## ISO/DTR 18986

ISO/TC 249<del>/WG 1</del>

Secretariat: SA

Date: 2023-<del>08-07</del>10-26

Traditional Chinese medicine——Report on <u>Panax ginseng</u> in <u>terms of the global</u> industry and standardization development <u>of Panax ginseng</u>

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This document was prepared by Technical Committee ISO/TC 249, Traditional Chinese medicine. - b23e-ca0acfe | 86ef/iso-dtr-18986

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#### Introduction

Panax ginseng is the dried root and rhizome of Panax ginseng C. A. Meyer. It has the effects of greatly tonifying the original qi, restoring the pulse and relieving collapse, invigorating the spleen and benefiting lungthe lungs. It is used primarily for symptoms such as weakness, cold extremities with faint pulse and spleen deficiency with reduced appetite. [2[22]]. As a kind of rare and valuable Chinese materia medica, Panax ginseng has a long history forof medicinal use. Its single and compound formulas, as well as a variety of modern preparations, have been playingplay an important role in clinical practice. What's moreFurthermore, Panax ginseng is widely applied in many other fields such as food, health carehealthcare and daily chemical products.

As one of the Chinese materia medica with the largest global trade volume, *Panax ginseng* occupies a large market scale. It is mainly cultivated in China, <u>South-the Republic of</u> Korea, <u>North-the Democratic People's Republic of</u> Korea, Japan and <u>Russiathe Russian Federation</u>, among which China and <u>South-the Republic of</u> Korea are the main countries for its production and consumption, accounting for over 80 % of the total yield in the world. In addition, *Panax ginseng* is also widely used in Europe and the United States, as well as some other countries in Asia.

Panax ginseng has been recorded in many national pharmacopoeias and regional standards. However, due to the restrictions on different national regulations and different purposes for standard development, the requirements specified by such pharmacopoeias and regional standards differ in thetheir scope, items, indicators, and other aspects. The lack of unified quality and safety standards gives rise to some negative phenomena, such as unqualified and fake products available on the market, substandard products sold at high prices and confusion with specification and grades of Panax ginseng products. Besides In addition, the lack of unified standards in terms of the terminology and processing technology leads to the confusion with products' names and brings lots of difficulties not only to supervision authorities but also to consumers. Therefore, it is urgent to develop unified international standards International Standards for the whole industry chain of Panax ginseng, including the production, processing, marketing and other links.

Standards for the whole industry chain of *Panax ginseng* would guarantee the sustainable development of the industry and international trade in the aspects of promoting the manufacturing, improving the quality, regulating the market and protecting the consumers' interests. This document conducts a comprehensive survey on the industry of *Panax ginseng* in terms of its origin and application history, medicinal value, geographical distribution, cultivation methods, processing methods, the development of industry, international trade and standardization status. It will be beneficial to provide reference and information support for the follow-up formulation of the <u>international standards</u> for *Panax ginseng*.

Traditional Chinese medicine———Report on Panax

ginseng in terms of the global industry and standardization development of Panax ginseng

## 1 Scope

This document <u>reviews</u>reports on the global industry and standardization development of *Panax ginseng*. It includes its origin and application history, medicinal value, geographical distribution, cultivation and processing methods, industry development, international trade, current status and development demands for standardization.

## 2 Normative references

There are no normative references in this document.

## 3 Terms and definitions

For the purposes of this document, the following terms and definitions apply.

ISO and IEC maintain terminological terminology databases for use in standardization at the following addresses:

- ——ISO Online browsing platform: available <a href="https://www.iso.org/obphttps://www.is
- IEC Electropedia: available at <a href="https://www.electropedia.org/">https://www.electropedia.org/</a>

## 3.1

## fresh ginseng

ginseng root harvested from the field, before being washed

[SOURCE: ISO 19610:2017, 3.1]

## 3.2

## washed ginseng

raw ginseng root washed with drinking water to remove any foreign matter and which is then used as the starting material for manufacturing red ginseng

[SOURCE: ISO 19610:2017, 3.2]

## 3.3

## steamed ginseng

ginseng produced through a process of steaming the washed ginseng to gelatinize the starch content

[SOURCE: ISO 19610:2017, 3.3]

#### 3.4

## red ginseng

ginseng root from *Panax ginseng C.A.* Meyer, treated with steam and then dried and packaged as whole or cut roots

[SOURCE: ISO 19610:2017, 3.4]

3.5

origin

country of origin and province/state of the seed.

The Standards

[SOURCE: ISO 17217-1:2014(E) 9.D]

3.6

Document Preview

place at which ginseng was cultivated

[SOURCE: ISO 19610:2017, 3.9]

place of cultivation

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## 4 Origin and application history

## 4.1 Origin

Panax ginseng is one of the oldest relict plants on earth. The character "学" was first discovered in the oracle bone inscriptions of the Shang Dynasty of China, dating back to 3 500 years—ago. The character "" on the oracle bone was pictographic, showing the typical characteristics of the aboveground and underground parts of Panax ginseng. [444] Fan Zi Ji Ran [545] (《范子計然》), written in the Warring States Period (465 BC), clearly recorded the origin and properties of Panax ginseng, which is the earliest available document of Panax ginseng.

## 4.2 Application history in China

## 4.2.1 Qin and Han dynasties

The Qin and Han dynasties were the initial medicinal and clinical application stages of *Panax ginseng*. The medicinal value of *Panax ginseng* was well<u>As</u> recorded in *Sheng Nong's* 

Herbal Classics [6] (《神農本草經》). Panax ginseng has rich therapeutic effects, such as follows: "Panax ginseng, sweet in taste and cold in property, is mainly used to tonifycalming the five Zang-organs, to calm essence and spirit, to tranquilize the ethered soul and corporeal soul, to cease fright and palpitation, to improve improving vision, to soothe the heart and to promote wisdom. Long term taking of it will strengthen the body and prolongenhancing intelligence and prolonging life. It is either called Renxian or Guigai. It grows in mountains and valleys".

Medicine Inscribed Wooden Slips of the Han Dynasty in Wuwei<sup>[2][4]]</sup> (《武威漢代醫簡》), the earliest literature on the clinical application of Panax ginseng, summarized the medication rules of Panax ginseng in treating various diseases. In the Eastern Han Dynasty, the clinical application of Panax ginseng was expanded and it was widely used in various formulas. In the Treatise on Febrile Diseases [2][4][(《海寒論》), there were 23 of a total of 113 formulas containing Contained Panax ginseng.

## 4.2.2 Jin and Southern-Northern dynasties

The clinical application of *Panax ginseng* was further explored in the Jin and Southern-Northern dynasties. The single prescription of *Panax ginseng* ""Ginseng Powder" was first recorded in the Handbook of Prescription for Emergency (《 所後備急方》) in the Eastern Jin Dynasty. It is mainly used for the treatment of sudden reversed flow of qi and life-threatening diseases. *Annotation of Materia Medica* (《本草經集注》), written induring the Liang Dynasty, elaborated the therapeutic effects of *Panax ginseng* on diseases occurring in specific Zang-fu organs, which extended the scope of functions and indications of *Panax ginseng*. The cultivation of *Panax ginseng* in gardengardens was first recorded in the *Individual Biography of Shi Le from The Book of Jin* (《 音書 石勒別/傳》).

## 4.2.3 Tang and Song dynasties

The Tang and Song dynasties witnessed the rapid development of application and dissemination of *Panax ginseng. Tang Materia Medica*[12[12]] (《新修本章》), the first official pharmacopoeia of China, recorded in detail about the functions, indications, usage, properties, and main production areas of *Panax ginseng*. Besides In addition, a large number of prescriptions containing *Panax ginseng* is also included in *Prescriptions Worth Thousand Gold for Emergencies*[13[13]] (《備意子金要方》) and Supplement to Invaluable Prescriptions for Ready Reference [14[14]] (《行金翼方》), with 445 and 310 prescriptions, respectively. With the frequent foreign trade and exchange activities, *Panax ginseng has beenspread* more widely spread to abroadoverseas in the Tang Dynasty. *Panax ginseng produced* in the Bohai State (an ancient kingdom existed existing from 698 AD to 926 AD) was continuously paid to the Central Plainscentral plains as tributes. Master Jianzhen, an eminent Chinese monk in Tang Dynasty, propagated the herb of *Panax ginseng* and its application to Japan. This herb is still preserved in Shosoin of Japan today. 115[15] 1.

Traditional Chinese <u>Medicine medicine</u> writings flourished in the Song Dynasty. *Illustrated Classics of Materia Medica*[16]16]1 (《本草圖經》) first systematically described the

morphological characteristics of the original plant of  $Panax\ ginseng$  in the form of illustrations.  $Classified\ Materia\ Medica^{[12[+7]]}\ (《證類本草》)$  further explained the medicinal theory of  $Panax\ ginseng$ . In the Song Dynasty, the resources of  $Panax\ ginseng$  increased with the eastward expansion of its main production areas. Meanwhile, a quantity of  $Panax\ ginseng$  was imported through border trade to ensure domestic demand for medicinal use  $\frac{115[15]}{12}$ 

#### 4.2.4 Ming and Qing dynasties

The Ming Dynasty iswas the peak period of clinical theory and practical application of Panax ginseng. Biography of Panax ginseng (《人多傳》), the first monograph of Panax ginseng, comprehensively summarized its pharmacology and medicinal properties. On this basis, Compendium of Materia Medica[18[18]] (《本草綱目》) made a detailed collation of the prescriptions and indications of Panax ginseng, including 67 prescriptions applied to 15 kinds of diseases. However, due to the severe destruction of resources, the production areas of Panax ginseng moved northward. Thus, the supply of Panax ginseng in the Ming Dynasty fell into a difficult period. Under such circumstances, the government opened cross-border markets to import large quantities of Panax ginseng from the Liao Dynasty and the Joseon Dynasty of Korea in order to ease domestic demand for supply. At the same time, large-scale artificial cultivation of Panax ginseng has been was realized through seed propagation among the people.

In the Qing Dynasty, the theories and management of *Panax ginseng* were well developed. *Essential of Materia Medica*[20[20]] (《本章備要》) divided *Panax ginseng* into "raw" and "cooked" ginseng. *Ginseng spectrum*[21[24]] (《多譜》) discussed ginseng production, purchase and sales, identification of quality, processing methods and value in detail. *BesidesAt the same time*, the authorities tightened the control over *Panax ginseng* by establishing monopoly institutions and implementing various policies and systems for the management of *Panax ginseng*. In addition, the grades and specifications of *Panax ginseng* were clearly specified. In the late Qing Dynasty, the garden cultivation of *Panax ginseng* was rapidly developed to make up for the shortage of its natural resources.

## 4.2.5 Modern times

Panax ginseng has been greatly developed covering such aspects as modern basic research, industrial scale and regulation supervision. With the development of modern medicine, biology and chemistry, people have a growing understanding of the pharmacological effects of Panax ginseng. Meanwhile, remarkable achievements have been made in the research of the material basis of Panax ginseng. Supported by—the modern science and technology, the industry of Panax ginseng continues to expand. The manufactured products of Panax ginseng, such as general food, health functional food, cosmetics and drugs have become the main components of the international industry of Panax ginseng. In order to promote the rapid and healthy development of this industry, relevant regulations and standards have been successively issued by various countries.

Panax ginseng was one of the first Chinese materia medica included in Chinese Pharmacopoeia 1963, with a detailed description of its source, processing, usage and dosage. The industry development of Panax ginseng in the People's Republic of China can be roughly divided into four periods:

- Expansion period (1970-to 1990): A production model of high quality and high yield was achieved in terms of the planting of *Panax ginseng*. [23 [23].]
- Turbulent period (1990-<u>to</u> 2002): The conflict between the lands and forests on *Panax ginseng* cultivation led to a drop in price and a slowdown in the industry.
- Recovery period (2003-<u>to\_</u>2012): The government paid more attention to the industry of *Panax ginseng* and facilitated with <del>the</del>-modern technology to promote its industry development.
- Growth period (2012-to present): The industry of *Panax ginseng* developed rapidly after its approval as ""new resource food"—".

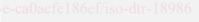
## 4.3 Application history in Korea

#### 4.3.1 The Samhan Period

According to <u>literature, [24,25]</u>. Iteratures [24,25]. Panax ginseng has already been cultivated and used to make medicine since the Samhan Period. It is recorded that Panax ginseng <u>grewgrown</u> on the Korean peninsula was traded to <u>the</u> Tang Dynasty, China, during the periods of Kings Jinpyeong (A.D. 627) and Sungdeok (A.D. 723) of the Silla Dynasty.

## 4.3.2 Goryeo Dynasty (Goryeo Period)

Panax ginseng was actively traded not only with China but also with Japan. In the reigning period of King Wu, Goryeo delivered ginseng as a gift when sending envoys to Japan to effectively handle trouble caused by Japanese pirates. Panax ginseng, which was used as medicine and tributes during the Samhan period, was regarded as a commodity that royal families trade for profit and was subject to taxation in the Goryeo Dynasty. In particular, it was one of the nation's representative items considered as significant in terms of diplomatic relations with China and Japan.



#### 4.3.3 Joseon Dynasty

The clinical application and processing &and cultivation method of *Panax ginseng* were greatly developed in the Joseon Dynasty. *Principles and Practice of Eastern Medicine*<sup>[27[27]]</sup> published during the reign of King Seonjo contains more than 170 prescriptions based on *Panax ginseng*. Moreover, Steamedsteamed ginseng in the Goryeo Dynasty was converted into pasam (steamed, compressed, and threaded ginseng) and pansam (steamed, compressed, and pasted ginseng) that are the ancestors of red ginseng. During the periods of Kings Sukjong and Jeongjo, the red ginseng production method was fully developed to enable the stable and long-distance transport.

In the Joseon Dynasty, trading ginseng privately was thoroughly controlled, so that ginseng was used as medicine or tributes by the royal families or those in authority for profit, [28][28]. However, the prices of *Panax ginseng* were sharply raised due to lack of wild ginseng, which requires the development of ginseng cultivation methods. *Imwongyeongjeji* shows that ginseng cultivation methods were fully established in the mid-Joseon Dynasty. [29][29][1]. Against this backdrop, the development of *Panax ginseng* and red ginseng cultivation methods greatly contributed to the financial health of the nation and the royal families.

#### 4.3.4 Modern times

Throughout the history of Korea, *Panax ginseng* has been traded as one of Korea's major commodities. [30[240]-1] In 1899, the Ginseng Policy Team (Samjeonggwa) initiated producing red ginseng under the leadership of the central government. In 1908, the *Red Ginseng Monopoly Act* and the *Ginseng Tax Act* were promulgated to lead the government to monopolize the red ginseng market. The *Ginseng Policy Handbook (Samjeongyoram)* [30[30] 1 published by the Ginseng Policy Team shows that research on ginseng cultivation methods, pest control, red ginseng manufacturing processes, and the age of *Panax ginseng* optimized for red ginseng manufacturing was systematically conducted.

In 1996, the *Red Ginseng Monopoly Act* and the *Ginseng Industry Act* were abolished and established, respectively, thereby transferring the authority to control *Panax ginseng* and red ginseng to the private sector for the first time in 600 years. The *Ginseng Industry Act* specifies matters required for the cultivation, manufacturing, and testing of *Panax ginseng* and ginseng products, leading *Panax ginseng* to be protected and promoted as a specialty-speciality and significantly increasing the number of related firms and red ginseng products. [30[30]]] After diverse animal and clinical tests, the functionality of *Panax ginseng* has been officially recognized in several areas by the Ministry of Food and Drug Safety. [31[31]]

## 4.4 Application history in other countries

Panax ginseng is mainly used for Kampo medicines and health food in Japan. In ancient Japan, Panax ginseng was frequently used by Kampo practitioners. [32[32]-] With the prosperity of Kampo medicine, prescriptions containing Panax ginseng were more widely

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used in the treatment of several diseases. Japan attaches great importance to the quality standards of Kampo medicine preparations and *Panax ginseng* varieties with high extract content and short leaching time are often used as raw materials.

Wild *Panax ginseng* is listed in the *Red Book of Primorsky Krai* as an endangered species in <u>Russia the Russian Federation</u>. Cultivated *Panax ginseng* has been planted in Siberia since the 19th century. In 1968, it was included in the State Pharmacopoeia of the Soviet Union. *Panax ginseng* is popular in <u>Russia the Russian Federation</u> as medicine, health products and food. [34]34].

#### 5 Medicinal value

#### 5.1 Traditional use

Panax ginseng has the effects of greatly tonifying the original qi, restoring the pulse and relieving collapse, invigorating spleen and benefiting lung, engendering fluid and nourishing blood, and quieting the spirit and sharpening the wits. It is used primarily for symptoms such as weakness, cold extremities with faint pulse and spleen deficiency with reduced appetite.

Both single and formulas of *Panax ginseng* can be used in traditional clinical application. Nevertheless, there are some incompatibilities. For example, *Panax ginseng* is <a href="mailto:antagonized">antagonized</a> by *Veratrum nigrum* root and rhizome, restrained by *Trogopterori* faeces and inhibited by *Raphanus sativus* seed. [18[18].]

#### 5.2 Modern application

## 5.2.1 Phytochemistry and pharmacological effects

Panax ginseng is mainly composed of saponins, polysaccharides, volatile oils, organic acids and esters, sterols, flavonoids, lignans, inorganic elements and vitamins. [35[45],] A variety of pharmacological actions also have been proven in Panax ginseng, including antifatigue, anti-stress, anti-mutagenesis, anti-oxidation, anti-cancer, as well as protection of the cardiovascular system. [36, etc[36],]

Ginsenosides are well believed as the primary bio-active ingredients in *Panax ginseng*. More than 170 kinds of ginsenosides have been reported [37[37]] which can be mainly divided into protopanaxadiol type (PPD), protopanaxatriol type (PPT), and oleanolic acid type (OA) according to their structure properties. Ginsenosides of different types possess varied of pharmacological characteristics. For example, PPT is recognized for the effects of stimulating central nervous system, improving memory, nourishing and promoting protein synthesis, whereas OA exerts the effects of anti-inflammation and anti-platelet release.

Besides, polysaccharides and volatile oils are also important ingredients of *Panax ginseng*. Up to now, there have been reports of 64 polysaccharides [38-[39]] and 160 volatile oils from *Panax ginseng*, [39[39]].] Ginseng polysaccharides are mainly composed of neutral and pectin-

like acidic polysaccharides, which play critical roles in enhancing immunity, anti-aging, anti-tumortumour and reducing liver injury. Ginseng volatile oils with special fragrance are primarily made up of sesquiterpenes, long-chain saturated acids and aromatic hydrocarbons. Among them, sesquiterpenes are the mostly constituents, while polyacetylenols such as panaxynol and panaxyloxynol are regarded as the principalcharacteristic compounds. [40[40]-]. Ginseng volatile oil has significant effects on antifungal, anti-inflammatory, central nervous system depression and bi-directional regulation of blood pressure.

In addition, the traditional processing methods of *Panax ginseng* could alter its chemical composition and pharmacological effects to achieve the purpose of efficiency enhancing. <u>Till now, various Various</u> processed *Panax ginseng* products have been exploited to better the therapeutic demands (see <u>Clause 8See clause 8 for more detailed information</u>).].

## 5.2.2 Modern TCM traditional Chinese medicine preparations

Ginsenoside monomer compounds, extract and preparations of *Panax ginseng* have been widely developed into drugs for the treatment of different diseases. Ginsenoside Rg3, 20 (S) -ginsenoside Rg3 and Ginsenoside Rd[41.42[41-42]] have been developed into new anticancer and antiviral drugs. Capsules developed from the fruits and total saponins of *Panax ginseng* leaves have been used as adjuvant medicine for the treatment of coronary heart disease, climacteric syndrome, diabetes, and <a href="mailto:tumours.[43.44]tumours.[43.44]">tumours.[43.44]</a> I Ginseng polysaccharide injection has been also mainly used for fighting various chronic infections, diabetes and various immune diseases.[45-[45]]

## 5.3 Healthcare and functional products UMENT Preview

Except for medicinal value, *Panax ginseng* has been widely developed into healthcare and functional products because of its rich nutrients. These products are mainly helpful for immunity enhancement, anti-aging, blood sugar control and auxiliary protective effects against liver injury, sleep improvement and anti-oxidation. [46[46]]

Functional foods with *Panax ginseng* include capsules, sticks, ampoules, pills, drinks, baked food, prepared food, and condiments and so on.—For example, ginseng polysaccharide fermented milk beverage [47,447] is used for improving immunity, preventing and treating hypertension, and fermented ginseng rice wine is used for antioxidation, with rare saponin CK produced and the content of ginsenoside increased significantly after fermentation.[48,449],]

The use of natural and non-toxic botanical sources has become the trend of cosmetics industrial development. Ginseng extracts and ginsenoside have been used in the field of cosmetics since the 1980s and nowadays have become the most widely used starting materials in plant cosmetics. [49[49],1] For example, moisturizing cream, whitening cream, facial mask, eye cream and shampoo made with *Panax ginseng* are mainly used to improve the antioxidant activity of skin cells, inhibit melanin transfer, improve skin colorcolour and reduce hair loss. [50.51[50.51],1]

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