

Nutritional Approaches In The Prevention And Management Of Hepatocellular Carcinoma: A Review

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ABSTRACT

Hepatocellular carcinoma (HCC) is one of the leading causes of cancer-related mortality globally. Emerging evidence indicates that dietary interventions can play a significant role in both the prevention and management of HCC, particularly among high-risk populations. This review explores the impact of nutritional factors on HCC progression, focusing on the protective roles of antioxidants, polyphenols, and specific functional foods. High intake of fruits, vegetables, omega-3 fatty acids, and plant-derived compounds has been associated with anti-carcinogenic effects, while excessive consumption of red meat, sugar, and saturated fats may elevate HCC risk. Furthermore, selected juices, herbal extracts, and bioactive food combinations exhibit hepatoprotective and anti-tumor properties in both in vitro and in vivo studies. This comprehensive review underscores the potential of evidence-based dietary strategies in reducing liver cancer burden and improving the prognosis of HCC patients.

Keywords: Hepatocellular carcinoma, nutrition, antioxidants, functional foods, liver cancer, phytochemicals.

1. INTRODUCTION

Hepatocellular carcinoma (HCC) is the most common primary malignancy of the liver and ranks among the leading causes of cancer-related deaths worldwide. Its incidence is rising steadily, particularly in regions experiencing a surge in obesity, type 2 diabetes, and metabolic syndrome—all of which are closely linked to nonalcoholic fatty liver disease (NAFLD) and its progressive form, nonalcoholic steatohepatitis (NASH) [1,2]. While viral hepatitis (HBV, HCV) and alcohol abuse remain prominent etiological factors, increasing attention has been directed toward modifiable lifestyle determinants, particularly dietary patterns, as key contributors to liver cancer risk [3–5].

Emerging evidence suggests that dietary components may play a dual role in both the prevention and progression of HCC. Certain foods rich in antioxidants, fiber, polyphenols, and unsaturated fatty acids exhibit hepatoprotective effects by modulating inflammation, oxidative stress, and insulin resistance—factors known to contribute to hepatic carcinogenesis [6–9]. In contrast, high consumption of red and processed meats, sugar-sweetened beverages, and refined carbohydrates has been associated with an increased risk of HCC through pathways involving lipotoxicity, chronic inflammation, and gut microbiome dysbiosis [10–12].

Beyond prevention, nutritional status significantly influences treatment outcomes and survival in patients diagnosed with HCC. Malnutrition, sarcopenia, and micronutrient deficiencies are prevalent in this population and are associated with poorer prognosis and treatment tolerance [13–15]. Therefore, optimizing diet quality may not only reduce the risk of developing HCC but also support improved clinical outcomes in patients undergoing therapy.

Given the growing burden of HCC and the potential of dietary strategies as cost-effective, non-invasive interventions, a comprehensive understanding of the relationship between diet and liver cancer is essential. This review aims to synthesize current evidence on dietary patterns, specific nutrients, and functional foods associated with HCC risk and prognosis, highlighting potential dietary recommendations for clinical and public health implementation.

1.1 DIETARY PATTERNS AND RISK OF HEPATOCELLULAR CARCINOMA

Epidemiological evidence suggests that dietary patterns significantly influence the risk of HCC. A systematic review by George et al. (2021) found that diets rich in fruits, vegetables, whole grains, and unsaturated fats are associated with reduced HCC risk, while high intake of processed foods, red meat, and sugary beverages is linked to increased incidence [1]. Similarly, Shu et al. (2023) performed a meta-analysis which concluded that the Western dietary pattern (characterized by high meat, fat, and sugar intake) elevates HCC risk, whereas the Mediterranean diet offers protective effects [2].

Baharvash et al. (2023) confirmed that consumption of whole grains, legumes, and green leafy vegetables was inversely associated with HCC risk, while diets high in saturated fats, processed meat, and alcohol showed a strong positive association [3]. This suggests that modifying dietary habits may play a crucial preventive role.

1.2 INFLUENCE OF SPECIFIC NUTRIENTS AND FOOD GROUPS

Márquez et al. (2024) reviewed how dietary components such as antioxidants, flavonoids, and polyphenols modulate liver carcinogenesis. The review emphasized the benefits of micronutrients like vitamin C, vitamin E, and selenium in reducing oxidative damage in hepatic tissues [4].

Coffee consumption, both caffeinated and decaffeinated, has been extensively studied. Kennedy et al. (2017) conducted a dose-response meta-analysis showing a significant inverse relationship between coffee intake and HCC risk, potentially due to chlorogenic acid and diterpenes that modulate liver enzyme activity [5].

Additionally, lemon juice, rich in vitamin C and flavonoids, demonstrated hepatoprotective effects in experimental models. Oyedepo et al. (2015) found that a lemon-sorghum (ogi) mixture could significantly restore liver function markers in rats with paracetamol-induced liver damage [6].

1.3 DIETARY INFLAMMATORY INDEX AND LIVER CANCER

Chronic inflammation is a key factor in liver carcinogenesis. Shetty et al. (2024) reported that a high Dietary Inflammatory Index (DII)—indicative of pro-inflammatory diets—was associated with a significantly higher risk of HCC, especially among cirrhotic patients [7]. Diets high in refined carbohydrates, red meat, and saturated fats elevate inflammatory markers like C-reactive protein, which may accelerate liver fibrosis and cancer progression.

2. NUTRITIONAL STATUS IN HCC PATIENTS

Poor nutritional status is common in HCC patients due to metabolic disturbances, anorexia, and systemic inflammation. Gamal et al. (2025) conducted a cross-sectional study which highlighted deficiencies in protein, zinc, and vitamins A, D, and E in HCC patients, underlining the need for tailored nutritional interventions [8].

2.1 ANTIOXIDANT AND HEPATOPROTECTIVE PROPERTIES OF PLANT EXTRACTS

Several plant-based foods have shown hepatoprotective and anticancer properties. For instance, Liu et al. (2006) demonstrated that *Bupleurum kaoi* leaf infusion had potent antioxidative and liver-protective activity in vitro [9]. Rambutan fruit extract exhibited cytotoxic effects on HepG2 liver cancer cells, indicating its chemotherapeutic potential (Perumal et al., 2021) [10].

Garcinia dulcis [12], *Vismia* species [15], and *Opuntia* fruits [28] have also been studied for their polyphenol-rich content and their ability to induce apoptosis in liver cancer cells. Green tea polyphenols (Darvesh & Bishayee, 2013) and curcumin from turmeric are widely recognized for their anti-inflammatory and anticancer actions in hepatic tissues [16, 34].

2.2 FERMENTED AND FUNCTIONAL FOODS

Fermented foods such as noni juice and Moringa tea have gained attention for their role in liver health. Guo et al. (2020) demonstrated that fermented noni juice attenuated alcohol-induced liver damage by enhancing antioxidant enzyme activity [17]. Similarly, Elzamzamy&Elkewawy (n.d.) showed that untreated fruit juice supplemented with Moringa tea exerted hepatoprotective effects against carbon tetrachloride (CCl₄) toxicity in rats [27].

2.3 CITRUS AND FRUIT JUICES IN LIVER PROTECTION

Citrus fruits like lemon, orange, and mandarin are rich in vitamin C and flavonoids. Bekkouch et al. (2022) found that a combination of lemon and ginger juices significantly reduced oxidative liver injury in rats [23]. Sugiura et al. (2006) demonstrated that Satsuma mandarin consumption improved hepatic antioxidant status in diabetic rat models [24].

3. HARMFUL DIETARY COMPONENTS

High intake of red and processed meats, saturated fats, and alcohol has been associated with an elevated risk of HCC. Talamini et al. (2006) [25] and Freedman et al. (2010) [33] found that animal fat and meat consumption significantly raised liver cancer risk in European and American cohorts. Zhang et al. (2007) warned against star fruit juice in hepatic-compromised individuals due to its inhibitory effects on liver cytochrome P450 enzymes [22].

3.1 WEIGHT LOSS AND LIFESTYLE MODIFICATION

Weight loss and lifestyle changes have been found effective in reducing liver fat and fibrosis, which are precursors to HCC. Vilar Gomez et al. (2015) demonstrated that a 10% weight reduction in obese NAFLD patients significantly improved liver enzyme profiles and inflammatory markers [29]. Incorporating plant-based diets, exercise, and behavioral changes is essential in HCC prevention and management.

3.2 SUMMARY OF NATURAL ANTICANCER AGENTS

Natural dietary agents like saffron [30], phloretin [14], garlic sulfur compounds [32], and flavonoids from herbal extracts [34] exhibit strong anticancer potential by inhibiting cell proliferation, inducing apoptosis, and reducing inflammation in hepatic tissues.

Table 1: Recommended Dietary Components for Liver Cancer (HCC) Patients

Food Item / Component	Key Benefits	Supporting Studies
Coffee (Regular/Decaf)	Antioxidant, anti-inflammatory, reduces HCC risk	[1], [4], [5]
Citrus Fruits (Lemon, Orange)	Liver detoxification, antioxidant, vitamin C	[6], [23], [24]
Garlic	Anticancer sulfur compounds, enhances detox enzymes	[13], [31], [32]
Green Tea	Rich in polyphenols, anti-proliferative properties	[16], [26]
Saffron	Potent anticancer properties	[30]
Berries / Grapes	Rich in resveratrol and anthocyanins	[1], [34]
Vegetables (especially cruciferous)	Contains indoles and glucosinolates for liver protection	[3], [4], [35]
Whole Grains	High fiber, reduces inflammation and insulin resistance	[2], [7], [29]
Fermented Foods (e.g., noni juice)	Antioxidant, probiotic potential	[17], [27]
Legumes & Pulses	Lowers inflammatory markers, supports liver function	[1], [2], [8]
Omega-3 Rich Foods (Flaxseed, Fish)	Anti-inflammatory, may slow tumor growth	[4], [34], [35]
Turmeric / Curcumin	Anti-inflammatory and anti-cancer agent	[11], [34]
Water / Hydration	Essential for liver detoxification and function	Clinical observation

4. CONCLUSION

Nutritional status and dietary composition are emerging as significant modifiable risk factors in the prevention and management of hepatocellular carcinoma (HCC). Numerous studies have identified specific macronutrients and micronutrients—including fiber, polyphenols, antioxidants, and unsaturated fatty acids—as being inversely associated with the risk of HCC. Dietary components such as fruits, vegetables, legumes, coffee, and green tea have demonstrated protective effects, while high intake of red/processed meats, saturated fats, refined carbohydrates, and sugar-sweetened foods has been linked to increased risk.

In individuals with underlying liver disorders, particularly non-alcoholic steatohepatitis (NASH), dietary intervention plays a crucial role in halting disease progression to cirrhosis and ultimately HCC. Optimizing the metabolic profile through targeted nutritional strategies can enhance both prevention and prognosis.

Despite promising observational evidence, there remains a pressing need for well-structured, prospective interventional studies to validate the role of specific dietary modifications in reducing HCC incidence and improving overall survival in affected patients. Future research should focus on the integration of nutritional therapies into comprehensive clinical guidelines for HCC risk reduction and patient care.

nutritional status and food composition are important risk factors for HCC. They also play a significant role in the prognosis

of HCC patients. Several macronutrient and micronutrient components have been discovered to be inversely associated to the risk of HCC. It is critical to improve the metabolic state in individuals with non-alcoholic steatohepatitis in order to avoid disease progression to liver cirrhosis or HCC. However, data from well-designed perspective interventional trials aimed at reducing HCC incidence or extending survival in HCC patients based on dietary adjustment is still needed.

REFERENCES

- [1] E. S. George, S. Sood, A. Broughton et al., “The Association between Diet and Hepatocellular Carcinoma: A Systematic Review,” *Nutrients*, vol. 13, 2021, p. 172, doi:10.3390/nu13010172
- [2] W. Shu, L. Liu, J. Jiang, Q. Yao, “Dietary Patterns and Hepatocellular Carcinoma Risk: A Systematic Review and Meta-Analysis of Cohort and Case-Control Studies,” *Nutrition & Metabolism*, Aug 2023.
- [3] A. Baharvash et al., “Analysis of Specific Food Groups and Liver Cancer Risk: Systematic Review & Meta-Analysis,” *Nutrition Journal*, 2023.
- [4] A. Márquez et al., “The Influence of Diet and Its Components on the Risk of Hepatocellular Carcinoma: A Review,” *Cancers (MDPI)*, vol. 16, no. 5, 2024, Art. 1030.
- [5] Kennedy, O. J., Roderick, P., Buchanan, R., Fallowfield, J. A., Hayes, P. C., & Parkes, J. (2017). Coffee, including caffeinated and decaffeinated coffee, and the risk of hepatocellular carcinoma: a systematic review and dose–response meta-analysis. *BMJ open*, 7(5), e013739.
- [6] Oyedepo, T. A., Ajayeoba, T. A., Babarinde, S. O., Morakinyo, A. E., & Oyetayo, B. O. (2015). Antioxidant and hepatoprotective potentials of lemon juice and sorghum ogi (lemon-ogi) mixture against paracetamol-induced liver damage in rats. *health*, 38.
- [7] G. Shetty et al., “Association between Dietary Inflammatory Index and HCC Risk in Cirrhosis,” *Eur. J. Cancer Prev.*, 2024.
- [8] E. Gamal et al., “Assessment of the Nutritional Status of HCC Patients: Cross-Sectional Study,” *Egyptian Liver Journal*, vol. 15, Art. 19, Jun 2 2025.
- [9] Liu, C. T., Chuang, P. T., Wu, C. Y., Weng, Y. M., Chen, W., & Tseng, C. Y. (2006). Antioxidative and in vitro hepatoprotective activity of Bupleurum kaio leaf infusion. *Phytotherapy Research: An International Journal Devoted to Pharmacological and Toxicological Evaluation of Natural Product Derivatives*, 20(11), 1003-1008.
- [10] Perumal, A., AlSalhi, M. S., Kanakarajan, S., Devanesan, S., Selvaraj, R., & Tamizhazhagan, V. (2021). Phytochemical evaluation and anticancer activity of rambutan (*Nephelium lappaceum*) fruit endocarp extracts against human hepatocellular carcinoma (HepG-2) cells. *Saudi Journal of Biological Sciences*, 28(3), 1816-1825.
- [11] A. C. Choudhary et al., “Emerging Antioxidants in Cancer Therapy: Clinical & Experimental Review,” *ArXiv*, 2020.
- [12] Abu Bakar, M. F., Ahmad, N. E., Suleiman, M., Rahmat, A., & Isha, A. (2015). *Garcinia dulcis* fruit extract induced cytotoxicity and apoptosis in HepG2 liver cancer cell line. *BioMed Research International*, 2015.
- [13] Arnault, I., Haffner, T., Siess, M. H., Vollmar, A., Kahane, R., & Auger, J. (2005). Analytical method for appreciation of garlic therapeutic potential and for validation of a new formulation. *Journal of pharmaceutical and biomedical analysis*, 37(5), 963-970.
- [14] Ma, L., Wang, R., Nan, Y., Li, W., Wang, Q., & Jin, F. (2016). Phloretin exhibits an anticancer effect and enhances the anticancer ability of cisplatin on non-small cell lung cancer cell lines by regulating expression of apoptotic pathways and matrix metalloproteinases. *International Journal of Oncology*, 48(2), 843-853.
- [15] Lizcano, L. J., Siles, M., Trepiana, J., Hernández, M. L., Navarro, R., Ruiz-Larrea, M. B., & Ruiz-Sanz, J. I. (2015). Piper and Vismia species from Colombian Amazonia differentially affect cell proliferation of hepatocarcinoma cells. *Nutrients*, 7(1), 179-195.
- [16] Darvesh, A. S., & Bishayee, A. (2013). Chemopreventive and therapeutic potential of tea polyphenols in hepatocellular cancer. *Nutrition and cancer*, 65(3), 329-344.
- [17] Guo, M., Mao, B., Sadiq, F. A., Hao, Y., Cui, S., Yi, M., ... & Zhao, J. (2020). Effects of noni fruit and fermented noni juice against acute alcohol induced liver injury in mice. *Journal of Functional Foods*, 70, 103995.
- [18] Yantih, N., Harahap, Y., Sumaryono, W., Setiabudy, R., & Rahayu, L. (2017). Hepatoprotective activity of pineapple (*Ananas comosus*) juice on isoniazid-induced rats. *Journal of Biological Sciences*, 17(8), 388-393.
- [19] “Hepatocellular Carcinoma: Nature Rev. Disease Primers 2020,” *Nature Rev. Dis. Prim.*, Apr 2020.
- [20] “Lifestyle & Pharmacologic Approaches to Prevent Metabolic-Associated Liver Disease,” *ScienceDirect*, 2025.

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- [21] Zhang, J. W., Liu, Y., Cheng, J., Li, W., Ma, H., Liu, H. T., ... & Yang, L. (2007). Inhibition of human liver cytochrome P450 by star fruit juice. *Journal of Pharmacy & Pharmaceutical Sciences*, 10(4), 496-503.
- [22] Bekkouch, O., Dalli, M., Harnafi, M., Touiss, I., Mokhtari, I., Assri, S. E., ... & Amrani, S. (2022). Ginger (*Zingiber officinale* Roscoe), Lemon (*Citrus limon* L.) Juices as Preventive Agents from Chronic Liver Damage Induced by CCl₄: A Biochemical and Histological Study. *Antioxidants*, 11(2), 390.
- [23] Sugiura, M., Ohshima, M., Ogawa, K., & Yano, M. (2006). Chronic Administration of Satsuma Mandarin Fruit (*Citrus unshiu* M ARC.) Improves Oxidative Stress in Streptozotocin-Induced Diabetic Rat Liver. *Biological and Pharmaceutical Bulletin*, 29(3), 588-591.
- [24] P. Talamini et al., "Food groups and risk of hepatocellular carcinoma: A multicenter case-control study in Italy," *Int. J. Cancer*, vol. 119, pp. 2916-2921, 2006.
- [25] "Green Tea for Cancer Prevention: Cochrane Review," *Cochrane Database Syst. Rev.*, 2020 .
- [26] Elzamzamy, F. M., &Elkewawy, H. E. Hepatoprotective effect of untreated fruit juice supplemented with moringa tea on CCl₄-induced damage in rats.
- [27] González-Ponce, H. A., Martínez-Saldaña, M. C., Rincón-Sánchez, A. R., Sumaya-Martínez, M. T., Buist-Homan, M., Faber, K. N., ... & Jaramillo-Juárez, F. (2016). Hepatoprotective effect of *Opuntia robusta* and *Opuntia streptacantha* fruits against acetaminophen-induced acute liver damage. *Nutrients*, 8(10), 607.
- [28] G. Vilar-Gomez et al., "Weight Loss Lifestyle Impact on NAFLD Biomarkers," *Gastroenterology*, Aug 2015
- [29] Amin, A., Hamza, A. A., Bajbouj, K., Ashraf, S. S., & Daoud, S. (2011). Saffron: a potential candidate for a novel anticancer drug against hepatocellular carcinoma. *Hepatology*, 54(3), 857-867.
- [30] Arnault, I., Haffner, T., Siess, M. H., Vollmar, A., Kahane, R., & Auger, J. (2005). Analytical method for appreciation of garlic therapeutic potential and for validation of a new formulation. *Journal of pharmaceutical and biomedical analysis*, 37(5), 963-970.
- [31] Iciek, M., Kwiecień, I., Chwatko, G., Sokołowska-Jezewicz, M., Kowalczyk-Pachel, D., & Rokita, H. (2012). The effects of garlic-derived sulfur compounds on cell proliferation, caspase 3 activity, thiol levels and anaerobic sulfur metabolism in human hepatoblastoma HepG2 cells. *Cell biochemistry and function*, 30(3), 198-204.
- [32] N. Freedman et al., "Association of meat and fat intake with liver disease and hepatocellular carcinoma in the NIH-AARP cohort," *J. Natl. Cancer Inst.*, vol. 102, pp. 1354-1365, 2010.
- [33] Huang, W. Y., Cai, Y. Z., & Zhang, Y. (2009). Natural phenolic compounds from medicinal herbs and dietary plants: potential use for cancer prevention. *Nutrition and cancer*, 62(1), 1-20.
- [34] Sacco, R., Conte, C., Marceglia, S., Mismas, V., Bresci, G., Romano, A., & Giacomelli, L. (2016). Beneficial and detrimental effects of natural dietary products on the risk of hepatocellular carcinoma, and their roles in its management. *Hepatoma Research*, 2, 53-61.
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