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MATERNAL NUTRITION AND FETAL RENAL HEALTH – A SYSTEMATIC REVIEW

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Abstract:

Background: The kidneys are fundamental to homeostasis, which underscores how important the kidneys are to develop in fetal life. Renal organogenesis is dependent on maternal nutrition, which impacts fetal health as well as long-term renal function.

Objective: This systematic review aimed to establish a relationship between maternal nutrition and fetal renal health while making distinctions between the macronutrients, micronutrients and breastfeeding and fetal kidney health and development.

Methods: A systematic search of the literature was performed using databases (Pubmed, Scopus and Web of Science), combining keywords like "maternal nutrition," fetal kidney development," and "breastfeeding." Inclusion criteria were developed as per the PICOS framework to include suitable studies that considered maternal dietary factors and outcomes, and more specifically, fetal renal health.

Results: There is evidence that adequate maternal intake of macronutrients (carbohydrates, protein, and healthy fats) is associated with nephrogenesis. Historically, important micronutrients such as folic acid, vitamin D, zinc and magnesium, are important to renal development, and lack of these nutrients is associated with congenital anomalies and increased risk of pediatric and adult chronic kidney disease. Breastfeeding extends the healthy kidneys for neonates, developing a pathway to a lifetime of health. Socioeconomic issues often hinder breastfeeding and other health areas for vulnerable populations. The evidence still is less for this important field.

Conclusion: Maternal nutrition significantly impacts fetal kidney development and overall health. Adequate dietary intake during pregnancy, combined with supportive breastfeeding practices, can mitigate risks of kidney-related congenital defects and chronic diseases later in life. Public health interventions promoting nutritional education, diversity in maternal diets, and breastfeeding are essential for optimizing fetal development and ensuring long-term renal health in children.

Keywords: mother, nutrition, food, fetus, kidney, function

Introduction:

The kidneys represent pivotal organs of the human body, essential for homeostasis with regulating fluid and electrolyte concentrations, correction of acid-base balance, and removal of wastes. Thus, their development-phase, especially during fetal life, is important for prompt renal functioning and may further better long-term aspects of health. The fetal kidneys begin their formation early in gestation, and this complicated formation process ¹can be impacted by variables, among which maternal nutrition seems significant.

Studies suggest an ever so important effect of maternal nutrition on fetal organogenesis and the complex formation of kidneys. Nutrients themselves contribute to cellular differentiation and proliferation as well as the development of organs. Among others, value is placed on the proper balance of these macronutrients: proteins, carbohydrates, and fats-proper for the realization of fetal growth and organ maturation. If there is a lack of adequate intake of any of these, little nephron formation can take place, thereby beginning a cascade that later on will manifest itself as renal problems, such as hypertension and chronic kidney disease-a CKD.

In addition to maternal dietary patterns and overall dietary quality, it is also important to consider dietary diversity and variety in the diet consumed. Higher dietary intakes of fruits, vegetables, lean protein sources, and healthy fats are associated with improved fetal health outcomes, while higher intakes of refined sugars, unhealthy fats, and processed foods can negatively affect fetal growth and development and can have implications for fetal kidney development. Micronutrients, including vitamins and minerals, are also very important for fetal kidney development. Deficits of micronutrients, including folic acid, vitamin D, zinc, and magnesium, have been significantly associated with a variety of congenital kidney defects and kidney function. As a line of evidence continues to emerge that links maternal nutrition and dietary choices to fetal renal health, it becomes important to understand the biological and mechanistic processes attached to dietary decisions during pregnancy. This review is aimed at summarizing current findings on maternal nutrition and its relationship to fetal renal health, as well as emphasizing the previously mentioned need for guidelines and interventions to improve nutrition for pregnant mothers²⁻⁶ to help optimize fetal development and long-term health for children.

Methodology:

A comprehensive literature search was carried out across peer-reviewed databases, such as PubMed, Scopus, and Web of Science. The search strategy used a combination of keywords like "maternal nutrition," "fetal kidney development," "renal health," and "fetal nutrition." The objective was to gather relevant studies that examine the correlation between maternal nutrition and its effect on fetal renal health.

The inclusion criteria followed the PICOS framework, which offers a systematic way of writing down the search parameters:

P (Population): Research conducted on pregnant women, their diet, and nutrition.

I (Intervention): Studies that evaluated maternal nutrients (macronutrients and micronutrients) or eating patterns, including intervention or observational studies that investigated the effect of a particular component of diet on fetal renal growth.

C (Comparison): Studies with comparisons between various eating patterns or nutritional conditions, including sufficient vs. insufficient nutrient intake.

O (Outcome): Studies that documented quantifiable outcomes associated with fetal renal health, including nephron counts, indicators of kidney function, congenital anomalies, or later renal health in offspring.

S (Study Design): Comprised human clinical trials, observational studies, and appropriate animal models but not studies that were aimed at pediatric renal health alone without regard to maternal nutrition. A complete selection of studies fitting these specifications was returned by the search. Later on, the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) reporting guidelines were used in order to provide systematic review transparency, rigor, and reliability. PRISMA is a reporting framework for systematic reviews that highlights key elements like

information flow throughout the research process, study selection, and findings synthesis. This systematic strategy facilitated a strong critique of the literature, making sure that important information concerning maternal nutrition and fetal renal well-being is properly addressed and communicated.

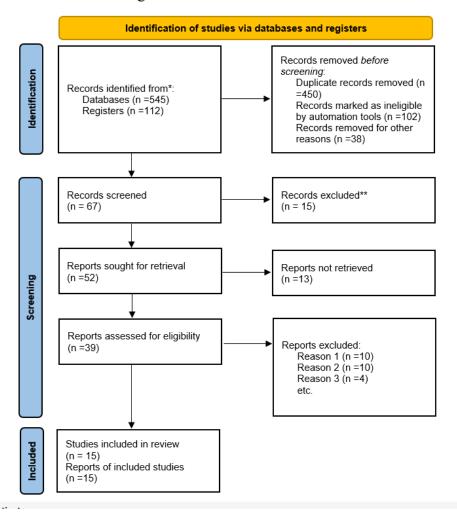


Figure 1 with PRISMA statement.

Results:

Breastfeeding⁷ extends the essential protective benefits in terms of renal health in the neonate and becomes a foundation for lifelong health. Socioeconomic barriers to breastfeeding must be eliminated to maximize its protective effects, particularly on precariously healthy populations.

Macronutrients

Macronutrients such as carbohydrates, proteins, and fats are central to fetal development and the development of organs, including the kidneys. Each macronutrient has a unique and vital function in terms of sustaining the physiological processes required for healthy fetal development.

- 1. Carbohydrates: Carbohydrates are the main energy supply for the mother and the growing fetus. Sufficient intake of carbohydrates will provide the fetus with enough energy to grow, especially at critical developmental stages. Inadequate carbohydrate intake may result in maternal hypoglycemia, which could affect fetal energy resources and compromise organogenesis, such as renal development.
- 2. Proteins: Proteins are necessary for hormone and enzyme synthesis, and for cellular function and structure. Maternal protein-energy malnutrition during pregnancy has been associated with poor fetal kidney development outcomes. Research emphasizes how insufficient protein consumption can result in fewer numbers of nephrons, which are vital for sustaining kidney function through life. A reduced

nephron complement at birth corresponds to an increased risk of hypertension and chronic kidney disease in adulthood.

3. Fats: Healthy fats, especially omega-3 and omega-6 fatty acids, are essential to cell membrane stability and the formation of the central nervous system, which in turn can affect kidney health. Proper intake of fat is crucial for the absorption of fat-soluble vitamins (A, D, E, and K), which play a significant role in all developmental processes, including those occurring in the kidneys.⁹⁻¹¹

Micronutrients

Micronutrients, such as vitamins and minerals, play a major role in fetal development, especially with reference to renal function. Proper maternal consumption of these nutrients is associated with proper kidney maturation and function in the offspring. This section addresses various important micronutrients, their functions in fetal renal development, and the implications of deficiencies according to current studies.

1. Folic Acid

Folic acid, a B-vitamin (B9), plays a role in DNA synthesis, cell division, and fetal growth generally. Deficiency during pregnancy has been shown to be harmful, with research showing a strong correlation between reduced folate levels and congenital malformations, such as renal anomalies. There are reports of folic acid supplementation prior to conception and in early pregnancy decreased the risk of neural tube defects and other congenital abnormalities, ¹²which frequently encompass renal defects. Data from animal models indicate that folate deficiency can cause defects in the proliferation and differentiation of nephron progenitor cells, resulting in decreased nephron number and enhanced vulnerability to kidney diseases in later life.

2. Vitamin D

Vitamin D has a central function in the control of calcium and phosphate metabolism, yet it is not limited to skeletal function. Recent studies point to its essential function in nephrogenesis—nephron development in the fetus. Arshad et al. reported that maternal vitamin D deficiency might result in renal dysplasia in animal models. This dysplasia is evident as structural and functional abnormalities in the kidney, with implications for the regulation of blood pressure and fluid homeostasis in the offspring. In humans, observational research has demonstrated that low maternal serum concentrations of vitamin D are associated with augmented risks for congenital renal anomalies and prenatal developmental defects. These observations highlight the necessity to ensure that pregnant women have proper vitamin D levels for optimal fetal kidney development.

3. Zinc

Zinc is an essential trace mineral for many cellular processes, such as DNA synthesis, protein synthesis, and cell division. Zinc deficiency in pregnancy can negatively impact fetal development, most notably kidney development. It has been established through research that zinc is crucial in nephrogenesis and a deficiency in zinc is linked with defective kidney development. For example, studies have established that pregnancy zinc supplementation enhances fetal outcomes, especially in at-risk populations for micronutrient deficiencies. In animal research, mothers who are deficient in zinc tend to have low nephron numbers and altered renal function in their offspring, potentially laying them open to chronic kidney disease in adulthood.

4. Magnesium

Magnesium is another biochemically important micronutrient that plays a role in numerous biochemical processes, including energy metabolism and enzymatic activity. It is also crucial for the regulation of vascular health and maintenance of normal kidney function. Maternal deficiency in magnesium has been reported to be associated with negative fetal outcomes, such as renal developmental impairment. Evidence indicates that magnesium is involved in the control of cellular signaling pathways involved in nephron development. Studying pregnant women has indicated that

magnesium deficiency would result in abnormal renal vascularization and kidney malformations, as evidenced in animal models. In addition, maternal magnesium deficiency could be responsible for hypertension in offspring and thus increase the risk for adult kidney disease. 13-15

5. Interrelationships and Implications

The interrelationships between these micronutrients are essential to the comprehension of their functions in fetal renal well-being. As an example, folate and vitamin D metabolism can be mediated by levels of magnesium, demonstrating the value of a balanced maternal diet. The simultaneous deficiency of several micronutrients may result in a cumulative risk for fetal renal anomalies.

Omega-3 Fatty Acids

Omega-3 fatty acids, which are polyunsaturated fats, have been recognized for their beneficial effects on health, especially in fetal development and outcome. Essential fatty acids, mostly from fish and fish oil supplements, are involved in vital biological processes such as the regulation of inflammation, cell membrane fluidity, and cardiovascular function. New evidence is emerging pointing to the significance of omega-3 fatty acids in renal development, especially their action on the kidneys.

A number of studies have supported a positive relationship between omega-3 fatty acid consumption in pregnant women and better fetal outcome. One study explained that pregnancy supplementation with omega-3 fatty acids is related to increased renal vasculature development and glomerular function in the developing fetus. A single significant study revealed that maternal omega-3 fatty acid consumption led to elevated renal blood flow as well as enhanced nephron maturation in animal models. More specifically, the children of mothers who were given omega-3 supplements had more functionally mature nephrons, which is essential for long-term kidney health.¹⁶

In addition, evidence indicates that omega-3 supplementation can lower the frequency of kidney problems in offspring. As an example, a randomized controlled trial showed a lowering of hypertension and proteinuria rates in children whose mothers had increased omega-3 consumption during pregnancy. This highlights the possible preventive effect omega-3 fatty acids have against the development of chronic kidney diseases in the future.¹⁷

With these findings, omega-3 fatty acids are a critical element of maternal diet, not just for fetal development but also for establishing long-term kidney health in children. Government health policies that encourage omega-3-rich diets during pregnancy may result in significant improvements in fetal and long-term kidney health outcomes.

Dietary Patterns and Their Impacts on Fetal Kidney Development

Mediterranean Diet

The Mediterranean diet, centered on high intake of fruits, vegetables, whole grains, legumes, nuts, and healthy fats (like olive oil), is well known for its health advantage, especially when pregnant. It has been demonstrated through epidemiological research that a robust correlation exists between the adherence to this eating pattern and improved fetal outcomes, such as improved renal function. Studies observe that the anti-inflammatory and antioxidant properties inherent in the Mediterranean diet can enhance vascular function and optimize placental function, which benefits fetal development.

Several studies emphasize how the common nutrients in the Mediterranean diet, including omega-3 fatty acids and dietary fiber, are associated with better maternal metabolic profiles. This diet has been linked with reduced rates of gestational diabetes and hypertension, which are known to have detrimental effects on fetal kidney development. Second, the high consumption of minerals and vitamins from fruits and vegetables promotes important developmental processes like nephrogenesis, ensuring proper nephron formation. Overall, Mediterranean diet adherence could offset risks of developmental kidney disease, providing a solid foundation for long-term renal function in offspring.

2. Western Diet

The Western diet, on the other hand, with its high intake of sugars, un healthy trans and saturated fats, and processed foods, holds paramount threats to both maternal and fetal health. Studies point out that

such diets are associated with higher rates of gestational diabetes and obesity, both of which can prove harmful to fetal kidney development. Maternal obesity not only impacts nutrient delivery but also affects maternal metabolic activities, compromising placental function and nutrient transport to the fetus.

Excessive intake of detrimental dietary components may cause nephrotoxicity, and there is evidence that excessive sugar and unhealthy fats may directly affect fetal renal function. For example, animal models show that maternal overnutrition during a Western dietary regimen can lead to structural defects in renal tissues and dysfunction in nephron development. Moreover, excessive sugar intake can lead to oxidative stress and inflammation, both of which can impair fetal kidney function and predispose to chronic kidney diseases in adult life.

Briefly, pregnancy diet plays a great role in determining the renal health of the fetus. A Mediterranean diet has protective effects, while a Western diet has considerable risks, highlighting the importance of public health efforts that encourage good maternal nutrition for healthy fetal growth.

The Indian diet:

The Indian diet is heterogeneous and differs greatly from region to region, generally composing a high rate of whole grains, legumes, fruits, vegetables, and spices, with moderate levels of dairy foods and good fats (e.g., mustard or olive oil). In contrast to the Mediterranean and Western diets, some beneficial and adverse characteristics are revealed with respect to fetal renal health.

Positive Factors

Rich in Whole Foods: The Indian diet is big on whole, unprocessed foods, which tend to be higher in vitamins, minerals, and fiber than processed foods seen so often in Western diets. Lentils, beans, and whole grains are great sources of essential nutrients such as folic acid, iron, and magnesium—nutrients essential for fetal development and nephrogenesis.

Spices with Medicinal Properties: Indian traditional spices like ginger and turmeric are known to have anti-inflammatory compounds. Their consumption by pregnant women could improve fetal development and maternal health and result in healthier kidneys for the fetus.

Balanced Macronutrients: Balanced macronutrients obtained from animal as well as plant sources are part of the Indian diet, offering the necessary proteins and good fats. Although conventional diets do lean more towards carbohydrate sources, the addition of legumes and dairy makes it easier to attain a balanced intake. 18-20

Negative Factors

High Salt and Processed Food: In modern-day diets, particularly in cities, diets high in sodium and unhealthy fats from processed food are increasingly being consumed. These types of diets can contribute to possible health complications such as hypertension that can have adverse effects on renal function in the fetus.

Limited Nutritional Needs Awareness: There can be a lack of awareness about specific nutritional requirements during pregnancy in certain areas. This might lead to deficiencies in important micronutrients that are required for proper fetal growth and development. The sample menu given in Table 1 is just a primer and not completely evidence based.

The thrifty phenotype hypothesis:

The thrifty phenotype hypothesis proposes ²¹that early-life environmental factors, particularly undernutrition during fetal development and infancy, can permanently alter an individual's metabolism, increasing their risk of developing chronic diseases such as type 2 diabetes and metabolic syndrome later in life. This happens because the body responds to reduced dietary availability by establishing a "thrifty phenotype" - a metabolic state that efficiently saves energy - which can be deleterious when food is plentiful in adulthood.

Time of Day	Primary Recipe		Prep Time		
	Spinach & Egg Omelet with Whole	Berries & Chia		Protein, Folate,	Eggs and spinach boost folate and choline; chia seeds add omega-3s.
Snack (10–11	Avocado on Rye Crackers with Pumpkin Seeds	Almond Milk,	5–10	Healthy Fats, Zinc,	Avocados and seeds support vascular and kidney tissue development.
Lunch (12–1	Grilled Salmon Bowl with Quinoa & Steamed Broccoli	Chickpea & Sweet	30 min	Protein, Vitamin	Salmon helps nephron development; chickpeas for plant-based iron.
Afternoon Snack (3–4 PM)		Cottage Cheese with Pineapple		Protein, Vitamin	Vitamin C boosts iron absorption; dairy adds B vitamins.
Dinner (6–8		with Bell Peppers	30–40	Folate, Iron,	Lentils and carrots support fetal kidney cell development.
	Warm Turmeric Almond Milk	Nut Butter	5 min	Vitamin D, Healthy Fat	Supports calcium absorption and nighttime fetal growth processes.

Table 1 with sample menu with alternates

Analytical discussion:

The interaction between maternal nutrition and fetal kidney health exemplifies the complex mechanisms of the relationship between diet, gestational outcomes, and future health consequences for the offspring. As illustrated in the abstracts, maternal nutritional status is critical in directing fetal development, especially kidney development, which is critical for homeostasis throughout life. Recent literature has emphasized the adverse effects of inadequate maternal nutrition on the kidney structure and function of offspring, particularly the deficiencies of nutrient intake and/or nutrient status including folate, vitamins A and D and protein. Inadequate maternal nutrition has correlated with a reduced nephron number, related to lifelong kidney function and risk for chronic diseases. The lifetime risk posed by inadequate nephron numbers places the offspring at increased risk of experiencing metabolic syndromes and kidney dysfunction as an adult, and emphasizes the need for early nutritional interventions.

In addition, certain studies note the significance of carbohydrate quality in pregnancy. A nutrition pattern comprised of higher amounts of low-quality carbohydrates during pregnancy can worsen pregnancy outcomes, along with facilitate unfavorably metabolic changes in offspring. This is in line with a new body of research suggesting to consider a diet pattern rather than a specific nutrient. Healthcare professionals can tailor holistic intervention strategies of maternal nutrition and improve pregnancy related health outcomes with a dietary pattern approach. Regarding the positive aspects of the unique dietary patterns discussed, the whole foods fundamental to an Indian diet, the overall higher

amounts fibrous fruits, vegetables, and legumes could improve overall intake of food components important for fetal growth & development (Indian Diet Section). Traditional Indian spices that had anti-inflammatory potential might also provide a modest additional benefit for maternal health & fetal outcomes (Indian Diet Section). However, we also noted Western food habits such as salt and processed foods, and the dual nature of dietary patterns where traditional Indian diets are impacted by the new way of life and affect health outcomes, especially in urban landscapes with few dietary principles.

The review of omega-3 fatty acids in suggests a nutrient deficiency that is widespread among pregnant women because they are restricted from eating seafood. This highlights the urgent need for public health strategies for encouraging a wide variety of sources of omega-3 in maternal diets to support both fetal neurodevelopment and overall health. Further complicating maternal nutrition is the additive effect of oxidative stress that was discussed. High-fat diets that negatively impact placental function raise questions about dietary choice and the immediate and downstream impact on fetal development.

To summarize, maternal nutrition has an essential and direct effect on the fetal kidney development and kidney health overall. Sufficient levels of macronutrients and micronutrients are necessary to promote nephrogenesis and to reduce the chance of congenital kidney defects and chronic disease in offspring. Dietary patterns, such as the Mediterranean diet, are beneficial, while western diets are detrimental. Along with the importance of varied and nutrient-rich maternal diets, public health initiatives should aim to improve knowledge and access to appropriate nutrition during pregnancy, thereby establishing the foundation for children's healthy growth and long-term kidney health.

All the authors have contributed significantly.

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