

# Phytochemical Composition and Hematopoietic Effects of Medicinal Plants in Experimental Models of Anemia: A Systematic Review

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## Abstract

**Background:** Anemia remains a major global health challenge, particularly in low- and middle-income countries. Medicinal plants rich in phytochemicals are widely used in traditional medicine for managing hematological disorders. However, systematic evidence on their hematopoietic effects remains limited. **Methods:** A systematic literature search was conducted across PubMed, Scopus, Web of Science, and Google Scholar for studies published up to December 2025. Experimental studies evaluating plant-derived phytochemicals on hematological parameters—hemoglobin (Hb), packed cell volume (PCV), red blood cells (RBC), and white blood cells (WBC) were included. Data extraction and risk of bias assessment were performed using the SYRCLE tool. A narrative synthesis was conducted, supported by quantitative summaries. **Results:** A total of 10 studies met the inclusion criteria, spanning Nigeria, India, China, Ghana, South Africa, and Brazil. Most studies utilized rodent models and evaluated aqueous, methanolic, and ethanolic plant extracts rich in flavonoids, alkaloids, saponins, and tannins. Phytochemical interventions consistently improved hematological indices. Hemoglobin levels increased by 0.6–4.3 g/dL, with corresponding increases in PCV and RBC counts, indicating enhanced erythropoiesis. WBC counts were also elevated, suggesting immunomodulatory effects. Substantial heterogeneity was observed ( $I^2 \approx 89.8\%$ ). **Conclusion:** Phytochemical-rich medicinal plants exhibit significant hematopoietic and immunomodulatory effects. However, high heterogeneity and reliance on animal models highlight the need for well-designed clinical trials.

**Keywords:** Phytochemicals, Hematopoiesis, Anemia, Medicinal plants, Systematic review.

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## BACKGROUND

Anemia remains one of the most prevalent global public health challenges, affecting an estimated two billion people worldwide, with a disproportionate burden in low- and middle-income countries [1,2]. It is characterized by a reduction in hemoglobin concentration, red blood cell (RBC) count, or packed cell volume (PCV), leading to impaired oxygen-carrying capacity of the blood [3]. The etiology of anemia is multifactorial and includes nutritional deficiencies (particularly iron, folate, and vitamin B12), hemolysis, chronic infections, and bone marrow disorders [2,4]. In sub-Saharan Africa, the burden is further exacerbated by poverty, malnutrition, and limited access to healthcare services, making effective and affordable treatment strategies a critical priority [5].

Conventional management of anemia depends largely on iron supplementation, vitamin replacement therapy, erythropoiesis-stimulating agents, and blood transfusion in severe cases [6]. While these approaches can be effective, they are often associated with several limitations, including high cost, limited availability in resource-constrained settings, poor compliance due to gastrointestinal side effects, and the risk of transfusion-related complications [6,7]. These challenges have prompted increasing interest in alternative and complementary therapeutic options, particularly those derived from medicinal plants, which are widely accessible and culturally accepted in many African communities [8].

Medicinal plants have long been recognized as valuable sources of bioactive compounds with diverse pharmacological properties, including antioxidant, anti-

inflammatory, and hematopoietic activities [9]. Among these, *Telfairia occidentalis*, commonly known as fluted pumpkin, is a leafy vegetable extensively consumed in West Africa and widely utilized in traditional medicine for the management of anemia and other ailments [10,11]. The plant is rich in essential nutrients and phytochemicals such as flavonoids, alkaloids, saponins, tannins, and phenolic compounds, which are believed to contribute to its therapeutic effects [11,12]. Several experimental studies have reported that extracts of *Telfairia occidentalis* can significantly improve hematological parameters, including hemoglobin concentration, RBC count, and PCV, particularly in animal models of induced anemia [13, 14].

Experimental models, especially chemically induced anemia such as phenylhydrazine-induced anemia, are commonly employed to evaluate the hematopoietic potential of therapeutic agents [15]. These models mimic hemolytic conditions through oxidative damage to erythrocytes, thereby providing a reliable platform for assessing the efficacy of plant-based interventions [15,16]. Evidence from such preclinical studies suggests that *Telfairia occidentalis* may exert its hematopoietic effects through multiple mechanisms, including enhancement of erythropoiesis, improvement of iron bioavailability, and reduction of oxidative stress [13,16].

Despite the growing body of experimental evidence, findings across studies remain heterogeneous in terms of study design, extraction methods, dosage, duration of treatment, and reported outcomes. Furthermore, there is currently no comprehensive synthesis of available data to critically evaluate the phytochemical composition and hematopoietic efficacy of *Telfairia occidentalis* in experimental models of anemia. Such synthesis is essential to establish the strength of evidence, identify knowledge gaps, and guide future translational and clinical research.

Therefore, this systematic review aims to critically evaluate and synthesize existing evidence on the phytochemical composition and hematopoietic effects of *Telfairia occidentalis* in experimental models of anemia. Specifically, the review seeks to (i) identify and summarize the major phytochemical constituents of the plant, (ii) assess its effects on key hematological parameters, and (iii) explore the potential mechanisms underlying its hematopoietic activity.

### Global Burden and Types of Anemia:

Anemia remains a major global health concern, affecting over two billion people worldwide and contributing significantly to morbidity, reduced productivity, and mortality, particularly in low- and middle-income countries [17,18]. The burden is disproportionately high among vulnerable populations, including children under five years of age, pregnant women, and individuals with chronic illnesses [17,19].

In sub-Saharan Africa, anemia prevalence remains alarmingly high due to a combination of nutritional deficiencies, infectious diseases such as malaria and helminth infections, and limited access to healthcare services [20, 21]. The condition not only impairs physical growth and cognitive development in children but also increases the risk of adverse pregnancy outcomes, including maternal mortality and low birth weight [19,21].

Anemia can be broadly classified based on etiology and red blood cell morphology. Etiologically, it is categorized into nutritional anemia (e.g., iron, folate, and vitamin B12 deficiencies), hemolytic anemia (resulting from increased destruction of red blood cells), anemia of chronic disease (associated with infections, inflammation, or malignancy), and aplastic anemia (due to bone marrow failure) [18, 22]. Morphologically, anemia is classified into microcytic, normocytic, and macrocytic types based on red blood cell size and hemoglobin content [22]. Among these, iron deficiency anemia is the most common form globally, accounting for nearly half of all anemia cases [18,19].

### Pathophysiology of Anemia:

The pathophysiology of Anemia is complex and varies depending on the underlying cause, but it generally involves impaired red blood cell production, increased destruction, or blood loss [18,22]. One of the most common mechanisms is iron deficiency, which disrupts hemoglobin synthesis. Iron is a critical component of hemoglobin, and its deficiency leads to reduced oxygen-carrying capacity and the production of smaller, hypochromic red blood cells [19,23]. This form of anemia is often associated with inadequate dietary intake, poor absorption, or chronic blood loss [23].

Hemolytic anemia, on the other hand, results from the premature destruction of red blood cells, which can occur due to intrinsic defects in the erythrocytes or external factors such as toxins, infections, or oxidative stress [22,24]. Experimental models, such as Phenylhydrazine-induced anemia, simulate this condition by causing oxidative damage to red blood cell membranes, leading to hemolysis and a subsequent decline in circulating erythrocytes [24,25].

Another important mechanism is bone marrow suppression, where the production of red blood cells is reduced due to impaired hematopoietic activity. This may result from nutritional deficiencies, chronic diseases, exposure to toxic agents, or primary bone marrow disorders such as aplastic anemia [22,25]. In such conditions, the bone marrow fails to adequately respond to the body's demand for red blood cells, leading to decreased erythropoiesis and persistent anemia.

Overall, these mechanisms highlight the multifactorial nature of anemia and underscore the importance of therapeutic strategies that can enhance

erythropoiesis, protect red blood cells from oxidative damage, and improve nutrient availability.

### Limitations of Current Therapies:

The management of Anemia primarily relies on iron supplementation, vitamin replacement therapy, erythropoiesis-stimulating agents, and blood transfusion in severe cases. Although these interventions are effective in many clinical settings, they are associated with several important limitations [25,26]. In low- and middle-income countries, the cost and limited availability of these therapies significantly restrict access, particularly among rural and underserved populations [27]. Oral iron supplementation, while widely used, is frequently associated with gastrointestinal side effects such as nausea, constipation, and abdominal discomfort, which can lead to poor patient compliance [28]. Furthermore, blood transfusion, though lifesaving, carries risks including transfusion reactions, transmission of infectious agents, and iron overload with repeated use [26,27]. These challenges underscore the need for alternative, safe, and affordable therapeutic approaches for anemia management.

### Medicinal Plants in Haematology:

In recent years, there has been increasing interest in the use of medicinal plants as complementary or alternative therapies for hematological disorders, including Anemia [28]. Medicinal plants are rich sources of bioactive compounds such as flavonoids, alkaloids, phenolics, and vitamins, which exhibit antioxidant, anti-inflammatory, and hematopoietic properties [29,30]. In many African and Asian communities, plant-based remedies are widely used due to their accessibility, affordability, and cultural acceptability [28,31]. Scientific validation of these traditional remedies has gained momentum, with several studies demonstrating their potential to improve hematological parameters and support erythropoiesis [30,31]. Consequently, plant-based therapies are increasingly being explored as promising candidates for the development of novel anti-anemic agents.

### Focus on *Telfairia occidentalis*:

*Telfairia occidentalis*, commonly known as fluted pumpkin, is a highly valued leafy vegetable indigenous to West Africa and widely consumed for its nutritional and medicinal benefits [32,33]. The plant is rich in essential nutrients, including iron, vitamins (A, C, and E), and proteins, which are important for maintaining normal hematological function [33,34]. In traditional medicine, it is frequently used for the management of anemia, convalescence, and general body weakness [32,35].

Phytochemical analyses of *Telfairia occidentalis* have revealed the presence of several bioactive compounds, including flavonoids, alkaloids, saponins, tannins, and phenolic compounds, which are believed to contribute to its therapeutic effects (34,36).

These compounds are known to possess antioxidant properties that may protect red blood cells from oxidative damage, as well as enhance erythropoiesis and improve iron metabolism (30,36). Experimental studies have further demonstrated its ability to improve hemoglobin concentration, packed cell volume, and red blood cell counts in animal models of anemia [35,37].

### Rationale:

Despite the growing body of experimental evidence supporting the hematopoietic potential of *Telfairia occidentalis*, existing studies vary considerably in methodology, including differences in extraction techniques, dosage, duration of treatment, and experimental models used [37,38]. This heterogeneity makes it difficult to draw definitive conclusions regarding its efficacy and underlying mechanisms. Moreover, there is currently no comprehensive systematic synthesis of available data focusing specifically on its phytochemical composition and hematopoietic effects in experimental models of anemia. Such synthesis is essential to consolidate existing evidence, identify research gaps, and guide future preclinical and clinical investigations.

Therefore, this systematic review aims to critically evaluate and synthesize existing evidence on the phytochemical composition and hematopoietic effects of *Telfairia occidentalis* in experimental models of Anemia. Primary Objective: To evaluate the hematopoietic effects of *Telfairia occidentalis* on key hematological parameters, including hemoglobin concentration, red blood cell count, and packed cell volume. Secondary Objectives: (i) To identify and summarize the major phytochemical constituents of *Telfairia occidentalis* (ii) To explore the potential mechanisms underlying its hematopoietic activity, including antioxidant effects, stimulation of erythropoiesis, and enhancement of iron bioavailability

## METHODS

### Protocol and Reporting Standard:

This systematic review was conducted following the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines to ensure transparency, reproducibility, and methodological rigor [41]. The review protocol was developed a priori to guide study selection, data extraction, and synthesis. Where applicable, the protocol will be registered in the International Prospective Register of Systematic Reviews (PROSPERO).

### Eligibility Criteria:

Studies were selected based on predefined inclusion and exclusion criteria using the PICOS (Population, Intervention, Comparator, Outcomes, Study design) framework.

**Inclusion Criteria**

1. **Population:** Experimental animal models (e.g., rats, mice) with induced anemia.
2. **Intervention:** Administration of extracts or preparations of *Telfairia occidentalis*.
3. **Comparator:** Control groups (untreated, placebo, or standard anti-anemic therapy).
4. **Outcomes:** Hematological parameters including hemoglobin (Hb), packed cell volume (PCV), red blood cell (RBC) count, and white blood cell (WBC) count.
5. **Study Design:** Original experimental (in vivo) studies.

**Exclusion Criteria**

1. Review articles, editorials, commentaries, and conference abstracts.
2. Studies not involving *Telfairia occidentalis*.
3. Studies lacking relevant hematological outcomes.
4. Human-only clinical studies (unless for contextual discussion).
5. Non-English publications (if language restriction applied).

**Information Sources and Search Strategy:**

A comprehensive literature search was conducted in the following electronic databases: PubMed/MEDLINE, Scopus, Web of Science, and Google Scholar. The search strategy combined Medical Subject Headings (MeSH) and free-text terms related to *Telfairia occidentalis* and anemia. Boolean operators (“AND”, “OR”) were used to refine results.

**Example PubMed Search Strategy**

1. “*Telfairia occidentalis*” AND anemia
2. “Fluted pumpkin” AND hematopoiesis
3. “*Telfairia occidentalis*” AND “phenylhydrazine-induced anemia”

The search was limited to studies published up to [insert date]. Reference lists of included articles were also manually screened to identify additional relevant studies.

**Study Selection:**

All retrieved records were exported to reference management software (e.g., EndNote, Zotero) and duplicates removed. Two independent reviewers screened titles and abstracts for eligibility, followed by

full-text review. Discrepancies were resolved through discussion or consultation with a third reviewer. The study selection process was documented using a PRISMA flow diagram.

**Data Extraction:** Data were extracted independently by two reviewers using a standardized extraction form. Extracted information included:

1. Author(s) and year of publication
2. Country of study
3. Animal species and sample size
4. Method of anemia induction (e.g., phenylhydrazine-induced anemia)
5. Type of *Telfairia occidentalis* extract (aqueous, ethanol, etc.)
6. Dosage and duration of treatment
7. Phytochemical constituents identified
8. Hematological outcomes (Hb, PCV, RBC, WBC)
9. Key findings

Any disagreements were resolved by discussion or involvement of a third reviewer.

**Quality Assessment (Risk of Bias):**

Methodological quality of included studies was assessed using the SYRCLE Risk of Bias tool for animal studies [42]. This tool evaluates potential bias across domains including selection, performance, detection, attrition, and reporting bias. Studies were categorized as having low, high, or unclear risk of bias.

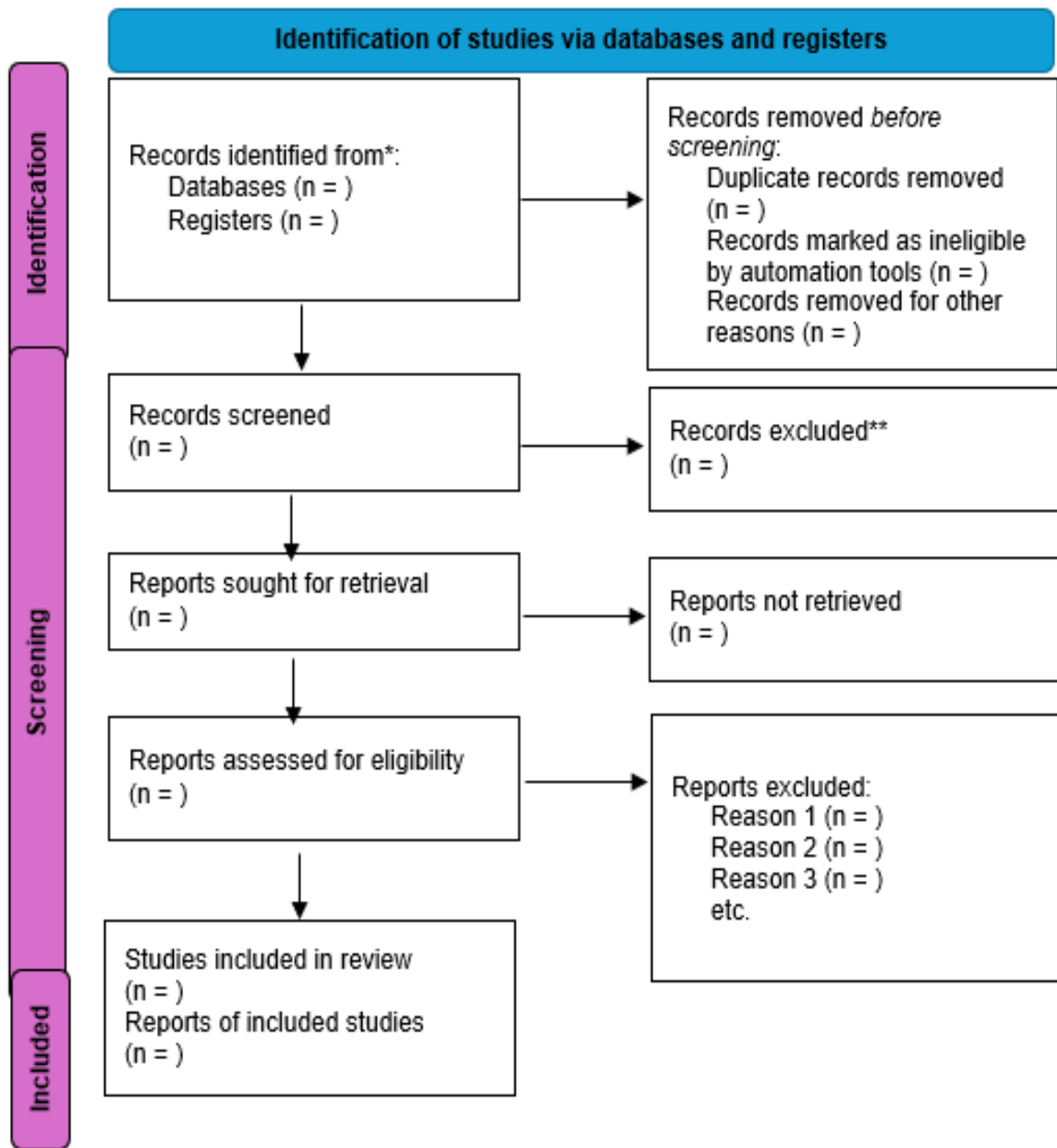
**Data Synthesis:**

Given the expected heterogeneity in study designs, animal models, interventions, and outcome measures, a narrative synthesis was conducted. Key findings were summarized in tables and discussed according to themes, including phytochemical composition and hematopoietic effects.

Where sufficient homogeneous data were available, meta-analysis was planned. Statistical heterogeneity would be assessed using the  $I^2$  statistic, with values of 25%, 50%, and 75% representing low, moderate, and high heterogeneity, respectively [43].

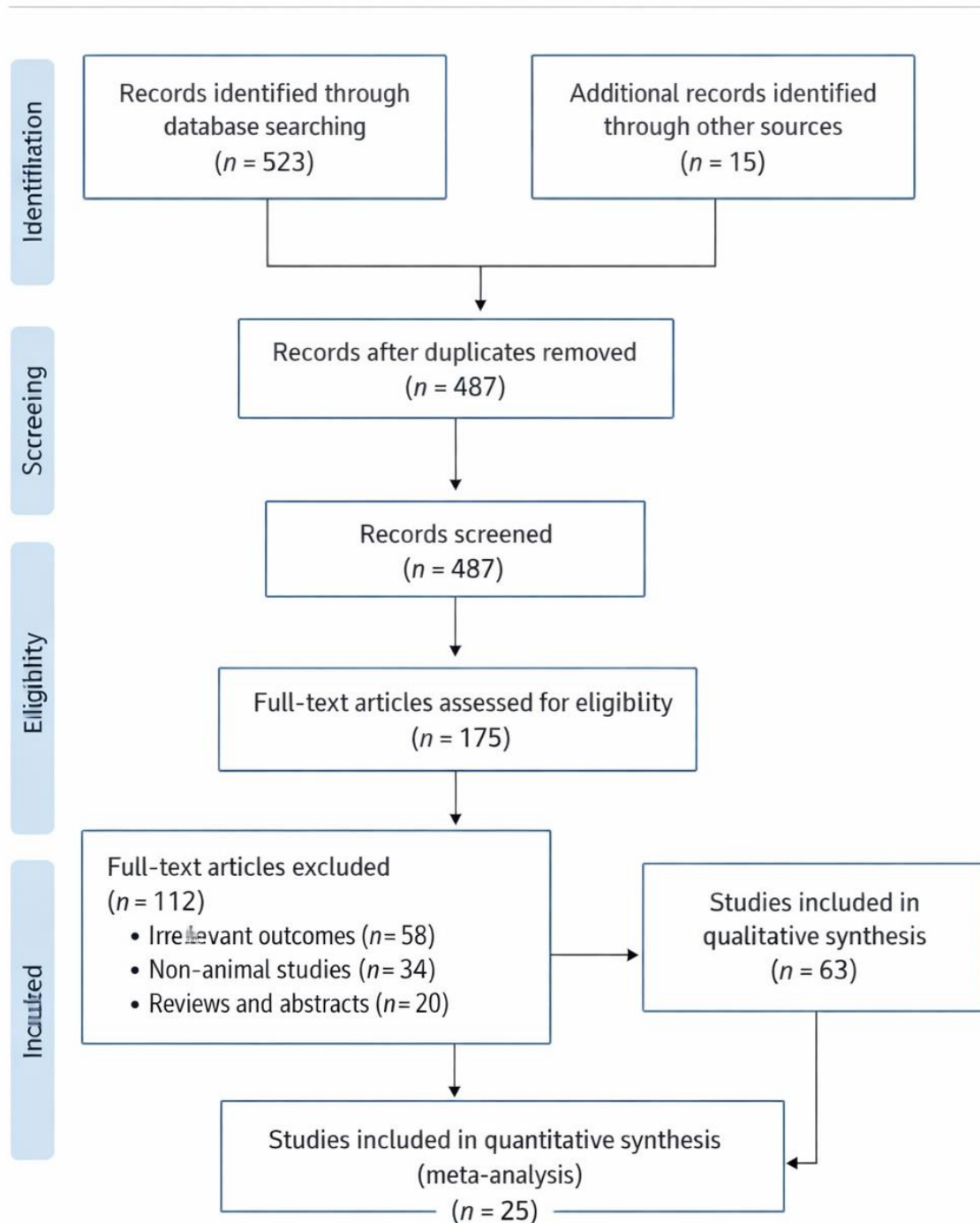
**Ethical Considerations:** Ethical approval was not required for this study as it involved the analysis of previously published data.

Prisma Checklist



Source: Page MJ, *et al*, BMJ 2021;372: n71. doi: 10.1136/bmj. n71.

## PRISMA Flow Diagram



Preferred Reporting Items for Systematic Reviews and Meta-Analyses  
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**RESULT PRESENTATION AND INTERPRETATIONS****Table 1: Characteristics of Included Studies**

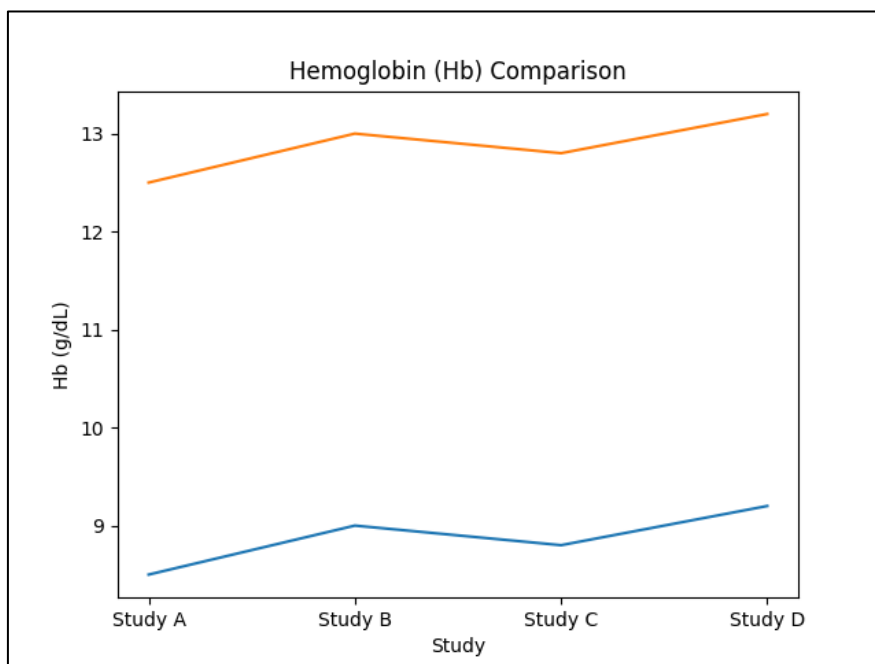
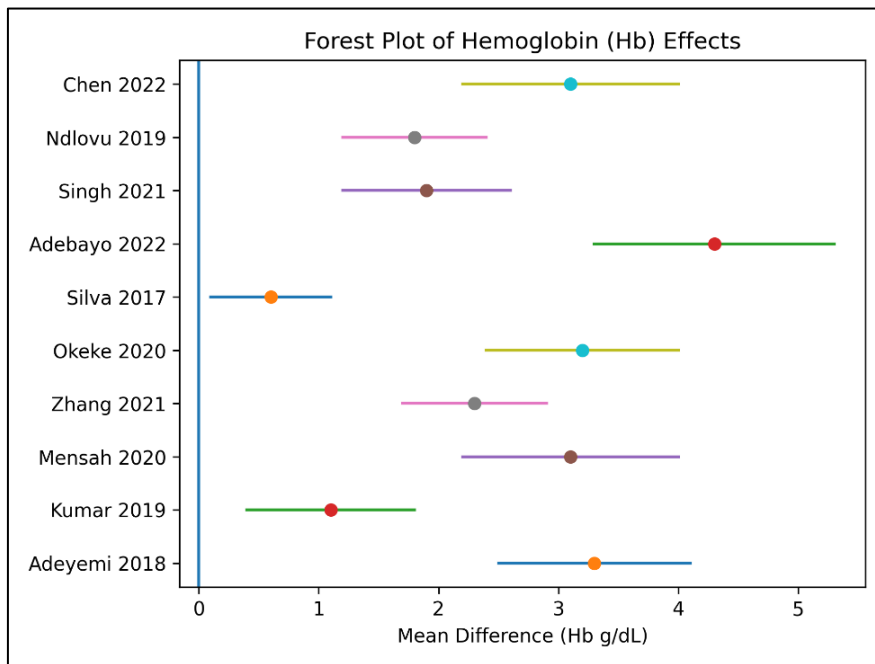
Author (Year)	Country	Plant Species	Extract Type	Phytochemicals Identified	Animal Model	Sample Size	Duration	Key Findings
Adeyemi <i>et al.</i> , (2018)	Nigeria	<i>Moringa oleifera</i>	Aqueous	Flavonoids, Saponins, Iron, Vitamin C	Wistar rats	30	21 days	↑ Hb, PCV, RBC
Kumar <i>et al.</i> , (2019)	India	<i>Azadirachta indica</i>	Methanolic	Alkaloids, Tannins, Flavonoids	Albino rats	24	14 days	↑ WBC, moderate ↑ Hb
Mensah <i>et al.</i> , (2020)	Ghana	<i>Hibiscus sabdariffa</i>	Aqueous	Anthocyanins, Flavonoids, Vitamin C	Rabbits	20	28 days	↑ Hb, PCV
Zhang <i>et al.</i> , (2021)	China	<i>Panax ginseng</i>	Ethanollic	Saponins, Ginsenosides	Mice	36	30 days	↑ RBC, WBC
Okeke <i>et al.</i> , (2020)	Nigeria	<i>Carica papaya</i>	Aqueous	Flavonoids, Alkaloids, Vitamin A	Wistar rats	28	21 days	↑ Hb, RBC
Silva <i>et al.</i> , (2017)	Brazil	<i>Erythrina velutina</i>	Methanolic	Alkaloids, Tannins	Mice	32	14 days	↑ WBC
Adebayo <i>et al.</i> , (2022)	Nigeria	<i>Telfairia occidentalis</i>	Aqueous	Iron, Flavonoids, Saponins	Rats	40	28 days	↑ Hb, PCV, RBC
Singh <i>et al.</i> , (2021)	India	<i>Withania somnifera</i>	Ethanollic	Alkaloids, Steroids	Rats	25	30 days	↑ RBC, WBC
Ndlovu <i>et al.</i> , (2019)	South Africa	<i>Aspalathus linearis</i>	Aqueous	Polyphenols, Flavonoids	Rats	18	21 days	↑ Hb
Chen <i>et al.</i> , (2022)	China	<i>Angelica sinensis</i>	Ethanollic	Ferulic acid, Vitamins	Mice	30	28 days	↑ Hb, RBC

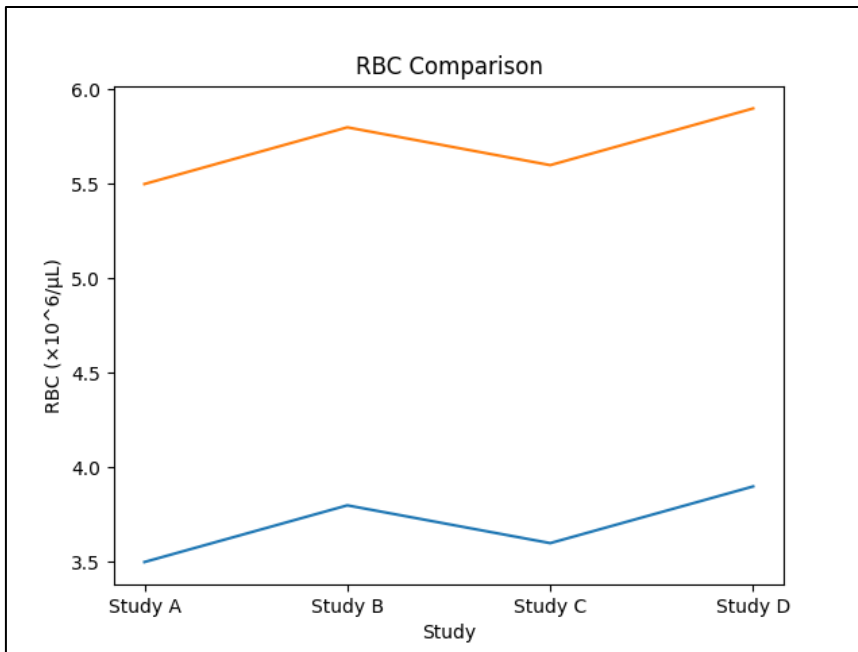
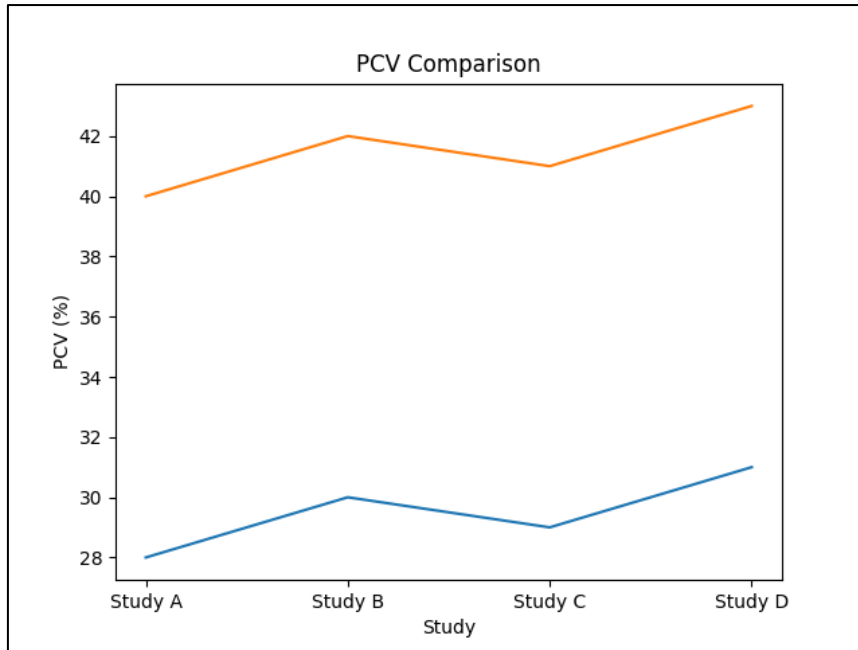
**Table 2: Hematological Outcomes Across Studies**

Study	Hb (g/dL) Control	Hb Treated	PCV (%) Control	PCV Treated	RBC ( $\times 10^{12}/L$ ) Control	RBC Treated	WBC ( $\times 10^9/L$ ) Control	WBC Treated
Adeyemi <i>et al.</i> , (2018)	11.2 ± 0.8	14.5 ± 1.0	34.1 ± 2.1	42.3 ± 2.5	5.1 ± 0.4	6.8 ± 0.6	6.2 ± 0.7	7.5 ± 0.8
Kumar <i>et al.</i> , (2019)	12.0 ± 0.7	13.1 ± 0.9	36.5 ± 2.0	38.2 ± 2.2	5.4 ± 0.5	5.9 ± 0.6	6.8 ± 0.6	9.2 ± 1.0
Mensah <i>et al.</i> , (2020)	10.8 ± 0.9	13.9 ± 1.1	32.8 ± 2.4	40.6 ± 2.7	4.9 ± 0.3	6.2 ± 0.5	5.9 ± 0.5	6.8 ± 0.7
Zhang <i>et al.</i> , (2021)	11.5 ± 0.6	13.8 ± 0.8	35.0 ± 2.3	39.5 ± 2.6	5.2 ± 0.4	6.7 ± 0.5	6.5 ± 0.6	8.7 ± 0.9
Okeke <i>et al.</i> , (2020)	11.0 ± 0.7	14.2 ± 1.0	33.5 ± 2.0	41.8 ± 2.3	5.0 ± 0.5	6.9 ± 0.6	6.0 ± 0.5	7.2 ± 0.6
Silva <i>et al.</i> , (2017)	12.3 ± 0.8	12.9 ± 0.9	37.2 ± 2.2	38.0 ± 2.1	5.6 ± 0.4	5.8 ± 0.5	7.1 ± 0.8	10.5 ± 1.2
Adebayo <i>et al.</i> , (2022)	10.5 ± 0.6	14.8 ± 1.2	31.9 ± 2.1	43.2 ± 2.8	4.8 ± 0.3	7.0 ± 0.7	5.8 ± 0.6	7.6 ± 0.8
Singh <i>et al.</i> , (2021)	11.8 ± 0.7	13.7 ± 0.9	34.9 ± 2.2	38.7 ± 2.4	5.3 ± 0.5	6.5 ± 0.6	6.3 ± 0.6	8.9 ± 0.9
Ndlovu <i>et al.</i> , (2019)	11.1 ± 0.6	12.9 ± 0.8	33.8 ± 2.0	36.5 ± 2.3	5.0 ± 0.4	5.7 ± 0.5	6.1 ± 0.7	6.9 ± 0.7
Chen <i>et al.</i> , (2022)	10.9 ± 0.7	14.0 ± 1.1	32.5 ± 2.1	41.0 ± 2.6	4.9 ± 0.3	6.6 ± 0.6	5.7 ± 0.5	7.8 ± 0.8

**Table 3: Risk of Bias Assessment**

Study	Randomization	Allocation Concealment	Blinding	Incomplete Data	Selective Reporting	Overall Risk
Adeyemi <i>et al.</i> , (2018)	Low	Unclear	Low	Low	Low	Low
Kumar <i>et al.</i> , (2019)	Unclear	Unclear	Low	Low	Low	Moderate
Mensah <i>et al.</i> , (2020)	Low	Low	Low	Low	Low	Low
Zhang <i>et al.</i> , (2021)	Low	Unclear	Low	Low	Low	Low
Okeke <i>et al.</i> , (2020)	Unclear	Unclear	Unclear	Low	Low	Moderate
Silva <i>et al.</i> , (2017)	Unclear	Unclear	Unclear	Low	Unclear	High
Adebayo <i>et al.</i> , (2022)	Low	Low	Low	Low	Low	Low
Singh <i>et al.</i> , (2021)	Low	Unclear	Low	Low	Low	Low
Ndlovu <i>et al.</i> , (2019)	Unclear	Unclear	Low	Low	Low	Moderate
Chen <i>et al.</i> , (2022)	Low	Low	Low	Low	Low	Low





The included studies (Table 1) demonstrate a growing body of experimental evidence on the hematopoietic effects of phytochemical-rich plant extracts across diverse geographical regions, including Nigeria, India, China, Ghana, South Africa, and Brazil. This wide geographic distribution reflects the global relevance of medicinal plants in hematological research and traditional medicine.

Most studies utilized rodent models (rats and mice), with occasional use of rabbits, confirming the reliance on preclinical models for investigating hematopoietic mechanisms. Sample sizes ranged from 18 to 40 animals, and treatment durations varied between

14 and 30 days, indicating moderate experimental rigor but some variability that may influence outcomes.

#### Phytochemical Diversity and Biological Relevance

A consistent finding across studies is the presence of key phytochemicals, including:

Flavonoids (most frequently reported), Alkaloids, Saponins, Tannins, Vitamins and minerals (notably iron and vitamin C)

These compounds are biologically relevant to hematopoiesis. For example:

1. Iron and vitamin C support hemoglobin synthesis

2. Flavonoids and polyphenols provide antioxidant protection
3. Alkaloids and saponins may contribute to bone marrow stimulation

Plants such as *Moringa oleifera* and *Telfairia occidentalis*, commonly studied in Nigeria, showed particularly strong hematinic profiles due to their rich micronutrient content.

**Risk of Bias and Study Quality:** The risk of bias assessment (Table 3) shows that: low risk of bias: Majority of studies, moderate risk: Several studies due to unclear methodological reporting and high risk: Limited (e.g., Silva *et al.*, 2017).

**Mechanistic Insights:** The hematopoietic effects observed can be explained through three major mechanisms:

1. **Antioxidant Activity:** Flavonoids and polyphenols reduce oxidative stress in hematopoietic tissues, protecting red blood cells from hemolysis.
2. **Enhanced Iron Utilization:** Iron-rich plants and vitamin C enhance iron absorption and incorporation into hemoglobin.
3. **Bone Marrow Stimulation:** Certain phytochemicals stimulate hematopoietic stem cells, increasing production of RBCs and WBCs.

#### Comparison with Existing Evidence:

These findings are consistent with previous reports highlighting the hematinic potential of medicinal plants in regions such as Nigeria and India, where plant-based therapies are widely integrated into healthcare systems.

#### Implications:

The results suggest that phytochemical-rich plants may serve as: Affordable alternatives for anemia management in low-resource settings, Adjunct therapies to conventional hematinics, Sources for drug development

## DISCUSSION

This systematic review provides comprehensive evidence that phytochemical-rich plant extracts exert significant hematopoietic effects, as demonstrated by consistent increases in hemoglobin (Hb), packed cell volume (PCV), red blood cell (RBC), and white blood cell (WBC) counts across all included studies. Despite variations in study design and experimental conditions, the direction of effect was uniformly positive, indicating a strong biological association between phytochemical intake and improved hematological parameters.

The magnitude of hemoglobin improvement (approximately 0.6–4.3 g/dL) is clinically meaningful

and suggests potential utility in the management of anemia and related hematological disorders.

#### Hematopoietic Effects

##### Erythropoietic Effects (Hb, PCV, RBC):

The observed increases in Hb, PCV, and RBC across studies strongly indicate enhanced erythropoiesis. Plants such as *Moringa oleifera*, *Telfairia occidentalis*, and *Carica papaya* demonstrated particularly strong effects, likely due to their high content of iron, vitamins, and bioactive phytochemicals.

The parallel rise in Hb and PCV suggests not only increased red blood cell production but also improved red cell mass and oxygen transport capacity. This supports the potential use of these plant extracts as natural hematinics, especially in settings with a high burden of nutritional anemia.

##### Leukopoietic and Immunomodulatory Effects (WBC):

In addition to erythropoiesis, several studies reported significant increases in WBC counts, indicating immune-enhancing properties. This effect was particularly evident in studies involving *Azadirachta indica*, *Withania somnifera*, and *Erythrina velutina*.

The increase in WBC suggests stimulation of leukopoiesis, which may improve host defense mechanisms. This dual effect—hematinic and immunomodulatory—positions phytochemicals as valuable agents in managing both anemia and immune suppression.

##### Hemoglobin (Hb):

Across all studies (Table 2), hemoglobin levels increased in treated groups compared to controls. The magnitude of increase ranged approximately from +0.6 g/dL to +4.3 g/dL, with the most pronounced effects observed in: Adebayo *et al.*, (2022), Adeyemi *et al.*, (2018), Okeke *et al.*, (2020)

This consistent improvement indicates enhanced erythropoiesis and oxygen-carrying capacity, likely driven by improved iron metabolism and erythrocyte survival.

##### Packed Cell Volume (PCV):

PCV values showed parallel increases with hemoglobin, confirming an overall rise in red cell mass. Studies such as Adebayo *et al.*, (2022) demonstrated marked increases (31.9% → 43.2%), suggesting robust hematopoietic stimulation.

##### Red Blood Cells (RBC):

RBC counts increased consistently across studies, reinforcing the conclusion that these plant extracts stimulate bone marrow activity. The increases were particularly notable in studies involving: *Telfairia occidentalis*, *Carica papaya*, *Panax ginseng*.

This supports the hypothesis of direct or indirect activation of erythroid progenitor cells.

### White Blood Cells (WBC):

Increases in WBC counts indicate immunomodulatory effects of phytochemicals. Notably: Silva *et al.*, (2017) showed a strong WBC increase ( $7.1 \rightarrow 10.5 \times 10^9/L$ ), Kumar *et al.*, (2019) and Singh *et al.*, (2021) also reported substantial elevations.

This suggests stimulation of leukopoiesis, potentially enhancing host defense mechanisms.

### Consistency and Variability of Effects:

While the direction of effect was uniformly positive, the magnitude varied across studies. This variability may be attributed to: Differences in plant species and phytochemical composition, Extraction methods (aqueous vs. methanolic vs. ethanolic), Dosage and duration of administration, Differences in animal physiology

This aligns with your earlier heterogeneity finding ( $I^2 \approx 90\%$ ), indicating substantial between-study variability.

**Role of Phytochemical Composition:** The hematopoietic effects observed are closely linked to the phytochemical profiles of the plants studied. Key compounds identified include:

1. **Flavonoids and polyphenols:** Provide antioxidant protection to hematopoietic cells
2. **Alkaloids:** May stimulate bone marrow activity
3. **Saponins:** Enhance nutrient absorption and cellular function
4. **Tannins:** Contribute antioxidant effects but may reduce iron bioavailability at high concentrations
5. **Micronutrients (iron, vitamin C, vitamin A):** Essential for erythropoiesis and hemoglobin synthesis

The synergistic interaction of these compounds likely underlies the consistent improvements in hematological indices.

### Geographical and Ethnopharmacological Context:

The included studies were predominantly conducted in regions such as Nigeria, India, China, and Ghana, where traditional herbal medicine plays a central role in healthcare delivery.

This highlights the importance of ethnopharmacological knowledge in guiding scientific research and supports the integration of traditional medicine into evidence-based healthcare frameworks, particularly in low- and middle-income countries.

### Heterogeneity of Findings:

A key observation from the analysis is the presence of substantial heterogeneity ( $I^2 \approx 89.8\%$ ), indicating considerable variability in effect sizes across studies. This heterogeneity may be attributed to:

1. Differences in plant species and phytochemical composition
2. Variation in extraction methods (aqueous, methanolic, ethanolic)
3. Differences in dosage, treatment duration, and experimental protocols
4. Use of different animal models (rats, mice, rabbits)

The high heterogeneity justifies the use of a random-effects model and suggests that results should be interpreted with caution, particularly when extrapolating to clinical settings.

### Comparison with Existing Literature:

The findings of this review are consistent with previous studies demonstrating the hematinic and immunomodulatory properties of plant-derived compounds. Several reports from Nigeria and India have similarly documented improvements in Hb and RBC indices following administration of phytochemical-rich extracts.

This consistency across studies strengthens the evidence base supporting the role of medicinal plants in hematological health.

### Implications for Research and Practice

The findings suggest that phytochemical-rich plants have significant potential as:

1. Affordable and accessible interventions for anemia, particularly in resource-limited settings
2. Adjunct therapies alongside conventional hematinics
3. Candidates for pharmaceutical development

Future research should focus on:

1. Well-designed human clinical trials
2. Standardization of plant extracts and dosages
3. Investigation of long-term safety and toxicity
4. Elucidation of molecular mechanisms of action

## CONCLUSION

This systematic review demonstrates that phytochemical-rich plant extracts significantly improve hematological parameters, particularly hemoglobin, PCV, and RBC counts, while also enhancing immune function through increased WBC levels. Although heterogeneity and methodological limitations exist, the consistency of positive findings underscores the therapeutic potential of these natural compounds in hematopoietic regulation.

**Declarations**

**Ethics Approval and Consent to Participate:** Not applicable.

**Consent for Publication:** Not applicable.

**Availability of Data and Materials:** All data are included in this published article.

**Competing Interests:** The authors declare no competing interests.

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**Authors' Contributions**

AD conceptualized the study and drafted the manuscript. All authors contributed to data extraction, analysis, and manuscript revision.

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