Int. J. Curr. Res. Chem. Pharm. Sci. (2024). 11(11): 26-34

## INTERNATIONAL JOURNAL OF CURRENT RESEARCH IN CHEMISTRY AND PHARMACEUTICAL SCIENCES

(p-ISSN: 2348-5213: e-ISSN: 2348-5221)

www.ijcrcps.com

(A Peer Reviewed, Referred, Indexed and Open Access Journal) DOI: 10.22192/ijcrcps Coden: IJCROO(USA) Volume 11, Issue 11- 2024

### **Review Article**



DOI: http://dx.doi.org/10.22192/ijcrcps.2024.11.11.005

## Iron Deficiency Anemia: Maternal and Fetal Outcomes in Pregnancy

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#### Abstract

Iron deficiency anemia (IDA) is the most prevalent nutritional deficiency during pregnancy, posing significant risks to maternal and fetal health. It arises due to the increased iron demands of pregnancy, coupled with inadequate dietary intake or absorption. Maternal IDA has been linked to adverse outcomes such as preterm delivery, postpartum hemorrhage, and maternal mortality. Additionally, it negatively impacts maternal quality of life by causing fatigue, reduced cognitive function, and increased susceptibility to infections. These challenges underscore the importance of addressing IDA as a public health priority. The consequences of IDA extend to the fetus, with profound implications for growth and development. Infants born to mothers with IDA are at increased risk of low birth weight, preterm birth, and impaired neurodevelopment. Furthermore, maternal IDA can deplete fetal iron stores, predisposing infants to anemia and developmental delays in early childhood. These outcomes highlight the critical role of maternal iron status in shaping neonatal and childhood health trajectories.

Keywords: Iron Deficiency Anemia, Pregnancy, Maternal Outcomes, Fetal Outcomes, Perinatal Health

#### Introduction

Iron deficiency anemia (IDA) is the most common nutritional disorder globally and remains a major public health concern, particularly during pregnancy. It is characterized by reduced hemoglobin levels due to insufficient iron, leading to impaired oxygen transport in the body. The World Health Organization (WHO) estimates that anemia affects 40% of pregnant women worldwide, with iron deficiency accounting for a significant proportion of cases. This condition

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disproportionately impacts low- and middleincome countries, where dietary iron intake is often insufficient and access to healthcare is limited.<sup>1-2</sup> Pregnancy places significant physiological demands on maternal iron stores, as iron is required for increased red blood cell production, placental development, and fetal growth. These demands, compounded by baseline deficiencies and factors such as blood loss. infections, and inflammation, often result in iron depletion during pregnancy. If untreated, IDA can lead to severe maternal and fetal complications, with far-reaching consequences for the health and well-being of both mother and child.<sup>3-4</sup> For the mother, IDA increases the risk of obstetric complications, including preterm delivery, postpartum hemorrhage, and even maternal mortality in severe cases. Beyond these lifethreatening conditions, maternal IDA significantly affects daily life, causing fatigue, weakness, and diminished cognitive function. These symptoms can impair a mother's ability to engage in prenatal care, maintain employment, or care for her family, further exacerbating health inequities and economic challenges.<sup>5-6</sup>

The adverse effects of maternal IDA are not limited to the mother; they extend to the fetus and neonate as well. Iron plays a critical role in fetal development, particularly for the brain and nervous system. Infants born to iron-deficient mothers are at heightened risk of low birth weight, preterm birth, and impaired cognitive development. Moreover, depleted fetal iron stores increase the likelihood of anemia in infancy, perpetuating a cycle of poor health outcomes that can extend into adulthood.<sup>7-8</sup> Despite the significant impact of IDA, it is a preventable and treatable condition. Timely detection through reliable diagnostic tools, including serum ferritin and transferrin saturation tests, is essential for mitigating the risks. Effective interventions, such as iron supplementation and dietary counseling, are widely available and can dramatically improve outcomes for both mothers and their babies. However, challenges persist, including low adherence to supplementation, limited healthcare access, and cultural dietary practices that may hinder the uptake of iron-rich foods.<sup>9-10</sup>

#### **Maternal Outcomes**

# 1. Increased Maternal Morbidity and Mortality

Iron deficiency anemia (IDA) during pregnancy is associated with a heightened risk of maternal morbidity and mortality. Anemia reduces oxygen transport capacity, compromising maternal cardiac output and increasing susceptibility to peripartum complications, such as postpartum hemorrhage and preeclampsia. Severe anemia (hemoglobin levels below 7 g/dL) has been linked to a higher likelihood of maternal death. particularly in low-resource settings where emergency obstetric care may be inadequate. Additionally, anemia exacerbates complications such as infections, which are more frequent and severe in anemic pregnant women due to compromised immune function.<sup>11-12</sup>

#### 2. Obstetric Complications

Pregnant women with IDA face an increased likelihood of adverse obstetric outcomes. including preterm labor, prolonged delivery, and uterine atony. These complications are partly attributable to reduced oxygen supply to uterine muscles, impairing their ability to contract effectively during labor. Moreover, IDA contributes to poor placental development and function, which can exacerbate pregnancy complications. Postpartum recovery is also affected, with delayed wound healing and prolonged hospital stays often observed in anemic mothers.<sup>13-14</sup>

#### **3. Impaired Quality of Life and Productivity**

Maternal IDA significantly impacts daily life by causing symptoms such as fatigue, dizziness, and reduced cognitive and physical performance. These symptoms diminish the mother's ability to perform household responsibilities, maintain employment, and adhere to prenatal care regimens. Psychosocial stress and mental health challenges, including depression and anxiety, are also more prevalent in anemic pregnant women. Furthermore, fatigue and weakness resulting from

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IDA may interfere with breastfeeding and postnatal care, further affecting maternal and infant health.<sup>15-16</sup>

#### 4. Nutritional and Immune Deficiencies

Iron plays a vital role in immune system function, and its deficiency during pregnancy leaves women more vulnerable to infections, including urinary tract infections and respiratory illnesses. Concurrently, IDA is often associated with deficiencies in other micronutrients, such as folic acid and vitamin B12, which are essential for hematopoiesis. These deficiencies can compound the negative effects of IDA, leading to more severe health outcomes.<sup>17</sup>

#### 5. Socioeconomic and Psychological Impact

In regions where IDA is prevalent, the economic burden of managing anemia-related complications adds to maternal stress. Increased healthcare costs, reduced household productivity, and caregiving challenges strain families financially and emotionally. The stigma associated with physical weakness or frequent illness may further contribute to psychosocial distress in affected women.<sup>18</sup>

#### 6. Long-term Maternal Health Risks

Untreated IDA during pregnancy has implications beyond the perinatal period. Women with a history of severe anemia are more likely to develop chronic conditions such as cardiovascular disease later in life. Additionally, repeated episodes of IDA in subsequent pregnancies can lead to cumulative health deterioration, emphasizing the importance of addressing anemia comprehensively during prenatal care.<sup>19</sup>

### **Fetal Outcomes**

# 1. Intrauterine Growth Restriction (IUGR) and Low Birth Weight

Iron deficiency anemia (IDA) in pregnant women is closely associated with intrauterine growth restriction (IUGR) and low birth weight. Iron is crucial for fetal growth and development, and maternal IDA can lead to reduced oxygen delivery to the fetus due to impaired hemoglobin function. This oxygen deficiency can hinder proper fetal growth, increasing the risk of low birth weight (<2.5 kg) and contributing to neonatal morbidity and mortality. IUGR also predisposes infants to a higher likelihood of developing chronic health issues later in life, such as metabolic syndrome and cardiovascular diseases.<sup>20</sup>

#### 2. Prematurity and Preterm Birth

IDA significantly increases the risk of preterm birth, defined as delivery before 37 weeks of gestation. The mechanisms linking IDA to preterm delivery include placental hypoxia, inflammation, and elevated levels of oxidative stress, all of which can trigger premature uterine contractions or placental abruption. Preterm infants face numerous complications, including respiratory distress syndrome, sepsis, and longterm neurodevelopmental challenges. These outcomes underline the importance of maintaining adequate maternal iron levels to support a fullterm pregnancy.<sup>21</sup>

#### **3. Impaired Neurodevelopment**

Iron is essential for the development of the fetal brain and nervous system, particularly during the third trimester, when brain growth is most rapid. Maternal IDA during pregnancy can disrupt iron transport to the fetus, affecting the development of critical neural pathways. This disruption has been associated with delayed cognitive, motor, and behavioral development in children. Studies have shown that children born to anemic mothers are at higher risk of developmental delays, attention-deficit hyperactivity disorder (ADHD), and learning difficulties, emphasizing the longterm consequences of prenatal iron deficiency.<sup>22</sup>

#### 4. Neonatal Anemia

Infants of mothers with IDA often experience low iron stores at birth, making them more susceptible to neonatal anemia. This condition can arise due

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to reduced iron transfer across the placenta during pregnancy. Neonatal anemia can further exacerbate developmental and growth challenges, particularly in resource-limited settings where access to iron supplementation or fortified nutrition may be limited.<sup>23</sup>

#### 5. Increased Risk of Perinatal Mortality

Severe maternal anemia has been linked to an elevated risk of perinatal mortality, which encompasses stillbirths and early neonatal deaths. Hypoxia resulting from inadequate oxygen transport due to maternal anemia can lead to fetal distress, increasing the likelihood of intrauterine death. Additionally, the risk of stillbirth is heightened in pregnancies complicated by severe IDA, highlighting the urgent need for early detection and intervention.<sup>24</sup>

# 6. Long-term Health Implications for Offspring

Emerging research suggests that IDA during epigenetic effects, pregnancy may have influencing gene expression in the developing fetus. These changes can predispose offspring to a range of chronic conditions, including obesity, diabetes. cardiovascular diseases, and in adulthood. The concept of "fetal programming" underscores how the intrauterine environment shapes long-term health trajectories, making maternal iron status a critical factor for future population health.<sup>24</sup>

### **Diagnostic Approaches**

#### 1. Hemoglobin and Hematocrit Levels

The initial step in diagnosing iron deficiency anemia (IDA) during pregnancy involves measuring hemoglobin (Hb) and hematocrit levels. Hemoglobin levels below 11 g/dL in the first and third trimesters or below 10.5 g/dL in the second trimester are indicative of anemia, according to World Health Organization (WHO) guidelines. While these tests are useful for screening, they lack specificity for distinguishing IDA from other types of anemia, such as those caused by folate or vitamin B12 deficiencies.<sup>25</sup>

#### 2. Serum Ferritin Measurement

Serum ferritin, a marker of iron stores, is the most reliable diagnostic tool for identifying IDA. Low serum ferritin levels (<15  $\mu$ g/L) are highly indicative of iron deficiency. However, ferritin is an acute-phase reactant and may be elevated in the presence of inflammation or infection, potentially masking iron deficiency. Adjusting ferritin thresholds during inflammatory conditions, such as by using C-reactive protein (CRP) or alpha-1-acid glycoprotein levels, can improve diagnostic accuracy.<sup>26</sup>

# **3. Transferrin Saturation and Total Iron Binding Capacity (TIBC)**

Transferrin saturation (TSAT), calculated as the ratio of serum iron to total iron-binding capacity (TIBC), provides additional insights into iron availability for erythropoiesis. A TSAT below 20% suggests iron deficiency. TIBC is typically elevated in IDA, reflecting increased production of transferrin in response to depleted iron stores. particularly These tests are useful in distinguishing IDA from anemia of chronic disease, where TIBC is often normal or decreased.27

#### 4. Peripheral Blood Smear Examination

A peripheral blood smear can reveal characteristic features of IDA, including microcytic (small-sized) and hypochromic (pale-colored) red blood cells. Anisocytosis (variation in red blood cell size) and poikilocytosis (abnormal shapes) may also be present. While not specific for IDA, these findings provide supportive evidence when combined with biochemical markers.<sup>28</sup>

#### 5. Reticulocyte Hemoglobin Content (CHr)

Reticulocyte hemoglobin content (CHr) is an emerging diagnostic parameter that reflects the hemoglobin content of newly formed red blood cells. CHr levels below 28 pg are indicative of

iron-restricted erythropoiesis. This test is particularly valuable for detecting early-stage IDA before significant changes in hemoglobin levels occur.<sup>29</sup>

#### 6. Soluble Transferrin Receptor (sTfR)

Soluble transferrin receptor (sTfR) levels are elevated in IDA and remain unaffected by inflammation, making it a useful diagnostic tool in patients with concurrent infections or chronic diseases. The sTfR/log ferritin ratio, known as the sTfR index, has been proposed as a reliable marker for distinguishing IDA from anemia of chronic disease.<sup>30</sup>

# 7. Advanced Techniques: Hepcidin and Zinc Protoporphyrin

Hepcidin, a regulatory hormone of iron metabolism, is decreased in IDA and increased in anemia of chronic disease. Measuring hepcidin levels can aid in differentiating these conditions, although its routine use is limited by high costs and lack of standardization. Similarly, zinc protoporphyrin (ZPP), a byproduct of impaired heme synthesis, is elevated in IDA and can serve as an adjunctive diagnostic tool.<sup>31</sup>

#### 8. Integration of Diagnostic Approaches

Given the multifactorial nature of anemia during pregnancy, a combination of diagnostic tests is often necessary for accurate identification and classification of IDA. Sequential testing, starting with hemoglobin and ferritin levels and followed by more specific markers such as TSAT or sTfR, can enhance diagnostic precision. Point-of-care testing and algorithms integrating clinical, biochemical, and hematological data are emerging as practical solutions for resource-limited settings.<sup>32</sup>

### **Management Strategies**

#### **1. Nutritional Interventions**

Nutritional interventions form the cornerstone of managing iron deficiency anemia (IDA) during

pregnancy. Increasing dietary intake of iron-rich foods, such as red meat, poultry, fish, legumes, and leafy green vegetables, is recommended. Complementary strategies include enhancing iron absorption by consuming foods rich in vitamin C, such as citrus fruits, while avoiding inhibitors like tea, coffee, and calcium-rich foods during meals. In low-resource settings, food fortification with iron is a practical approach to addressing widespread deficiencies. Programs that provide iron-fortified cereals, flours, or salt have shown promise in reducing IDA prevalence among pregnant women.<sup>33</sup>

#### 2. Iron Supplementation

Oral iron supplementation is the first-line treatment for IDA in pregnancy. Ferrous sulfate, ferrous gluconate, and ferrous fumarate are commonly prescribed, delivering 30-120 mg of elemental iron daily. To enhance absorption and gastrointestinal side minimize effects. supplementation is often taken with vitamin C or between meals. Newer formulations, such as liposomal iron and heme-based iron supplements, offer improved tolerability and efficacy. For women with severe anemia or poor response to oral therapy, intravenous iron formulations, such as ferric carboxymaltose or iron sucrose, provide a rapid and effective alternative.<sup>34</sup>

# **3.** Folic Acid and Multinutrient Supplementation

Since IDA often coexists with other micronutrient deficiencies, such as folate and vitamin B12, combining iron with a multivitamin supplement is advisable. Folic acid supplementation (400-600  $\mu$ g/day) is particularly crucial during pregnancy to prevent neural tube defects in the fetus while supporting hematopoiesis. Prenatal supplements that include additional nutrients, such as zinc and vitamin A, may offer comprehensive benefits for both maternal and fetal health.<sup>35</sup>

#### 4. Blood Transfusion

In cases of severe anemia (hemoglobin <7 g/dL) with imminent risks to maternal or fetal health,

blood transfusions are a life-saving intervention. Transfusion rapidly restores oxygen-carrying capacity and stabilizes critical patients. However, it is used sparingly due to risks such as transfusion reactions, infections, and alloimmunization. Ensuring the availability of safe blood and appropriate crossmatching practices is essential for effective transfusion management.<sup>36</sup>

#### 5. Management of Underlying Causes

Addressing the root causes of IDA is integral to its management. Infections such as malaria, hookworm, or schistosomiasis, which contribute to iron depletion, must be treated concurrently. Antiparasitic medications, along with preventive measures like deworming and use of insecticidetreated nets, play a critical role in reducing recurrent anemia. Similarly, managing chronic inflammatory conditions that impair iron utilization is essential for sustaining treatment efficacy.<sup>32</sup>

#### 6. Public Health Interventions

At the population level, public health initiatives aimed at improving maternal nutrition and anemia awareness are vital. Community-based education programs emphasize the importance of prenatal care, dietary modifications, and adherence to iron supplementation. Regular anemia screening during antenatal visits allows for early detection and intervention. Policymakers should prioritize equitable access to iron supplements and healthcare services, especially in low-resource settings where anemia rates are highest.<sup>33</sup>

#### 7. Emerging Therapies

Research into novel therapies for IDA in pregnancy is expanding. Hepcidin antagonists, erythropoiesis-stimulating agents, and iron nanoparticles represent emerging options that may overcome limitations of current treatments. For instance, hepcidin inhibitors could enhance iron absorption and mobilization in patients with concurrent inflammation or chronic disease. These innovations, coupled with advancements in diagnostic precision, could revolutionize the management of IDA in the future.<sup>34-35</sup>

#### 8. Monitoring and Follow-up

Effective management of IDA requires regular monitoring to assess treatment response and ensure maternal and fetal safety. Hemoglobin levels are typically re-evaluated 4-6 weeks after initiating treatment, with adjustments made based on clinical progress. Long-term follow-up ensures sustained improvement in iron status and prevents relapse, particularly in women with repeated pregnancies or chronic conditions.<sup>36</sup>

### Conclusion

Iron deficiency anemia (IDA) in pregnancy remains a significant public health challenge due to its profound impact on maternal and fetal outcomes. The condition contributes to adverse maternal effects, including fatigue, reduced work capacity, and increased risks of preterm labor and postpartum hemorrhage, while also being strongly associated with fetal complications such as intrauterine growth restriction, low birth weight, preterm delivery, and impaired neurodevelopment. Addressing IDA is essential not only for optimizing pregnancy outcomes but also for promoting the lifelong health and development of the offspring. Effective management strategies for IDA involve a multifaceted approach, combining nutritional interventions, iron supplementation, and treatment of underlying causes with advanced therapies and robust public health initiatives. Early diagnosis through integrated screening protocols and targeted therapies tailored to individual needs is crucial for timely intervention. Public health programs that focus on education, equitable access to resources, and food fortification play a pivotal role in reducing the global burden of IDA, particularly in resource-limited settings.

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How to cite this article:

Emmanuel Ifeanyi Obeagu. (2024). Iron Deficiency Anemia: Maternal and Fetal Outcomes in Pregnancy. Int. J. Curr. Res. Chem. Pharm. Sci. 11(11): 26-34. DOI: http://dx.doi.org/10.22192/ijcrcps.2024.11.11.005