

# Iron supplementation to children

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**Date:** 30 November 2021

## Description of condition and intervention

Iron deficiency is one of most common causes of anemia in children and an important cause of disease burden in low- and middle-income countries (LLMICs). Iron deficiency among children is often caused by inadequate intake or absorption of dietary iron, increased need in periods of growth, or infection by intestinal helminths. Iron is an essential nutrient for development and cell growth in the immune and neural systems, as well as in regulation of energy metabolism and exercise. The economic costs of iron deficiency anemia from annual physical productivity losses have been calculated to be around US\$ 2.32 per capita, or 0.57% of gross domestic product in LLMICs. The WHO has consistently recommended oral iron supplementation as one of the interventions that can reduce the prevalence of anemia. Iron is required for the survival and virulence of many pathogens. Concerns have been expressed on a possible increased risk of malaria with iron interventions in malaria-endemic areas, particularly among iron-replete children. The intervention assessed in this Evidence Brief is iron supplementation in children, 10–12.5 mg elemental iron given as drops/syrups daily 6-23 months and 25 mg elemental iron per week intermittently 24-59 months. (*Source: WHO 2016, WHO 2011*). This evidence brief assesses effects and costs for three interventions being analyzed in FairChoices: DCP analytical tool (For an overview of other interventions, see appendix below and the separate evidence briefs for these):

|                        |  |
|------------------------|--|
| <b>NUTR01-02-01-01</b> | <b>Daily iron supplementation for children 6 to 23 months</b>        |
| <b>NUTR01-02-01-02</b> | <b>Daily iron supplementation in children health center</b>          |
| <b>NUTR01-02-02</b>    | <b>Intermittent iron supplementation in children (24 -59 months)</b> |

## International guidelines for iron supplementation

| Organization             | Indications/recommendations   | Applicability in LIC & Lower MIC settings    |
|--------------------------|---|--|
| <a href="#">WHO 2016</a> | Daily iron supplementation: 10-12.5mg elemental iron supplementation in infants and young children (age 6-23 months) given as drops/syrups for three consecutive months in a year. This 10-12.5mg elemental iron corresponds to 50– | If prevalence of anemia in infants and young |

NUTRITION: Provision of iron supplementation to children  
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 (DCP4 ID: NUTR02-01 and NUTR02-02)  
 Cluster: Nutrition

**FairChoices**  
 DCP Analytic Tool

|                          |   |   |
|--------------------------|---|---|
|                          | 62.5 mg of ferrous sulfate heptahydrate, 30–37.5 mg of ferrous fumarate or 83.3–104.2 mg of ferrous gluconate.  | children is 40% or higher   |
| <a href="#">WHO 2011</a> | Intermittent iron supplementation: Every week 25 mg of elemental iron suggested in form of drops/syrup for three months duration, alternated with three months of no supplementation. This 25 mg of elemental iron corresponds to 75 mg of ferrous fumarate, 125 mg of ferrous sulfate heptahydrate or 210 mg of ferrous gluconate. | If prevalence of anemia in preschool children (24-59 months) is 20% or higher |

Source: Organization GWH. Guideline: Daily iron supplementation in infants and children 2016

## Intervention attributes

### Type of interventions & delivery platform

Table 1: Type of interventions & delivery platform

| Intervention                      | Type       | Delivery platform |
|-----------------------------------|------------|-------------------|
| Daily iron supplementation        | Prevention | Community         |
| Daily iron supplementation        | Prevention | Health center     |
| Intermittent iron supplementation | Prevention | Community         |

## Equity

In addition to considerations like cost-effectiveness and health systems factors, dimensions of equity can be relevant for priority setting. The opportunity for a long and healthy life varies according to the severity of a health condition that individuals might have, so there are inequities in individuals' opportunities for long and healthy lives based on the health conditions they face. Metrics used to estimate the severity of illness at an individual level can be used to help prioritize those with less opportunity for lifetime health. FairChoices: DCP

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Analytics Tool uses Health adjusted age of death (HAAD), which is a metric that estimates the number of years lived from birth to death, discounting years lived with disability. A high HAAD thus represents a disease less severe in terms of lifetime health loss, while a low HAAD represents a disease that is severe on average, causing early death or a long period of severe disability. It is also possible to estimate the distribution of HAAD across individuals with a health condition. FairChoices shows for each intervention an average HAAD value of the conditions that are affected by respective interventions that have health effects. Additionally, a plot shows HAAD values for around 290 conditions (Johansson KA et al 2020).

## Population in need of interventions

Table 2: Population in need of interventions

| Intervention                               | Treated population   |  | Affected population  |  | Disease state addressed |
|--|--|--|--|--|-------------------------|
|  | Treated age  | Treated fraction   | Affected age   | Affected fraction  |                         |
| Daily iron supplementation (Community)     | 0 to 1 years (From six months of age when breastfeeding is not adequate) | According to Henriksen et al. Treated fraction will be 0.5 | 0 to 1 years (From six months of age when breastfeeding is not adequate) | Those with the condition, both genders: According to Henriksen et al. Treated fraction is 0.5 for community level (Unpublished work in progress)     | Dietary iron deficiency |
| Daily iron supplementation (Health center) | 0 to 1 years (From six months of age when breastfeeding is not adequate) | According to Henriksen et al. Treated fraction will be 0.5 | 0 to 1 years (From six months of age when breastfeeding is not adequate) | Those with the condition, both genders: According to Henriksen et al. Treated fraction is 0.5 for health center level (Unpublished work in progress) | Dietary iron deficiency |

|                                   |                                  |   |                                  |   |                         |
|-----------------------------------|----------------------------------|---|----------------------------------|---|-------------------------|
| Intermittent iron supplementation | 2 to 5 years (From 24-59 months) | According to Henriksen et al. Treated fraction will be 1(all) | 2 to 5 years (From 24-59 months) | Those with the condition, both genders:<br>According to Henriksen et al. Treated fraction is 1 for community level (Unpublished work in progress) | Dietary iron deficiency |
|-----------------------------------|----------------------------------|---|----------------------------------|---|-------------------------|

## Disease stage addressed

Delay or prevent development of iron deficiency anemia

## Intervention effectiveness and safety

Table 3: Effectiveness and safety of daily iron supplementation in 6-23 months children

| Effect of intervention   |   | Certainty of evidence |
|--|---|-----------------------|
| Prevalence of <b>iron deficiency</b> (population: 6-23 months children)<br>Pasricha et al 2013 found that with daily iron supplementation relative risk of iron deficiency anemia was 0.14 (95% CI: 0.10–0.22) | Pasricha et al 2013.<br>Demonstrates high evidence of daily iron supplementation for prevention of iron deficiency anemia, but high uncertainty of evidence on child growth | ⊕⊕⊕⊕<br>Very high     |

Table 3.1: Effectiveness and safety of intermittent iron supplementation in 24-59 months children

| What happens?  | No intervention | With intervention | Certainty of evidence | Transferability of evidence |
|--|-----------------|-------------------|-----------------------|-----------------------------|
| Effects on children 24-59 months due to the condition  |                 |                   |                       |                             |
| Prevalence<br>De-Regil et al 2011 found that children receiving intermittent iron supplementation had a lower risk of anemia (RR=0.51, 95% CI: 0.37 to 0.72) |                 |                   | ⊕⊕⊕⊖<br>Moderate      |                             |

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## Model assumptions

Table 4.0: Summary of model parameters and values used in FairChoices – DCP Analytical Tool

| Category   | Model parameter   | Notes                          |  |
|--|---|--------------------------------|--|
| Intervention   | Daily iron supplementation                                  |                                |  |
| Cost parameters  |   |                                |  |
| Treated population<br>Treated fraction<br>Treated fraction age | Children with dietary iron deficiency<br>0.5<br>0 to 1 year | Global Burden of Disease study |  |
| Effect calculation   |   |                                |  |
| Affected Population  | Children with dietary iron deficiency                       |                                |  |
| Affected gender  | Both gender   |                                |  |
| Affected fraction age  | 0 to 1 year   |                                |  |
| Affected fraction  | 0.5   |                                |  |
| Comparison   | Placebo   |                                |  |
| Prevalence Reduction (RRR)                                     | 0.86  |                                |  |

Table 4.1: Summary of model parameters and values used in FairChoices – DCP Analytical Tool

| Category   | Model parameter   | Notes                          |  |
|--|---|--------------------------------|--|
| Intervention   | Intermittent iron supplementation                           |                                |  |
| Cost calculation   |   |                                |  |
| Treated population<br><br>Treated fraction<br>Treated fraction age | Children with dietary iron deficiency<br>0.5<br>2 to 5 year | Global Burden of Disease study |  |
| Effect calculation   |   |                                |  |
| Affected Population  | Children with dietary iron deficiency                       |                                |  |
| Affected gender  | Both gender   |                                |  |
| Affected fraction age  | 2 to 5 year   |                                |  |
| Affected fraction  | 0.5   |                                |  |
| Comparison   | Placebo   |                                |  |
| Prevalence Reduction (RRR)   | 0.49  |                                |  |

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## Intervention Cost

The total unit cost is estimated to be **USD 4.87** (Year: 2020) per case per child by giving daily iron supplementation on a community-based level for the prevention of dietary iron deficiency anemia according to *Henriksen et al. (Unpublished work in progress)*

The total unit cost is estimated to be **USD 5.24** (Year: 2020) per case per child by giving daily iron supplementation on a health center level for the prevention of dietary iron deficiency anemia according to *Henriksen et al. (Unpublished work in progress)*

The total unit cost is estimated to be **USD 3.59** (Year: 2020) per case per child by giving intermittent iron supplementation on a community-based level for the prevention of dietary iron deficiency anemia according to *Henriksen et al. (Unpublished work in progress)*

## References

WHO 2016: Guideline: Daily iron supplementation in infants and children. Geneva: World Health Organization; 2016.

Pasricha et al 2013: Pasricha SR, Hayes E, Kalumba K, Biggs BA. Effect of daily iron supplementation on health in children aged 4-23 months: a systematic review and meta-analysis of randomised controlled trials. *Lancet Glob Health*. 2013;1(2):e77-e86.

De-Regil et al 2011: De-Regil LM, Jefferds ME, Sylvetsky AC, Dowswell T. Intermittent iron supplementation for improving nutrition and development in children under 12 years of age. *Cochrane Database Syst Rev*. 2011(12):CD009085.

Henriksen ES, Økeland J, Malawim O, Said S, Kaur G, Rava` MS, et al. Economic evaluation of nutritional interventions in Zanzibar: An analysis using FairChoices – DCP analytic tool.(Work in progress)

## Appendix

### Literature Review for effectiveness & safety

This literature search is an example of level 4 evidence(metaanalysis) for intervention inputs taken from DCP3. (Despite low significant level for efficacy)

Level of evidence of efficacy studies:

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1. Low (expert opinions, case series, reports, low-quality case control studies)
2. Moderate (high quality case control studies, low quality cohort studies)
3. High (high quality cohort studies, individual RCTs)
4. Very high (Multiple RCTs, metaanalysis, systematic reviews, clinical practice guidelines)

An overview of all NUTR interventions in FairChoices-DCP analytical tool (Interventions assessed in this evidence brief are marked in bold)

|                         |   |
|-------------------------|---|
| NUTR01-01               | Daily Iron Folic acid supplementation (pregnant women)                        |
| NUTR01-02               | Calcium supplementation, pregnancy  |
| NUTR01-03<br>households | Food and caloric supplementation to pregnant women in insecure                |
| NUTR01-04-02            | Promotion of breastfeeding and/ or complementary feeding                      |
| NUTR01-05               | Intermittent Iron-folic acid supplementation (Menstruating women)             |
| NUTR01-06               | Food to non-pregnant women in insecure households                             |
| <b>NUTR01-02-01-01</b>  | <b>Daily iron supplementation for children 6 to 23 months</b>                 |
| <b>NUTR01-02-01-02</b>  | <b>Daily iron supplementation in children health center</b>                   |
| <b>NUTR01-02-02</b>     | <b>Intermittent iron supplementation in children (24 -59 months)</b>          |
| NUTR01-02-03            | Vitamin A supplementation to children 6 to 59 months                          |
| NUTR01-02-04            | Zink to children 6 to 59 months   |
| NUTR01-02-05            | Food to children, if below basic food poverty line                            |
| NUTR01-03-01            | Management of severe acute malnutrition without medical complications         |
| NUTR01-03-02            | Management of severe acute malnutrition associated with medical complications |