

REVIEW

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Research progress in the phytochemistry and biology of *Ilex* pharmaceutical resources

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KEY WORDS

Ilex; Chemical components; Pharmacology; Chemotaxonomy; Phylogeny; Pharmaceutical and food resource **Abstract** *Ilex* is a botanical source for various health-promoting and pharmaceutically active compounds that have been used in traditional Chinese medicine and food for thousands of years. Increasing interest in *Ilex* pharmaceutical and food resources has led to additional discoveries of terpenoids, saponins, polyphenols (especially flavonoids), glycosides, and many other compounds in various *Ilex* species, and to investigation of their chemotaxonomy, molecular phylogeny and pharmacology. In continuation of our studies on *Ilex* pharmacology and phylogeny, we review the phytochemistry, chemotaxonomy, molecular biology and phylogeny of *Ilex* species and their relevance to health-promotion and therapeutic efficacy. The similarity and dissimilarity between *Ilex paraguariensis*, the source plant of mate tea, and the source plants of large-leaved Kudingcha (e.g., *Ilex kudingcha* and *Ilex latifolia*) are discussed. It is essential to utilize emerging technologies in non-*Camellia* tea studies to promote the sustainable utilization of *Ilex* resources and the identification and development of novel compounds with potential health and clinical utility. Systems biology and "-omics" technologies will play an increasingly important role in pharmaceutical and food research on the bioactive compounds of *Ilex* species.

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1. Introduction

Ilex, whose common name is holly, is the only living genus of almost 600 species in the family Aquifoliaceae¹. The species are evergreen and deciduous trees, shrubs and climbers from tropics to temperate regions worldwide. Various Ilex species are utilized by worldwide ethnic groups to make non-Camellia tea drinks for daily consumption and health promotion². Mate tea from *I. para*guariensis originated from the southern part of South America (Brazil, Argentina, Uruguay and Paraguay) and is now a popular health-promoting drink in western countries³. Large-leaved Kudingcha (bitter spikeleaf tea) is made from I. kudingcha and I. latifolia and has been consumed as a functional food in southern China for about 2000 years² (Table 1). Various compounds have been isolated from Ilex plants since the 1980's, and their chemistry and pharmacology has been reported^{2,4}. Polyphenol constituents of green tea, mate tea and large-leaved Kudingcha are listed in Table 2. Studies on the chemotaxonomy and molecular phylogeny have been carried out to facilitate further conservation and exploitation of *Ilex* pharmaceutical and food resources^{1,5}. In this brief review, we summarize the recent progress in phytochemical and biological research of *Ilex* plants.

2. Molecular taxonomy, molecular phylogeny, and genomics

The evolutionary history and diversification of the genus *Ilex*, comprised of 108 species, is deduced by analyzing two nuclear (ITS and *nepGS*) and three plastid (*rbcL*, *trnL-F* and *atpB-rbcL*) sequences¹. Nuclear and plastid trees are highly dissimilar and the nuclear tree is more attuned with current taxonomic classifications. Many Chinese species, including *Ilex cornuta*, *Ilex zhejiangensis*, *Ilex integra*, *Ilex hylonoma*, *I. latifolia*, *Ilex pernyi*

Name	Original region	Use history	Main application region	Traditional use		
<i>Camellia sinensis</i> (green tea)	South of the Yang Tze River, China	Around 3000 years	Worldwide, raw materials for drink, food, cosmetic and health care products	Sweet and bitter taste, cool in property, refreshing, reduce stress and thirst, reduce phlegm, help digestion, diuresis, detoxify, treat headache, blurred vision, somnolence, dyspepsia, dysentery		
Ilex paraguariensis (Mate tea)	South America	300–400 Years	drink and health care products in America, Europe and Asia	Hepatic protection, help digestion, anti-rheumatism, arthritis and inflammation, anti-obesity, hypertension and hypercholesterolemia		
I. latifolia, I. kudingcha (large-leaved Kudingcha)	South of the Yang Tze River, China	Around 2000 Years	Tea drink in China and southeast Asia	Sweet and bitter taste, cold, liver meridian, lung meridian and stomach meridian, heat clearing, reliev summer-heat, smooth the liver, dispel wind, improv eyesight, help produce saliva, treat sunstroke and high fever, headache, toothache, red eye, aphtha, polydipsia, diarrhea, dysentery, mastitis, relieve swelling and pain, astringency and hemostasis		
<i>Ligustrum robustum</i> (<i>L. purpurascens</i> , small-leaved Kudingcha)	Southwest China	Since ancient times	Tea drink and health care products in China and southeast Asia	Slightly bitter taste, sweet, slightly cold, heat clearing detoxifying, stress-reducing, relieve swelling and pair anti rheumatism and scald, furuncle and arthralgia therapy		
Litsea coreana (eagle tea)	Guizhou, Sichuan Of China	Since ancient times	Southwest China	Detoxify, detumescence, improve eyesight, help produce saliva, decrease thirst, prevent sunstroke		

 Table 2
 Polyphenol constituents of green tea, mate tea and large-leaved Kudingcha

Name	Total polyphenol	Caffeine	Total flavonoid(%)	Total saponin
Green tea	30%	1-4%	>0.4	NR
Mate tea	More abundant than green tea 178.32 mg/g spray-dried extract	0.7–2%	>1	0.35 g/L
Large-leaved Kudingcha	2-6%	No or trace (0.228-0.436 µg/g)	9–12	>20%
Small-leaved Kudingcha	20–30%	NR	>5	-
Eagle tea	> 30%	NR	>10	>1%

Modified from Ref. 2. NR: not reported.

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