The black pepper \( (Piper nigrum \text{ L}) \) vine and its extracts have been used as a folk medicine in a variety of cultures and are the source of the most commonly used spice worldwide. The chemical piperine is a major bioactive component present in black pepper (and white pepper as well) that has numerous reported physiological and drug-like actions. The scientific literature provides evidence that black pepper may have health benefits, particularly in enhancing digestive tract function. There is suggestive evidence that black pepper piperine may have nervous system benefits and may influence body energy usage in rats. Preliminary evidence in cell culture studies suggests that black pepper contains antioxidant constituents and possesses anti-inflammatory and antimicrobial properties. An overview of major uses for black pepper is presented here, and the strength of the evidence is graded. Nutr Today. 2010;45(1):43–47

Black pepper from the \( (Piper nigrum \text{ L}) \) vine (Figure 1) is the most commonly used spice worldwide, and its extracts have been used as a folk medicine in a variety of cultures. \( Piper nigrum \) is used for the production of both black pepper (from the unripened fruit) and white pepper (from the dehulled mature berry). In ancient Sanskrit literature, black pepper use for medicinal purposes was documented. In India, it was one of the most commonly used herbs in Ayurvedic medicine and has been considered for treatment of gastrointestinal disorders and, even more recently, of chronic malaria. Black pepper also was used for the treatment of epilepsy in traditional Chinese medicine. In the Middle Ages, black pepper was recorded as being used in seasoning and for concealing the flavor of salted, cured meat. This plant is indigenous to Southern India and also is cultivated on the islands of the Malay archipelago, as well as Madagascar and other islands off the coast of Africa. Popular names for this spice depend on the source. For example, Malabar black pepper comes from the Malabar coast of southwest India, and Lampong black pepper is from Sumatra, Indonesia. Black pepper currently finds multiple uses in flavorings, in perfumes, and in insecticide formulations.\(^1\text{-}\,^3\)

Overview

Black pepper constituents include fiber, essential oils, piperine, eugenol, the enzyme lipase, and minerals. Essential oil components include \( \alpha \) - and \( \beta \)-pinene, limonene, and \( \beta \)-caryophyllene.\(^2\text{-}\,^4\) Piperine and its isomers are the major factors responsible for the pungency and irritant action of black pepper. The chemical piperine, 1-piperoylpiperidine, is the major bioactive component present in both black and white peppers and individually has numerous reported physiological and drug-like actions similar to those reported for black pepper. The scientific literature provides evidence that black pepper may have health benefits, particularly in enhancing digestive tract

Figure 1. Black peppercorns.
function. It has been reported, for example, in rats and mice that black pepper and piperine can stimulate digestive enzymes, modify stomach secretions, alter gastrointestinal food transit time, and inhibit diarrhea.\textsuperscript{3–10} The acute effect of black pepper on the human stomach seems to be similar to that for aspirin, although the long-term effect of black pepper on the stomach is unknown.\textsuperscript{11}

Black pepper also has a substantial effect on enzyme systems that metabolize phytochemicals and drugs. Black pepper and piperine have been observed to inhibit cytochrome P450 enzymes, other phase I enzymes, the phase II detoxification enzyme uridine-5'-diphospho-(UDP)-glucuronyl transferase, and drug transporters in several tissues.\textsuperscript{9,12–18} This can lead to pronounced changes in the bioavailability of a natural compound or drug after ingestion. For example, coadministration of black pepper to rats along with curcumin, a phenolic constituent of turmeric, interferes with conjugation and metabolism of curcumin by UDP-glucuronyl transferase. Consequently, increased blood levels of curcumin have been noted in rats and humans.\textsuperscript{19} A similar increase in circulating levels of bioactive compounds associated with black pepper and piperine has been reported for other dietary factors such as the flavonoid (−)-epigallocatechin-3-gallate, beta carotene, and coenzyme Q\textsubscript{10}.\textsuperscript{20–23} In healthy human volunteers, piperine administration increased blood levels of the antiepileptic drug phenytoin, the antihypertensive drug propranolol, and theophylline, a drug used to treat respiratory conditions.\textsuperscript{24–27} The clinical consequences of concomitant black pepper intake on dietary factors and drugs, particularly those with narrow therapeutic windows, will likely depend on multiple factors, including the dose of black pepper and drugs, the chemical characteristics of the agents, and the individual physiological responses to the specific agent.\textsuperscript{28}

There is suggestive evidence that black pepper and piperine may have nervous system benefits and may influence body energy usage in rats. Preliminary evidence in cell culture studies suggests that black pepper contains antioxidant constituents and possesses anti-inflammatory and antimicrobial properties. Likewise, anticancer actions have been observed.

Examples of these uses for black pepper are presented in the “Summary of Research” section, and an effort is made to give an overview of the variety of scientific research on this topic. Points of view for rating of evidence in each category are based on consideration of cell culture, animal, and human clinical data from the peer-reviewed scientific literature. A higher rating was given when there were both preclinical and clinical data and there was consistency of findings among well-controlled human studies.

### Summary of Research

#### Potential Health Benefits of Black Pepper

<table>
<thead>
<tr>
<th>Scientific Findings for Select Uses</th>
<th>Rating</th>
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<tr>
<td>Alteration of digestive tract function: Black pepper (and its constituent piperine) is reported to act as a digestive tract stimulant in rats and mice, although the effect is not always consistent.\textsuperscript{5–9} In small clinical studies, intake of black pepper extract stimulated stomach secretions and affected the rapidity of movement of food through the digestive tract.\textsuperscript{10} Some of this effect of black pepper seems to be due to piperine activating specific drug receptors.\textsuperscript{26} Black pepper also may alter the bioavailability of certain food components and drugs, partly by altering the body’s systems controlling the metabolism and absorption of dietary constituents and drugs.\textsuperscript{20–25} This effect of black pepper has been reported to improve the bioavailability of some agents, although the clinical consequences may not necessarily be beneficial.</td>
<td>Emerging, suggestive</td>
</tr>
<tr>
<td>Metabolism and obesity: Piperine has been shown in animal studies to increase the body’s expenditure of energy. Apparently, piperine does this by affecting the production of hormone-like chemicals that regulate energy balance. This may have important implications for human body weight regulation and obesity, although, to date, there is little evidence to support such a benefit in humans.\textsuperscript{27}</td>
<td>Preliminary, inconclusive</td>
</tr>
<tr>
<td>Nervous system benefits: In mice, an extract of black pepper exhibited activity in suppressing convulsions.\textsuperscript{28–31} In one human study, inhalation of black pepper oil components improved the swallowing reflex in stroke patients, apparently by activating specific regions of the brain.\textsuperscript{32} A novel effect of inhalation of black pepper extract was the stimulation of respiratory tract sensations that apparently alleviated smoking withdrawal symptoms.\textsuperscript{33}</td>
<td>Preliminary, inconclusive</td>
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</table>
Animal studies have evaluated the toxicity of black pepper or piperine and report mixed results. One study observed that black pepper caused acute toxicity in rats and mice, whereas 2 studies reported no adverse effects of black pepper administration to rats. The reason for these disparate results is unknown, and it should be pointed out that the doses used were not representative of usual human intakes. Most reports suggest that black pepper and piperine are not genotoxic or immunotoxic.

Black pepper is considered Generally Recognized as Safe (GRAS) for use in foods as a flavoring agent. This Generally Recognized as Safe status of black pepper for use as a food flavoring agent reflects doses that are much lower than those used in animal and cell culture studies and is thus not meant to justify unrestricted use at these higher levels. In this regard, it has been estimated that consumption of black pepper in India is about 0.3 g/d, a value similar to that for Americans estimated from disappearance data published by the US Department of Agriculture. This amount is several-fold lower than those amounts of black pepper given in human studies (about 1.5 g/d). Interestingly, piperine has been administered in some human trials at levels of 5 to 20 mg, which, assuming piperine content is 6% of black pepper dry weight, is approximately the amount consumed in 83 to 333 mg of black pepper.

### Conclusions

Based on the current scientific evidence, more information is needed on the health benefits of black pepper, particularly in human subjects. There is suggestive evidence to support its benefit in improving the function of the digestive tract and increasing energy expenditure. More evidence from well-designed clinical trials is needed, particularly those that examine black pepper doses relevant to typical intakes.

There are numerous cell culture studies demonstrating the antioxidant effects of black pepper and its extracts and that it has anti-inflammatory properties. However, the bioavailability of these constituents of black pepper, once ingested by humans, and the subsequent improvement in appropriate biomarkers in humans are not well characterized.

There is very limited evidence that black pepper constituents may benefit symptoms of convulsions and stroke. Additional controlled human studies using several dietary doses of black pepper components are needed to confirm these purported and intriguing health benefits. Also, drug interactions after intakes of a wide range of black pepper and piperine doses need to be more carefully evaluated, so that interactions with medications can be better understood.
Thus, black pepper will continue to be a popular spice for enhancing food flavor, but its use in improving human health must await more definitive scientific evidence.

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