nutrition programmes can improve developmental outcomes, which cannot be fully promoted through nutrition interventions alone.[9] Breastfeeding is an example of an intervention that combines elements of nutrition with bonding.

**Building on sectoral services**

Multi-sector approaches include coordinated services across sectors, for example water and sanitation, ideally with unifying policies. Integrated approaches refer to integration across services with shared messages and opportunities for synergy, as discussed in Paper 1 of this Series.[1] Many sectoral interventions could serve as the basis for delivery of services that link policy level strategies of cash transfer, social policies, and income generation with programmatic interventions, such as parenting support, that could benefit childhood development (appendix pp 19–21). Sectors were not included in this review, as further research is needed to examine their effects on developmental outcomes. However, associations have been noted between these sectoral interventions and such outcomes as child nutritional status, growth, and health.[98–92]
Delivering multi-sectoral intervention packages to improve childhood development

The effect of interventions on early childhood development could be improved by taking into consideration the major insights we have gained over the past decade about how human development is affected by complex and multi-faceted experiences, starting with previous generations. Based on the science of early human development, we need to conceptualise meaningful integration of interventions through a coordinated approach. In instances in which sectoral interventions were combined with elements of nurturing care and protection—eg, the Care for Child Development Programme delivered by Lady Health Workers in Pakistan—the effect of the intervention on child outcomes increased significantly. This approach allows us to intervene with the family as a unit rather than the child alone. Furthermore, there are increasing opportunities to improve interventions by combining them with nurturing care and protection, through parenting support and skills programmes.

Previous attempts at creating packages of effective interventions have focused either on the temporal relevance of the interventions (ie, packaging interventions that co-occur during the same age period of the child) or on the delivery of the programmes through the same system (eg, maternal, newborn, and child health). Although it is important to consider these factors, we also need to incorporate nurturing care and protection into the packages and tailor them to unique sets of risks and adversities facing the young child population particular to the setting.

Based on our review, we propose three illustrative packages that build on these principles and the findings. These interventions affect different aspects of nurturing care and cover numerous domains and stages in the life course (figure).

Family support and strengthening package

There are three elements of family strengthening: (access to quality services (eg, antenatal care, immunisation, and nutrition); skills building (eg, positive and responsive parenting to reduce harsh discipline and promote stimulation); and support (eg, social protection, safety networks, and family support policies). These elements increase the likelihood that families are better able to provide nurturing care for their children. Each of these elements—services, skills, and support—have independent predictive effects, however significant positive effects are seen when they are combined with programmatic interventions (eg, social protection interventions). By creating a package of the three elements of services, support, and skill building, based on the age of the child and nature of bio-ecological and contextual risk factors, developmental outcomes could be substantially improved.

Multi-generational nurturing care package

This two generation package emphasises care and protection of the mother’s and father’s physical and mental health and wellbeing, while enhancing their capacity to provide nurturing care to their child. This package combines the essential interventions of health and nutrition for mother and child—primarily delivered by the health-care system from pre-conception up until the first 1000 days of a child’s life—and the elements of care, responsibility, stimulation, and protection. This package can be further strengthened with parental leave policies as discussed in Paper 3 of this Series. While the reviews did not specifically cover situations of conflict
and violence, this package is also relevant for humanitarian contexts (panel 3). Conflict, violence, and insecurity present a complex array of adversities. In these settings families, parents, and caregivers require a package of services that addresses their needs as well as the immediate and long-term needs of their children.

**Early learning and protection package**
This set of interventions integrates the support for young children with parental support and the facilitation of teachers’ and caregivers’ ability to create a nurturing environment in early childhood centres, classrooms, and community settings for learning. This package of interventions should include nurturing care and protection by enhancing teachers’ capacities to proving a nurturing, safe, and positive emotional climate, and should include greater attention to parental support. Long-term gains have been noted when early learning packages have included parenting support and protection for young children.21 This package needs to emphasise quality and family support through parental empowerment, guidance on nutrition and care, and child protection.

There are advantages of such integrated packages in terms of delivery; for example, one location can be used for the provision of key services for young children. Identification of platforms at community, clinic, and school levels can be used to coordinate the delivery of the packages targeting population segments and families in greatest need. For example, community platforms that mobilise antenatal and postnatal home visits by community health workers complement facility-based care and promote family contact with the health-care system at crucial times. Social protection platforms provide the opportunity for identification of families in need and delivery of packages of services that link these policies with programmatic interventions. Factors that affect the selection of intervention include the age group being targeted, the expertise of the sector, coverage, or an analysis of the most efficient and effective use of resources within a service for a particular context. More evaluation is needed to codify the interventions to consolidate them into essential packages and assess effectiveness, implementation quality, and cost-benefits of integrated, inter-sectoral, and multi-sectoral approaches for early childhood development packages. Delivery of multi-sectoral services involves challenges, including limited workforce capacity, demonstration of value added for including programmatic interventions of nurturing care, and political will. Some of these challenges are discussed in Paper 3 of this Series.109

**Future research areas**
Although there has been progress in the understanding of what interventions work, there are major gaps in knowledge. The particular set of risks faced by children in conflict is not well understood. There is also a lack of knowledge about the effectiveness of early childhood development interventions in conflict-affected and fragile countries. We need to improve our understanding of how to: better combine interventions through robust assessment of intervention outcomes and evaluations of integrated parenting, responsive care, stimulation, mental health, education and protection interventions that could be delivered through community platforms; use technology-based platforms to deliver effective interventions (appendix pp 29–31); and how to scale up using evidence-based approaches.

**Conclusion**
In this paper we call for meaningful integration across sectoral interventions, through programmatic packages that promote nurturing care and protection to improve developmental outcomes. We also call for better integration of evidence-based interventions within health-care and nutrition sectors. The results of our literature review suggest that successful, smart, and sustainable interventions to improve developmental outcomes need to: promote nurturing care and protection; be implemented as packages that target multiple risks; be applied at developmentally appropriate times during the life course; be of high quality; and build on existing delivery platforms to enhance feasibility of scaling up and sustainability. We have proposed illustrative packages that meet these requirements. The nature of these interventions will continue to progress as new understanding of early human development emerges. Although questions remain about scaling up interventions at a population level, as discussed by Richter and colleagues in Paper 3 of this Series,108 we are now at a historic juncture; the evidence is clear about what
needs to be done to improve the wellbeing of future generations, and the political commitment to this is strong, as expressed by the adoption of the Sustainable Development Goals. The science is clear and the evidence convincing that our earliest experiences matter; the Sustainable Development Goals provide a crucial opportunity for implementation. We must draw on this knowledge to take action to support parents, caregivers, and families in providing the nurturing care and protection that young children deserve.

Contributors
PRB and SJL conceptualised the review in consultation with the Early Childhood Development Series Steering Committee and wrote the first draft of the Series paper with substantial inputs from RP, ZAB, RP-R, MFG, and TV led the review of MCNH and nutrition interventions. NR, PI, and AKY led the review of early childhood education and parenting interventions. HM led the review of child maltreatment prevention interventions. LCHF led the review of social protection interventions. SGM, AC, AF, and VGM contributed to the scientific literature review of nurturing care and human development. TDW and HVA reviewed the literature on cumulative and protective risk factors. All authors and members of the review groups saw successive drafts of the paper and provided input. PRB, SJL, and KP prepared the final version of the Series paper, which all authors approved. PRB had final responsibility for the decision to submit for publication.

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Declaration of interests
PRB is employed by UNICEF. JFL has received several contracts, gifts, and grants focused on the impact of early child development programmes from UNICEF, the Anne Coxwell Egitim Vakfı (ACEV, Mother-Child Education Foundation), the UBS Optimus Foundation, and the Open Road Alliance. The other authors declare that they have no conflicts of interest. The views expressed are those of the authors and not necessarily those of UNICEF. Bill & Melinda Gates Foundation, and Conrad N Hilton Foundation. As corresponding author, PRB states that she had full access to all data and final responsibility to submit for publication.

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