Effects of iron supplementation and nutrition education on haemoglobin, ferritin and oxidative stress in iron-deficient female adolescents in Palestine: randomized control trial

Marwan Jalambo $^{\scriptscriptstyle 1,2}$, Norimah Karim $^{\scriptscriptstyle 2}$, Ihab Naser $^{\scriptscriptstyle 3}$ and Razinah Sharif $^{\scriptscriptstyle 2,4}$

¹Nutrition and Public Health Programme, Academic Department, Palestine Technical College, Palestine. ²Nutrition Science Programme, Faculty of Health Science, University Kebangsaan, Malaysia, Selangor, Malaysia. ³Clinical Nutrition Department, Faculty of Applied Science, Al-Azhar University, Gaza, Palestine. ⁴Centre for Healthy Ageing and Wellness, Faculty of Health Sciences, University Kebangsaan Malaysia, Selangor, Malaysia (Correspondence to: Razinah Sharif: razinah@ukm.edu.my).

Abstract

Background: Iron deficiency and iron-deficiency anaemia are associated with oxidative stress, but their role is largely unclear. Information is scarce on the effects of iron supplementation on biomarkers of oxidative stress in humans.

Aims: This study evaluated the effectiveness of iron supplementation and nutrition education on improving the levels of haemoglobin and ferritin, and decreasing oxidative stress among iron-deficient female adolescents in Gaza, Palestine.

Methods: A total 131 iron-deficient female adolescents were recruited and allocated randomly into 3 different groups. The iron supplementation group (A) received 200 mg of ferrous fumarate weekly during the 3-month intervention, the iron supplementation with nutrition education group (B) received iron supplements with nutrition education sessions, and the control group (C) did not receive any intervention. The levels of haemoglobin, ferritin and malonyl dialdehyde were measured at baseline, after 3 months (at which point the intervention was stopped), and then 3 months later. Trial registration number: ACTRN12618000960257.

Results: Haemoglobin levels increased significantly after supplementation in both groups A and B. At the follow-up stage (3 months after stopping the intervention), iron and haemoglobin levels in group B continued to increase and malonyl dialdehyde decreased. In Group A, haemoglobin, ferritin and malonyl dialdehyde levels decreased after 3 months of stopping the intervention. No changes were seen in Group C.

Conclusions: A nutrition programme should be adopted and integrated into comprehensive intervention programmes to target iron-deficiency anaemia among female adolescents in Palestine.

Keywords: Adolescent; female; anaemia, iron deficiency; oxidative stress; dietary supplements, Gaza

Received: 06/01/17; accepted: 21/06/17

Citation: Jalambo M; Karim N; Naser I; Sharif R. Effects of iron supplementation and nutrition education on haemoglobin, ferritin and oxidative stress in iron-deficient female adolescents in Palestine: randomized control trial. East Mediterr Health J. 2018;24(6):560–568. https://doi.org/10.26719/2018.24.6.560

Copyright © World Health Organization (WHO) 2018. Some rights reserved. This work is available under the CC BY-NC-SA 3.0 IGO license (https:// creativecommons.org/licenses/by-nc-sa/3.0/igo).

Introduction

Iron deficiency is the most common form of nutritional disorder worldwide, affecting more than 2 billion people globally (1). Iron deficiency is not the only cause of anaemia but where anaemia is prevalent, iron deficiency is the most common cause (2). Most adolescents suffer from iron deficiency and its adverse health effects (2). In Gaza, Palestine, the prevalence of anaemia was reported as 33.3% (3), while the prevalence of iron deficiency was 23.6% among adolescents (4). Adolescents of today are the adult population of tomorrow and therefore their health and wellbeing are important (5).

Pregnancy increases the risk of iron deficiency and the approach to iron-deficiency anaemia in pregnancy has changed recently, from providing nutritional supplements during pregnancy to taking steps earlier to ensure that women, especially adolescents, have adequate iron stores before conception (6). Previous studies have found that knowledge is one of the first steps to changing behaviour. Nutrition knowledge is therefore an essential basis for good dietary habits (7,8). Conversely, a lack of knowledge is a risk factor for malnutrition (9). Nutrition education programmes are needed to increase awareness in female adolescents about anaemia (10). Moreover, health education has proven to be very effective and has resulted in a substantial improvement in iron levels and nutrition knowledge (11,12).

Oxidative stress is defined as an imbalance between the oxidation and anti-oxidation systems, resulting in the excessive production of reactive oxygen species (13,14). Reactive oxygen species of erythrocytes is one of the principal causes of anaemia (15). Red blood cells are subjected to continuous oxidative stress during their lifetime. They are particularly susceptible to oxidative damage from high content of unsaturated fatty acid chains in the lipid bilayer combined with high oxygen

levels (16).

As far as we know, no studies have been carried out to assess the effects of ferrous fumarate supplementation on the biomarkers of oxidative stress in female adolescents with iron deficiency and iron deficiency anaemia. Therefore, the aim of this study was to evaluate the effects of iron supplementation and nutritional education on haemoglobin (Hb) and ferritin levels and oxidative stress in female adolescents (aged 15–19 years) with iron deficiency and iron-deficiency anaemia in Gaza, Palestine.

Methods

Study design and setting

This randomized control trial was conducted in Gaza, Palestine on female adolescents aged 15–19 years with iron deficiency and iron-deficiency anaemia (mild and moderate). The study was conducted from 25 October 2015 to 25 April 2016.

Sample size and selection

The sample size was calculated based on the formula for 2 means:

$$n = [(Z\alpha/2 + Z\beta)^2 \times 2(\sigma^2)]/(\mu_1 - \mu_2)^2$$

Where:

 μ_1 = mean change in Hb after 3 months from iron supplements = 1.23 g/dL (17).

 μ_2 = expected mean change in Hb after 3 months from iron supplements = 2 g/dL

 σ = standard deviation = 1.195

 $Z\alpha/2$: at 5% level of significance = 1.96

Z β : for a power of 80% = 0.84.

The calculation gave 38 respondents in each group. However, assuming a 15% drop-out from the trial, the number of participants was increased to 45 per group.

Eligibility

Participants were eligible for inclusion in the trial if they were unmarried and not pregnant. Participants were excluded if they: had severe anaemia (Hb < 8 g/dL); were suffering from acute or chronic infections that could affect their Hb and ferritin levels at the time of the blood sampling; had anaemia other than iron-deficiency anaemia; were underweight; were on medication; or were diagnosed with thalassemia trait.

Group allocation and intervention

This intervention programme was part of a 2-phase study in which anaemia and iron deficiency status were assessed among female adolescents attending secondary schools in Gaza (first phase).

In the first phase, a sample of female adolescents aged 15–19 years was selected from secondary schools in Gaza. As there are 5 governorates in Gaza, 1 school was selected from each governorate out of a total of 145 schools. Schools were listed by governorate and 1 was randomly

selected from each governorate. Similarly classes were selected randomly from grades 10, 11 and 12 in the selected schools, 1 or 2 class(es) from each grade according to the population of the governorate. All the girls in the selected classes (330 girls) were assessed for iron deficiency and iron-deficiency anaemia. Preliminary screening for iron deficiency and iron-deficiency anaemia (Hb < 12 g/dL and ferritin < 15 μ g/L) was done through vein blood samples. The girls were also interviewed in school by a female research assistant about their socioeconomic status and to determine if they met the eligibility criteria.

A total of 177 girls (54%) had iron deficiency or irondeficiency anaemia. Of these, 43 did not meet the eligibility criteria and were excluded. Therefore 135 students with iron deficiency or iron-deficiency anaemia were invited to participate in the second phase of the study. None was taking taking any supplements. Those who agreed (n = 131) were randomized into 3 groups (Figure 1).

- Group A (iron supplementation) included 45 participants who received 200 mg of ferrous fumarate once weekly for 3 months; this was given by the researchers during school hours.
- Group B (iron supplementation with nutrition education) included 44 participants who received iron supplements (ferrous fumarate) once weekly and 9 nutritional education sessions (1.5 hours/session) for 3 months.
- Group C (controls) included 42 participants who did not receive any intervention throughout the study period.

The purpose of the nutrition education was to teach the participants about the importance of good nutrition, with an emphasis on iron deficiency and iron-deficiency anaemia. The nutrition education intervention consisted of lectures, posters, videos, booklets, and brochures. Nutrition lectures were delivered by the researchers using simple methods and vocabulary to present topics on nutrition in Arabic such as food groups, the food pyramid, a balanced diet, iron absorption enhancers and inhibitors, good sources of iron, and ways to improve the absorption of iron from foods.

After 3 months of the intervention, the supplementation and education sessions were stopped for Groups A and B, and the groups were followed up after a further 3 months.

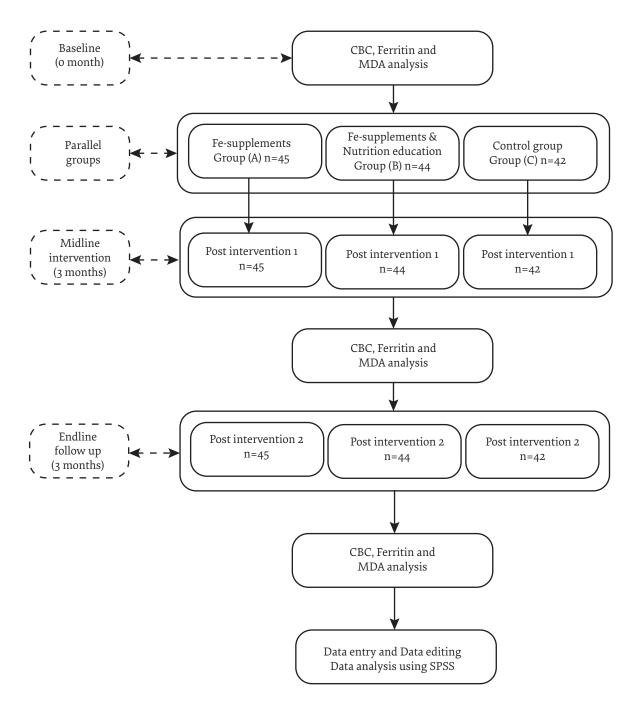
After the study was completed, the intervention was offered to the girls in the control group; they were given a packet of iron supplement tablets (20 tablets) and 2 sessions of nutrition education.

Measurements

Three blood assessments were done: at baseline (25 October 2015), after 3 months (first post-intervention, 25 January 2016), and after 6 months of baseline (second post-intervention (25 April 2016).

Complete blood count, serum ferritin and malonyl dialdehyde (MDA) levels were measured. About 5 mL of

Figure 1 Flow diagram of the progress of the groups through the stages of the study, at baseline, and after 3 and 6 months of the intervention (CBC = complete blood count, MDA = malonyl dialdehyde, Fe = Iron)



venous blood were drawn (0.5 mL in EDTA tubes and 3 mL in serum evacuated tubes) from each participant. All the samples were placed in tube racks and packed in an appropriate ice container (4 °C) and sent to a laboratory accredited by the Palestinian Ministry of Health. The samples were analysed on the same day. Blood samples were coded so assessors were unaware of the group a blood sample was from.

Anaemia was assessed by measuring Hb concentration in the complete blood count analysis (Horiba ABX Micros ES 60, France). Iron status was assessed by measuring the serum ferritin levels (Chemistry Autoanalyzer, Model: BS-120, Mindray Bio-Medical Electronic Co. Ltd, China) in duplicate, and recording the average results. An MDA adduct competitive ELISA kit (OxiSelectTM, Cell Biolabs, Canada) was used to measure MDA levels.

Body weight of each participant was measured by using a calibrated scale (Seca model 750 1017009, Germany. Students were weighed barefooted to the nearest 0.5 kg. Standing height also was measured without shoes to the nearest 0.5 cm with a stadiometer (Seca body meter 206, Germany), with the shoulders in a relaxed position and the arms hanging freely. The body mass index (BMI) for age was be calculated using the Anthro Plus program of the Centers for Disease Control and Prevention (version 7.1.5.2). Duplicate measurements of weight and height of the students were taken and the mean of value was determined.

Statistical analysis

All analyses were conducted using *SPSS*, version 22. Repeated measures analysis of variance (one-way repeated measures ANOVA) was used to evaluate changes in all the continuous variables (Hb, ferritin and MDA) over the study period. Repeated measures ANOVA measured the changes between the groups, within the group, and time and group interactions. A *P*-value < 0.05 was considered statistically significant.

Ethical considerations

Before data collection, written permission to carry out the study was obtained from the Helsinki Committee in Palestine and the Ministry of Education in Palestine (reference number PHRC/HC/3^/14). After reviewing the study protocol, the Human Ethics Committee of Universiti Kebangsaan Malaysia gave its ethical approval (reference number UKM1.5.3.5/244/NN-025-2015).

In keeping with social and cultural norms, a female interviewer was hired. Written consent was obtained from all the respondents if they were aged ≥ 18 years or from their parents if they were under 18 years. It was explained that they were free to withdraw from the study at any time.

Trial registration

This trial is registered under ACTRN number: AC-TRN12618000960257.

Results

Descriptive results of the study sample

Table 1 describes the baseline sociodemographic characteristics of the participants (age, region, educational grade), BMI, and blood analysis. Ferritin and MDA levels in group B were slightly higher than in the other 2 groups but no statistically significant differences were found. Thus, no adjustments were made in the outcome analysis on the basis of any of the variables.

Effect of nutrition education and supplementation on Hb, ferritin and MDA status

The main objective of the trial was to determine the effectiveness of iron supplementation and nutrition education on the levels of Hb, ferritin and MDA in female adolescents aged 15–19 years. The effect of the intervention was assessed using the 2 dimensions of repeated measure ANOVA; firstly, the within group effect to determine whether there was a difference in the blood parameters (Hb, ferritin and MDA), and secondly, to assess the interaction of treatment and time together.

Table 2 shows that the difference in the mean Hb concentration was statistically significant between within the same group at different intervals (P < 0.001). A post-hoc test using the Bonferroni correction showed that there was a statistically significant difference between all the timelines. The mean ferritin concentration also differed significantly between the timelines (P < 0.001). A post-hoc test using the Bonferroni correction showed that there was a statistically significant difference between the baseline and midline, and the baseline and end-line, but there was no statistically significant difference between the midline and end-line. There was a statistically significant difference between the 3 timelines in MDA concentrations also (P < 0.001). The Bonferroni test showed that the differences were statistically significant between the baseline and midline, and the midline and end-line, but not between the baseline and end-line.

The results of the mean ferritin concentration were statistically significant between the groups at midline and end-line (post-intervention and post-intervention 2) (P < 0.001).

Figures 2, 3 and 4 explain the differences in the effect of nutrition education intervention between the 3 groups after 6 months. The trend in the plot showed the adjusted mean levels (estimated marginal mean) of Hb, ferritin and MDA for zero months (baseline), 3 months (intervention period) and 6 months (follow-up without intervention). The mean Hb, ferritin and MDA levels

| Table 1 Characteristics and blood measurements of the groups at baseline (randomization) | | | | | | | | | |
|--|---------------------|---------------------|---------------------|---------------------|---------|--|--|--|--|
| Variable | Group A (n = 45) | Group B (n = 44) | Group C (n = 42) | Test value | P-value | | | | |
| Age (years) [mean (SD)] | 16.11 (0.89) | 16.37 (0.78) | 16.51 (0.89) | 2.422 ^a | 0.09 | | | | |
| Grade (10th, 11th, 12th) [No.] | 18, 12, 15 | 16, 10, 18 | 9, 15, 18 | 4.452 ^b | 0.34 | | | | |
| Region (NG, GC, MG, KH, RF) [No.] | 10, 10, 10, 9, 6 | 4, 15, 14, 6, 5 | 12, 10, 10, 7, 3 | 7.915 ^b | 0.44 | | | | |
| Body mass index (kg/m²) [mean (SD)] | 21.74 (3.88) | 22.63 (3.57) | 22.49 (4.13) | 20.405 ^a | 0.50 | | | | |
| Haemoglobin (g/dL) [mean (SD)] | 11.52 (0.96) | 11.45 (1.18) | 11.73 (0.89) | 0.800ª | 0.41 | | | | |
| Ferritin (µg/dL) [median (IQR)] | 8.60 (4.0) | 10.50 (4.5) | 9.50 (6.0) | 1.471 ^b | 0.47 | | | | |
| Malonyl dialdehyde (Pmol/mL) [median (IQR)] | 83.0 (43.0) | 92.0 (31.0) | 83.50 (63.0) | 1.076 ^b | 0.58 | | | | |

^aOne-way ANOVA F-test. ^bKruskal–Wallis chi-squared test.

NG: North Gaza, GC: Gaza City, MG: Middle Governorate, KH: Khanyounis, RF: Rafah.

SD = standard deviation, *IQR* = interquartile range.

| Blood analysis | Intervention groups | | | Mixed repeated-measure ANOVA | | |
|-----------------------------|---------------------|---------------------|---------------------|------------------------------|-----------------------|--|
| | Group A (n = 45) | Group B (n = 44) | Group C (n = 42) | Time effect | Interaction effect | |
| | Mean (SD) | Mean (SD) | Mean (SD) | ղ ք² (P-value) | ղ p² (P-value) | |
| Haemoglobin (g/dL) | | | | 0.373 (< 0.001)* | 0.177 (< 0.001)* | |
| Baseline | 11.52 (0.96) | 11.45 (1.18) | 11.73 (0.89) | | | |
| Midline | 12.46 (0.64) | 12.09 (1.02) | 11.98 (0.93) | | | |
| End-line | 12.04 (0.67) | 12.17 (0.94) | 11.74 (0.84) | | | |
| Ferritin (µg/L) | | | | 0.511 (< 0.001)* | 0.331 (< 0.001)* | |
| Baseline | 9.92 (3.16) | 9.19 (2.92) | 9.70 (3.48) | | | |
| Midline | 15.92 (5.43) | 14.75 (3.69) | 10.13 (5.25) | | | |
| End-line | 13.94 (4.61) | 15.73 (3.99) | 10.14 (4.14) | | | |
| Malonyl dialdehyde (Pmol/L) | | | | 0.437 (< 0.001)* | 0.186 (< 0.001)* | |
| Baseline | 90.14 (47.88) | 91.5 (27.33) | 96.23 (40.81) | | | |
| Midline | 129.8 (61.89) | 116.7 (35.33) | 112.0 (39.79) | | | |
| End-line | 91.2 (43.60) | 78.64 (25.62) | 107 (39.96) | | | |

*Significant at P < 0.05.

 ηp = partial eta, SD = standard deviation.

were almost equal for the 3 groups at the baseline. After 3 months of the intervention, the mean Hb, ferritin, and MDA levels increased in the 2 intervention groups but there were no changes in the control group. After 6 months of the baseline, the Hb and ferritin levels still increased in the nutrition education group, while the MDA level decreased in the same group. On the other hand, for the group that received iron supplementation alone, there was a decrease in the Hb and ferritin levels at the follow-up stage.

Discussion

This study evaluated the effects of an iron supplementation and nutrition education intervention on Hb, serum ferritin and MDA levels as indicators of oxidative stress among female adolescents with iron deficiency and iron-deficiency anaemia in Gaza, Palestine.

After 3 months of the intervention, mean Hb concentration had increased significantly in groups A and B. This finding is consistent with the results of studies in Malaysia and Ghana in 2012 (17,18).

The results of our study showed that weekly iron supplementation (200 mg of ferrous fumarate) led to an increase in Hb levels from a mean of 11.52 (SD 0.96) g/ dL to 12.46 (SD 0.64) g/dL after 3 months. This result is consistent with that of previous studies. For example, a study in India found an increase in mean Hb levels from 10.51 (SD 0.35) g/dL to 12.49 (SD 0.65) g/dL after 3 months of weekly iron supplementation with folic acid among

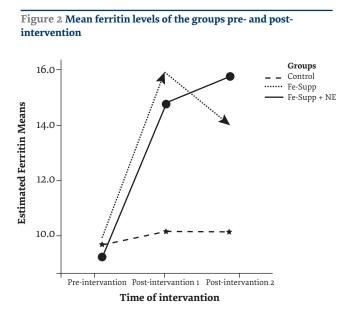


Figure 3 Mean haemoglobin levels of the groups pre- and postintervention

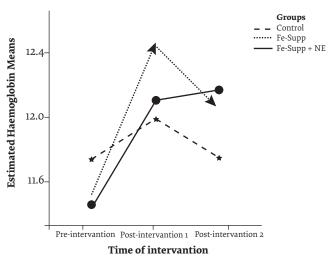
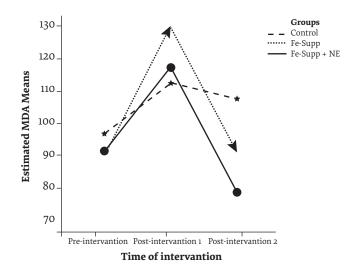


Figure 4 Mean malonyl dialdehyde levels of the groups pre- and post-intervention



female adolescents (19). Another study in India found the mean Hb level among female adolescents with anaemia increased from 10.80 g/dL to 12.65 g/dL after 3 months of once weekly iron supplementation (20). Similarly, the study in Ghana among students with anaemia aged 6–11 years reported an increase in mean Hb levels from 11.38 (SD 0.15) g/dL to 11.63 (0.13) g/dL after 10 weeks of a 5-day weekly iron supplementation (65 mg of ferrous fumarate) (17). A study in Peru also reported an increase in mean Hb levels with twice-weekly iron supplementation (60 mg of ferrous sulfate)—from 11.39 (SD 0.05) g/dL to 12.07 (SD 0.01) g/dL—among female adolescents with anaemia aged 12–18 years after 4 months (21).

No significant differences in Hb levels were seen between groups A, B and C (P = 0.57) before the intervention. At 3 months, a significant increase in the Hb levels was seen in groups A and B but not the control group. At 6 months, Hb levels in group A decreased, but in group B, Hb levels continued to increase, though at a lower rate, until the end of the study. These results are consistent with the results of a study conducted among adolescents in Malaysia (17). The improvement in the Hb levels in the participants in the iron supplementation group was most likely due to the iron supplementation, while the improved Hb levels in the participants in the iron supplementation with nutrition education group were most likely because of modifications to their dietary habits, which depended on a positive attitude and good practices.

The effect of the iron supplementation on iron levels was assessed using the mean ferritin concentration. There was a significant increase in the mean ferritin level in Groups A and B. This finding is consistent with the findings of the studies by Opoku and Menendez et al. (18,22). In our study, mean ferritin levels after 3 months of iron supplementation in group A changed from 9.9 (SD 3.16) μ g/L to 19.92 (SD 5.43) μ g/L. This change was higher than the change in a study in Malaysia where ferritin

levels increased from 34.3 (SD 2.49) μ g/L to 37.5 (SD 2.49) μ g/L among female adolescents aged 12–17 years with anaemia but not iron deficiency following weekly iron supplementation for 3 months (23). This difference in the amount of increase may be because the Malaysian study targeted non-iron-deficient female adolescents. In addition, according to the results of our study, the change in the ferritin levels after the iron supplementation was lower than the study by Opoku, in which an increase in the ferritin levels was reported from a mean of 14.17 (SD 0.64) μ g/L to 40.38 (SD 4.95) μ g/L following 5-day a week iron supplementation for 10 weeks among students aged 6–11 years with iron-deficiency anaemia (18). This difference may have been due to the age differences between the studied samples.

Information is scarce on the effects of iron supplementation on the biomarkers of oxidative stress in humans (24). The lack of epidemiological data on oxidative damage in healthy human populations is a serious gap in the distribution, correlation and causative factors of oxidative damage (25). Iron deficiency and irondeficiency anaemia are associated with oxidative stress, but their role in initiating stress is largely unclear. Also, oxidative stress induced by iron deficiency and irondeficiency anaemia may also be caused by an inadequate supply of oxygen to tissues, resulting in increased concentrations of inflammatory mediators that activate leukocytes (26,27). There are inconsistencies in the results of these studies. Previous studies reported that MDA levels increased significantly in iron-deficiency anaemia (28-30).

In iron-deficiency anaemia, oxidative stress increases with the generation of free radicals, while therapeutic doses of iron supplements increase oxidative stress and antioxidant supplementation reduces oxidative stress. This increase and reduction in oxidative stress was studied among adults (31). Unlike in adults, oxidative stress in iron-deficiency anaemia is not aggravated by iron supplementation among children aged 10 months to 16 years (31). Tiwari et al. concluded that iron supplementation is effective in improving Hb levels but at the cost of increased oxidative stress among women with anaemia (29). An increase in MDA levels has been reported after 13 weeks of iron supplementation (32). An increase was also found in women after only 4 weeks of iron supplementation (33). The MDA level correlated with the serum ferritin level, suggesting that the iron status, having been modified by iron supplementation, increased the biomarker of lipid peroxidation (MDA).

In contrast, the findings of our study differ from other studies that reported iron supplementation for 6 weeks or 12 weeks caused a decrease in MDA levels (24,30), while another study reported no significant effects on lipid peroxidation and iron-deficiency anaemia after iron supplementation (33). These different findings might be explained by the different study designs, sample sizes, biomarkers, the consideration of certain covariates in some studies, and the health status of the participants. The use of various dosages of iron supplements and the duration of the supplementation are other possible reasons (24).

In our study, at week 13 of iron supplementation, the MDA biomarker was 44% higher than at the baseline. This finding is consistent with that of previous studies, which reported that an increase in the level of oxidative stress was induced by iron supplementation with ferrous fumarate or ferrous sulfate (26,32). In a 70-day study of iron-deficient non-anaemic women who were given ferrous sulfate (98.0 mg Fe/day) for 8 weeks, there was a marked increase in the plasma MDA level. At week 6 of the supplementation, the MDA indicator was more than 40% higher than at baseline (26,32).

A limitation of this study is that we did not include an intervention a group that only received nutritional education to compare with the other groups.

Conclusions

Interventions and strategies are needed to control anaemia and iron deficiency in Palestinian adolescent girls. The implementation of nutrition education programmes, including about iron-deficiency anaemia, in Palestinian secondary schools is recommended. The Palestinian Ministry of Health, in conjunction with the Ministry of Education, should carry out assessments of anaemia and iron deficiency.

The school-based nutrition education programme was associated with improvements in the Hb and iron status, and knowledge, attitudes and practices among female adolescents in Gaza. This findings highlights the importance of nutrition education. Given that iron supplementation helps adjust iron deficiency temporarily, a combination of supplementation and education is recommended to correct iron deficiency, and maintain Hb within the normal range. Our findings are supported by several studies showing nutritional education is an effective tool in improving haematocrit, Hb, serum ferritin levels and anaemia status among adolescents (34,35). Future studies are needed to determine whether nutrition education alone would be sufficient to increase Hb and ferritin among iron-deficient female adolescents.

Oxidative stress increases with iron deficiency and iron-deficiency anaemia. However, doses of ferrous fumarate for 3 months can increase the oxidative status by increasing MDA levels. The goal should be to correct anaemia without increasing the oxidative stress. Therefore, further studies are needed in this regard.

Funding: None.

Competing interests: None declared.

Effets de la supplémentation en fer et de l'éducation nutritionnelle sur l'hémoglobine, la ferritine et le stress oxydatif chez des adolescentes carencées en fer (Palestine) : essai contrôlé randomisé

Résumé

Contexte : Les carences en fer et l'anémie ferriprive sont associées au stress oxydatif, mais leur rôle reste grandement méconnu. Les informations portant sur les effets de la supplémentation en fer sur les biomarqueurs du stress oxydatif chez l'homme sont peu nombreuses.

Objectifs : La présente étude a évalué l'efficacité de la supplémentation en fer et de l'éducation nutritionnelle sur l'amélioration des taux d'hémoglobine et de ferritine, ainsi que sur la réduction du stress oxydatif chez les adolescentes carencées en fer à Gaza (Palestine).

Méthodes : Au total, 131 adolescentes carencées en fer ont été recrutées et réparties de façon aléatoire en trois groupes distincts. Le groupe de supplémentation en fer (A) a reçu 200 mg de fumarate de fer sur une base hebdomadaire au cours d'une intervention de trois mois. Le groupe de supplémentation en fer avec éducation nutritionnelle (B) a reçu des suppléments en fer et a assisté à des sessions d'éducation nutritionnelle. Le groupe témoin (C) n'a bénéficié d'aucune intervention. Les taux d'hémoglobine, de ferritine et de malonyl-dialdéhyde ont été mesurés au début, après trois mois (stade auquel l'intervention a été interrompue) et ensuite trois mois plus tard. (Numéro d'enregistrement de l'essai : ACTRN12618000960257).

Résultats : Les taux d'hémoglobine des groupes A et B augmentaient significativement après une supplémentation. À l'étape de suivi (trois mois après l'arrêt de l'intervention), les taux de fer et d'hémoglobine du groupe B continuaient d'augmenter et les taux de malonyl-dialdéhyde diminuaient. Dans le groupe A, les taux d'hémoglobine, de ferritine et de malonyl-dialdéhyde diminuaient trois mois après l'arrêt de l'intervention. Aucun changement n'a été observé dans le groupe C.

Conclusions : Un programme de nutrition devrait être adopté et intégré aux programmes d'intervention globaux de façon à cibler l'anémie ferriprive chez les adolescentes en Palestine.

أثر مكمّلات الحديد و التثقيف التغذوي على مستويات الهيموجلوبين والفيريتين والإجهاد التأكسدي لدى المراهقات اللائي تعانين من نقص الحديد في فلسطين: تجربة ضابطة عشوائية

مروان جلمبو، نوريما كريم، إيهاب نصر، رازينة شاريف الخلاصة

الخلفية: يرتبط نقص الحديد وفقر الدم الناجم عن نقص الحديد بالإجهاد التأكسدي، إلا أن دور كل منها يظل غير واضح بدرجة كبيرة. ولا تتوفر سوى معلومات شحيحة بشأن آثار مكمّلات الحديد على العلامات البيولوجية للإجهاد التأكسدي في البشر.

الأهداف: عمدت هذه الدراسة إلى تقييم فعالية مكمّلات الحديد والتثقيف التغذوي على تحسن مستويات الهيموجلوبين والفيريتين، وتضاؤل مستوى الإجهاد التأكسدي في صفوف المراهقات اللائي تعانين من نقص الحديد في قطاع غزة بفلسطين.

طرق البحث: ضمت هذه الدراسة ١٣١ مراهقة ممن تعانين من نقص الحديد، تم توزيعهن بشكل عشوائي على ٣ مجموعات مختلفة. فتلقّت مجموعة مكمّلات الحديد (المجموعة ألف) ٢٠٠ ملليجرام من فومارات الحديد أسبوعيا خلال فترة التدخل الممتدة لثلاثة أشهر، وتلقت مجموعة مكمّلات الحديد والتثقيف التغذوي (المجموعة باء) مكمّلات للحديد مع عقد جلسات للتثقيف التغذوي، ولم تتلق المجموعة الضابطة (المجموعة جيم) أي تدخلات. وقيست مستويات الهيموجلوبين والفيريتين وثنائي ألدهيد المالون مقابل خط الأساس، أي بعد ثلاثة أشهر (وهي وقت توقف التدخل)،

النتائج: ارتفعت مستويات الهيموجلوبين ارتفاعا دالا بعد استخدام المكمّلات في كلتا المجموعتين ألف وباء. وفي مرحلة المتابعة (بعد ثلاثة أشهر من وقف التدخل)، استمرت مستويات الحديد والهيموجلوبين في المجموعة باء في الارتفاع وانخفض مستوى ثنائي ألدهيد المالون. وفي المجموعة ألف، انخفضت مستويات الهيموجلوبين والفيريتين وثنائي ألدهيد المالون بعد ثلاثة أشهر من إيقاف التدخل. ولم تشاهد أي تغييرات في المجموعة جيم.

الاستنتاجات: ينبغي اعتماد برنامج للتغذية وإدماجه في برامج التدخل الشاملة لاستهداف فقر الدم الناجم عن نقص الحديد في صفوف المراهقات في فلسطين.

References

- 1. Tulchinsky TH. Micronutrient deficiency conditions: global health issues. Public Health Rev. 2010;32(1):243–55. https://doi. org/10.1007/BF03391600
- 2. Prevention of iron deficiency anaemia in adolescents. role of weekly iron and folic acid supplementation. New Delhi: World Health Organization Regional Office for South East Asia; 2011 (http://apps.searo.who.int/PDS_DOCS/B4770.pdf?ua=1, accessed 6 February 2018).
- 3. Jalambo MO, Hamad A, Abed Y. Anemia and risk factors among female secondary students in the Gaza Strip. J Public Health (Bangkok). 2013;21(3):271–8. https://doi.org/10.1007/s10389-012-0540-9
- 4. Abunada O, Jalambo MO. Assessment of iron status and its relation to diet and lifestyle among adolescents in the Gaza-Palestine. Journal of Scientific Research & Reports. 2015;4(3):265–73. https://doi.org/10.9734/JSRR/2015/11593
- 5. Shekhawat S, Gupta PP, Gupta M, Kakkar M, Mathur M, Ahaluwalia N, et al. A study of nutritional and health status of adolescent girls (10-19 Years) In Jaipur City. J Evolution Med Dent Sci. 2014;3(16):4299–309. https://doi.org/10.14260/jemds/2014/2435
- Kaur M, Bassi R, Sharma S. Impact of nutrition education in reducing iron deficiency anemia in adolescent girls. Ind J Fund Appl Life Sci. 2011;1:222–8 (http://www.cibtech.org/J-LIFE-SCIENCES/PUBLICATIONS/2011/Vol%201%20N0.%204/13-30-JLS-Manjeet. pdf, accessed 4 May 2018).
- 7. Alderman H. Improving nutrition through community growth promotion: longitudinal study of the nutrition and early child development program in Uganda. World Dev. 2007;35(8):1376–89. https://doi.org/10.1016/j.worlddev.2007.04.003
- 8. Spendlove JK, Heaney SE, Gifford JA, Prvan T, Denyer GS, O'Connor HT. Evaluation of general nutrition knowledge in elite Australian athletes. Br J Nutr. 2012 Jun;107(12):1871–80. https://doi.org/10.1017/S0007114511005125 PMID:22018024
- 9. Ryoo E. Adolescent nutrition: what do pediatricians do? Korean J Pediatr. 2011 Jul;54(7):287-91. https://doi.org/10.3345/ kjp.2011.54.7.287 PMID:22025921
- 10. Kulkarni MV, Durge P, Kasturwar N. Prevalence of anemia among adolescent girls in an urban slum. Natl J Community Med. 2012;3(1):108–11.
- 11. Senanayake HM, Premaratne SP, Palihawadana T, Wijeratne S. Simple educational intervention will improve the efficacy of routine antenatal iron supplementation. J Obstet Gynaecol Res. 2010 Jun;36(3):646–50. https://doi.org/10.1111/j.1447-0756.2010.01197.x PMID:20598050
- 12. Akhtar S, Ismail T, Atukorala S, Arlappa N. Micronutrient deficiencies in South Asia current status and strategies. Trends Food Sci Technol. 2013;31(1):55–62. https://doi.org/10.1016/j.tifs.2013.02.005
- 13. Bakr A, Abul Hassan S, Shoker M, Zaki M, Hassan R. Oxidant stress in primary nephrotic syndrome: does it modulate the response to corticosteroids? Pediatr Nephrol. 2009 Dec;24(12):2375–80. https://doi.org/10.1007/s00467-009-1246-2 PMID:19644712

- 14. Fan A, Jiang X, Mo Y, Tan H, Jiang M, Li J. Plasma levels of oxidative stress in children with steroid-sensitive nephrotic syndrome and their predictive value for relapse frequency. Pediatr Nephrol. 2016 Jan;31(1):83–8. https://doi.org/10.1007/s00467-015-3195-2 PMID:26341250
- 15. Iuchi Y. Anemia caused by oxidative stress. In: Silverberg DS, editor. Anaemia. Croatia: Intech.Org; 2012:49–52 (https://www. intechopen.com/books/anemia, accessed 7 February 2018).
- 16. Clemens MR, Waller HD. Lipid peroxidation in erythrocytes. Chem Phys Lipids. 1987 Nov-Dec;45(2-4):251–68. https://doi. org/10.1016/0009-3084(87)90068-5 PMID:3319229
- 17. Yusoff H, Daud WN, Ahmad Z. Nutrition education and knowledge, attitude and hemoglobin status of Malaysian adolescents. Southeast Asian J Trop Med Public Health. 2012 Jan;43(1):192–200. PMID:23082570
- 18. Opoku SK. Iron deficiency in relation to congnitive function among Ghanian children of school going age in the Bosomtwi Atwima Kwanoma District of Ghana. [PhD thesis]. Ghana: Kwame Nkrumah University of Science & Technology; 2012.
- 19. Shobha S, Sharada D. Efficacy of twice weekly iron supplementation in anemic adolescent girls. Indian Pediatr. 2003 Dec;40(12):1186–90. PMID:14722370
- 20. Trivedi P, Palta A. Prevalence of Anaemia and impact of iron supplementation on anaemic adolescent school girls. Health Popul Perspect Issues. 2007;30:45–55.
- 21. Zavaleta N, Respicio G, Garcia T. Efficacy and acceptability of two iron supplementation schedules in adolescent school girls in Lima, Peru. J Nutr. 2000 Feb;130(2S) Suppl:462S-4S. https://doi.org/10.1093/jn/130.2.462S PMID:10721929
- 22. Menendez C, Schellenberg D, Quinto L, Kahigwa E, Alvarez L, Aponte JJ, et al. The effects of short-term iron supplementation on iron status in infants in malaria-endemic areas. Am J Trop Med Hyg. 2004 Oct;71(4):434–40. PMID:15516639
- 23. Tee E-S, Kandiah M, Awin N, Chong S-M, Satgunasingam N, Kamarudin L, et al. School-administered weekly iron-folate supplements improve hemoglobin and ferritin concentrations in Malaysian adolescent girls. Am J Clin Nutr. 1999 Jun;69(6):1249–56. PMID:10357747
- 24. Khoshfetrat MR, Mohammadi F, Mortazavi S, Rashidi A, Neyestani T, Kalantari N, et al. The effect of iron-vitamin C co-supplementation on biomarkers of oxidative stress in iron-deficient female youth. Biol Trace Elem Res. 2013 Jun;153(1-3):171–7. https:// doi.org/10.1007/s12011-013-9695-7 PMID:23686563
- 25. Block G, Dietrich M, Norkus EP, Morrow JD, Hudes M, Caan B, et al. Factors associated with oxidative stress in human populations. Am J Epidemiol. 2002 Aug 1;156(3):274–85. https://doi.org/10.1093/aje/kwf029 PMID:12142263
- 26. Koskenkorva-Frank TS, Weiss G, Koppenol WH, Burckhardt S. The complex interplay of iron metabolism, reactive oxygen species, and reactive nitrogen species: insights into the potential of various iron therapies to induce oxidative and nitrosative stress. Free Radic Biol Med. 2013 Dec;65:1174–94. https://doi.org/10.1016/j.freeradbiomed.2013.09.001 PMID:24036104
- 27. Grune T, Sommerburg O, Siems WG. Oxidative stress in anemia. Clin Nephrol. 2000 Feb;53(1) Suppl:S18-22. PMID:10746801
- 28. Sundaram RC, Selvaraj N, Vijayan G, Bobby Z, Hamide A, Rattina Dasse N. Increased plasma malondialdehyde and fructosamine in iron deficiency anemia: effect of treatment. Biomed Pharmacother. 2007 Dec;61(10):682–5. https://doi.org/10.1016/j.bio-pha.2007.06.013 PMID:17698317
- 29. Tiwari AK, Mahdi AA, Chandyan S, Zahra F, Godbole MM, Jaiswar SP, et al. Oral iron supplementation leads to oxidative imbalance in anemic women: a prospective study. Clin Nutr. 2011 Apr;30(2):188–93. https://doi.org/10.1016/j.clnu.2010.08.001 PMID:20888091
- 30. Kurtoglu E, Ugur A, Baltaci AK, Undar L. Effect of iron supplementation on oxidative stress and antioxidant status in iron-deficiency anemia. Biol Trace Elem Res. 2003 Winter;96(1-3):117–23. https://doi.org/10.1385/BTER:96:1-3:117 PMID:14716090
- 31. King SM, Donangelo CM, Knutson MD, Walter PB, Ames BN, Viteri FE, et al. Daily supplementation with iron increases lipid peroxidation in young women with low iron stores. Exp Biol Med (Maywood). 2008 Jun;233(6):701–7. https://doi.org/10.3181/0708-RM-233 PMID:18408148
- 32. Knutson MD, Lim AK, Viteri FE. A practical and reliable method for measuring ethane and pentane in expired air from humans. Free Radic Biol Med. 1999 Sep;27(5-6):560-71. https://doi.org/10.1016/S0891-5849(99)00095-7 PMID:10490276
- 33. Meral A, Tuncel P, Sürmen-Gür E, Özbek R, Oztürk E, Günay U. Lipid peroxidation and antioxidant status in beta-thalassemia. Pediatr Hematol Oncol. 2000 Dec;17(8):687–93. https://doi.org/10.1080/08880010050211402 PMID:11127401
- 34. Khoshnevisan F, Kimiagar M, Kalantaree N, Valaee N, Shaheedee N. Effect of nutrition education and diet modification in iron depleted preschool children in nurseries in Tehran: a pilot study. Int J Vitam Nutr Res. 2004 Jul;74(4):264–8. https://doi. org/10.1024/0300-9831.74.4.264 PMID:15580808
- 35. Moore JB, Pawloski L, Rodriguez C, Lumbi L, Ailinger R. The effect of a nutrition education program on the nutritional knowledge, hemoglobin levels, and nutritional status of Nicaraguan adolescent girls. Public Health Nurs. 2009 Mar-Apr;26(2):144–52. https://doi.org/10.1111/j.1525-1446.2009.00765.x PMID:19261153