

Review Article

J. Ginseng Res. Vol. 35, No. 4, 389-398 (2011)
<http://dx.doi.org/10.5142/jgr.2011.35.4.389>

Trends in Ginseng Research in 2010

Si-Kwan Kim^{1*} and Jeong Hill Park²

¹Department of Life Sciences, College of Biomedical and Health Sciences, Konkuk University, Chungju 380-701, Korea

²College of Pharmacy, Seoul National University, Seoul 151-742, Korea

A total of 470 papers directly related to research on the *Panax* species were retrieved by performing internet searches with the keywords *Panax* and ginseng as the search terms. The publications were categorized as follows: 399 research articles, 30 reviews, 30 meeting abstracts, 7 proceedings, and 4 letters. The majority of these publications were published by scientists from Korea (35.7%), China (32.3%), and the USA (11.3%). Scientists from a total of 29 nations were actively involved in conducting ginseng research. A total of 43.6% of the publications were categorized as pharmacodynamic studies. The effects of ginseng on cerebrovascular function and cancer were the two most common topics considered in the pharmacodynamic studies. More than half of the ginseng studies assessed the use of *P. ginseng*. A total of 23 countries participated in studies specifically related to *P. ginseng*, and more than 80% of these studies originated from Korea and China. A total of 50 topics within the pharmacodynamics category were examined in association with the use of *P. ginseng*.

Key words: Trends in ginseng research, 2010, Country of origin of research publications, Major research categories, Pharmacodynamics

INTRODUCTION

This review provides a comprehensive overview of the trends in ginseng research in 2010. A total of 588 scholarly works were retrieved when an internet search using the keywords *Panax* and ginseng as the search terms was performed. Brazilian (*Pfaffia glomerata* and *P. paniculata*), Indian (*Withania somnifera* L Dunal Solanaceae) and Siberian ginseng (*Eleutherococcus senticosus*) were excluded from this analysis even though the term 'ginseng' was used in their descriptions. Publications that had no direct relationship to ginseng research were also excluded. For example, articles on the identification of microbes isolated from fields used for field-cultivated ginseng were excluded from this study. However, an article that examines the microbially mediated bioconversion or degradation of ginsenosides by microorganisms isolated

from a ginseng cultivation field was included. The country from which the research and publication originated was categorized based on the country of residence of the corresponding author. Research categories were defined based on the major research topics examined.

PUBLICATION CATEGORIES

A total of 470 publications retrieved by the internet search were reviewed and classified, according to their country of origin and main research topics (Table 1). There were 399 formal research articles, which accounted for 84.9% of these 470 publications. There were 30 review articles and 30 meeting abstracts, which each comprised 6.4% of the publications. The 4 letters and 7

© This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (<http://creativecommons.org/licenses/by-nc/3.0/>) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

Received 01 Nov. 2011, Revised 22 Nov. 2011, Accepted 22 Nov. 2011

*Corresponding author

E-mail: skim@kku.ac.kr

Tel: +82-43-840-3574, Fax: +82-43-840-3929

Table 1. Types of publications

Publication type	No. of papers	Percentage
Research articles	399	84.9
Review papers	30	6.4
Meeting abstracts	30	6.4
Letters	4	0.9
Proceedings papers	7	1.5
Total	470	100

proceedings papers together accounted for slightly less than 2.5% of all of the publications.

RESEARCH PUBLICATIONS

A total of 29 nations were actively involved in conducting ginseng research (Table 2). Korea ranked first in terms of the number of publications, contributing 168 papers (35.7%), and China ranked second with 152 publications (32.3%).

In the second tier, there were 53 papers that originated from the United States of America (USA), 16 papers from Japan, and 13 publications each from Taiwan and Canada. In the third tier, India, Russia, Hong Kong,

Brazil, and Singapore published 9, 7, 4, 4, and 3 papers, respectively. Several other countries contributed 1 or 2 papers. Interestingly, the research output by Japanese scientists decreased dramatically in 2010 compared to previous years. On the other hand, several countries in the Middle East (Egypt and Iran), Europe (Turkey, Greece, Poland, and Slovakia) and South America (Uruguay) appear to have recently initiated ginseng research programs. A small number of scientists from Western Europe, including the United Kingdom, Italy, Switzerland, Austria, and Germany, showed some interest in ginseng research in 2010 as indicated by the research publication volume; the Scandinavian countries exhibited the least amount of interest.

MAJOR RESEARCH CATEGORIES AND THE CRITERIA FOR CATEGORIZATION

All of the 470 papers were sorted into research categories based on the following specific criteria (Table 3): publications detailing cultivation methods, tissue culture techniques, or plant pathology as the major research topic were placed into the ‘horticulture’ category; processing, bioconversion, extraction methods, and quality control

Table 2. Sorting of publications by country of origin

Origin of paper	No. of papers	Percentage
Korea	168	35.7
China	152	32.3
USA	52	11.3
Japan	16	3.4
Taiwan, Canada ¹⁾	13	2.8
India	9	1.9
Russia	7	1.5
Hong Kong, Brazil, United Kingdom ¹⁾	4	0.9
Singapore, Australia ¹⁾	3	0.6
Italy, Turkey, Austria, Pakistan, Egypt, Switzerland ¹⁾	2	0.4
Uruguay, Slovakia, Greece, Germany, Poland, Iran, Vietnam, Thailand, Denmark, New Zealand ¹⁾	1	0.2
Total	470	100

¹⁾The number of publications is the same for each country within the group.

Table 3. Research categories

Category	Area of research
Horticulture	Cultivation, tissue culture, plant pathology
Manufacture	Process, bioconversion, extraction methods, quality control
Analytical chemistry	Analysis of pesticide residues and ginsenosides
Pharmacodynamics	<i>In vitro</i> and <i>in vivo</i> biochemical and physiological studies; prevention of wrinkle formation and sunlight damage
Pharmacokinetics	<i>In vivo</i> metabolism of active compounds
Clinics	Clinical and epidemiological studies, pharmacovigilance
Marketing surveillance	Market share
Others	Nutrition, cultivation of mushroom, genetics, identification of ginseng species, quality control, gene expression

studies were categorized as ‘manufacture’ publications; analyses of pesticide residues and ginsenosides were categorized as ‘analytical chemistry’ research; *in vitro* and *in vivo* biochemical, pharmacological, and physiological studies and studies examining ginseng use for the prevention of wrinkle formation and sunlight damage were categorized as ‘pharmacodynamics’ publications; studies of the *in vivo* metabolism of active compounds were categorized as ‘pharmacokinetics’ research; clinical and epidemiological studies and pharmacovigilance studies were categorized as ‘clinics’ publications; and market share studies were categorized as ‘marketing surveillance’ research.

MAJOR RESEARCH CATEGORIES

A total of 205 papers related to pharmacodynamics research were published in 2010, accounting for 43.6% of the 470 ginseng-focused publications (Table 4). The second most abundant category was analytical chemistry, with a total of 72 publications (15.3% of all of the

Table 4. Sorting of papers by research categories

Category	No. of papers	Percentage
Horticulture	49	10.4
Manufacture	41	8.7
Analytical chemistry	72	15.3
Pharmacodynamics	205	43.6
Pharmacokinetics	33	7.0
Clinics	31	6.6
Marketing surveillance	4	0.9
Others	35	7.4
Total	470	100

Table 5. Subcategories of pharmacodynamic studies

Subcategory	Specification
Brain function (19)	Brain cell (6), brain function (5), nerve cell (3), depression (2), cognition improvement (2), Alzheimer (1)
Cerebrovascular function (39)	Hypertension (17), atherosclerosis (4), stenocardia (4), cerebral ischemic (3), blood vessel (3), cardiovascular (2), cardiac ischemia (2), blood circulation (1), hyperlipidemia (1), arrhythmia (1), improvement of lipid metabolism (1)
Antitumor effect (39)	Anticancer (37), stomach cancer (2)
Immune function and anti-inflammatory effect (26)	Anti-inflammatory (12), immune function (8), dermatitis (2), antiviral (1), peptic ulcer (1), allergy (1), AIDS (1)
Bone (7)	Arthritis (2), bone necrosis (2), spinal cord damage (1), osteoporosis (1), fracture (1)
Antistress, antifatigue and tonic effect (9)	Fatigue (4), stress (4), tonic effect (1)
Antioxidant and antiaging effect (22)	Antioxidant (13), oxidative stress (3), radiation protection (2), skin ageing (2), cosmetic (1), antiageing (1)
Liver and toxicity (12)	Liver toxicity (6), liver damage (5), toxicity (1)
Diabetes and obesity (24)	Diabetes (21), obesity (3)
Sexuality (3)	Male sexual function (1), climacteric disturbance (2)
Others (5)	Insomnia (2), eye and vision (1), hyperactivity (1), gene expression (1)

publications). There were 49 and 41 articles that were regarded as cultivation research and manufacture studies, respectively, accounting for 10.4% and 8.7% of all of the publications. Pharmacokinetic and clinical studies comprised 33 (7.0%) and 31 (6.6%) of the studies, respectively.

PHARMACODYNAMIC SUBCATEGORIES

The pharmacodynamics category was further subdivided into several specific research areas (Table 5). Ginseng research related to cancer prevention and therapy was the most common topic, accounting for 39 of the 205 pharmacodynamic studies. The majority of the cancer research was confined to *in vitro* cellular studies.

PANAX GINSENG AND RELATED SPECIES

There were 268 studies, representing 57% of the total publications in 2010, that focused on *Panax ginseng* as the primary research topic (Table 6). There were smaller numbers of articles that primarily examined *P. quinquefolium* (49), *P. notoginseng* (36), and *P. japonicus* (5), while a total of 112 studies were performed using ginsenosides of obscure origins. Studies on *P. japonicus* were rare, and the number of publications devoted to *P. ginseng* was more than five fold greater than the number of studies that focused on *P. quinquefolium*. Many of the papers classified as ‘others’ were mechanistic studies conducted using a single ginsenoside.

PANAX GINSENG

Researchers from a total of 23 countries performed

Table 6. Sorting of studies of *Panax ginseng* and other related species

Type of ginseng	No. of papers	Percentage
<i>P. ginseng</i>	268	57.0
<i>P. quinquefolium</i>	49	10.4
<i>P. notoginseng</i>	36	7.7
<i>P. japonicus</i>	5	1.1
Others	112	23.8
Total	470	100

studies focused on *P. ginseng*, but more than 80% of the studies originated from Korea and China (Table 7). Of the 268 papers published on *P. ginseng*, Korean and Chinese scientists published 154 (57.5%) and 68 papers (25.4%), respectively. Scientists from the USA and Japan each published 9 papers (3.4%). The numbers of publications by Japanese scientists decreased compared to prior years. In addition to the three main East Asian countries and the USA, scientists from other regions of Asia, the Americas, Europe, and the Middle East also conducted research on *P. ginseng*. Although scientists in New Zealand, Canada, India, Austria, Turkey, Hong Kong, Singapore, Pakistan, Denmark, Switzerland, Poland, Germany, Greece, Australia, and Uruguay only published one paper from each country, it appears that the interest in supporting *P. ginseng* research is increasing in these countries.

A broad spectrum of pharmacodynamic studies were performed with *P. ginseng* as the main focus. There were at least 50 distinct topics studied that related to *P. ginseng*, including the following: its roles in the detoxification of anticancer agents, immune function, antioxidant activity, allergies, obesity, blood vessels, peptic ulcers, dermatitis, cytotoxicity, brain neurons, hepatic disease, lipid metabolism, cancer, AIDS, stress, inflammation, sexuality, diabetes, fatigue, neurotoxicity, viral infections, cosmetics, tumors, oxidative stress, apoptosis, granulocyte growth, platelet aggregation, hyperactivity, insomnia, cognitive functions, enzyme activity, skin aging, car-

Table 7. Country of origin of *Panax ginseng* studies

Host country	No. of papers	Total
Korea	154 (57.5%)	154
China	68 (25.4%)	68
USA, Japan	9 (3.4%)	18
Russia	5 (1.9%)	5
Taiwan	4 (1.5%)	4
United Kingdom, Egypt	2 (0.7%)	4
New Zealand, Canada, India, Austria, Turkey, Hong Kong, Singapore, Pakistan, Denmark, Switzerland, Poland, Germany, Greece, Australia, Uruguay	1 (0.4%)	15
Total		268

Table 8. Country of origin of *Panax quinquefolium* studies

Host country	No. of papers	Percentage
USA	23	46.9
China	15	30.6
Canada	7	14.3
Japan, Austria, Taiwan, India ¹⁾	1	2.0
Total	49	100

¹⁾The number of publications is the same for each country within the group.

diac disease, gastritis, cardiac ischemia, nephrotoxicity, aging, brain ischemia, cardiovascular disease, vision, Alzheimer's disease, bone necrosis, arthritis, metabolic diseases, hyperlipidemia, photosensitivity, diabetic memory loss, cardiovascular endothelial cells, physical activity, and radiation injuries.

PANAX QUINQUEFOLIUM

A total of 49 publications that focused on the American ginseng, *P. quinquefolium*, were published in 2010. Scientists from 7 countries were actively involved in these studies (Table 8). American scientists published the largest proportion of the articles (46.9%) on this species, and the vast majority (91.8%) of the studies on *P. quinquefolium* were performed in the USA, China, and Canada. These studies originated from only a few countries. A relatively narrow spectrum of pharmacodynamic topics related to *P. quinquefolium* was examined, comprising the following 7 subjects areas: cardiovascular disease, cancer, antioxidant activity, radiation injury, diabetes, inflammation, and depression.

PANAX NOTOGINSENG

Chinese scientists performed almost all of the research concerned with *P. notoginseng*. Two groups from Korea and one group from Singapore also studied the effects of

Table 9. Country of origin of *Panax notoginseng* studies

Host country	No. of papers	Percentage
China	33	91.7
Korea	2	5.6
Singapore	1	2.8
Total	36	100

P. notoginseng (Table 9). The *P. notoginseng* subspecies is not a common research subject outside of China. Although there were fewer pharmacodynamic studies that featured *P. notoginseng* compared to *P. ginseng*, a relatively broad spectrum of research topics was addressed in studies concerned with *P. notoginseng* compared to *P. quinquefolium*. A total of 15 topics within the pharmacodynamic research category were explored in the studies of *P. notoginseng*, including arteriosclerosis, arthritis, the cardiovascular system, bone fractures, antioxidant activity, inflammation, obesity, cancer, hyperlipidemia, osteoporosis, fibrosis of the liver, Alzheimer's disease, cardiac ischemia, blood circulation, and cytotoxicity. In contrast, the pharmacodynamic studies of *P. japonicas* were limited to examinations of its effects on liver toxicity and α -glucosidase inhibition.

SUMMARY OF THE MAJOR FINDINGS OF SELECTED PUBLICATIONS

Scholey *et al.* [1] studied the effects of *P. quinquefolium* on neurocognitive function using a double-blind, placebo-controlled method and found a significant improvement in the performance of the working memory, reaction time and decision making processes, as well as increased calmness. Wang *et al.* [2] suggested that the administration of *P. ginseng* can significantly reverse learning impairments induced by scopolamine. Hao *et al.* studied the hepatic cytochrome P450-catalyzed metabolism of ginsenosides [3]. Cui *et al.* [4] performed antibody array experiments on precancerous colon cells and found that the consumption of ginseng maintains the environment of the colon in metabolic equilibrium. Cheng [5] reported that the ginsenoside Rh₂ aids in preventing and treating diabetic disorders. Park *et al.* [6] found that Korean red ginseng is clinically effective for treating dry mouth. Chung *et al.* [7] reported that Korean red ginseng improves vascular stiffness in patients with coronary artery disease. Musende *et al.* [8] attempted to treat prostate cancer patients with a combination of the ginsenoside Rh₂ and docetaxel, a chemotherapeutic agent. Sakamoto *et al.* [9] successfully expressed a single-chain

variable fragment antibody against the ginsenoside Re in silkworm larvae to establish an enzyme-linked immunosorbent assay to measure total ginsenosides for quality control purposes. Reay *et al.* [10] presented data that indicate that *P. ginseng* can improve certain aspects of the working memory and induce calmness in healthy young adults. Kim *et al.* [11] suggested that the anti-stress effects of *P. ginseng* are mediated via the down-regulation of the expression of the tyrosine hydroxylase and dopamine β -hydroxylase genes in an animal model of repeated immobilization stress. Li *et al.* [12] analyzed decocting-induced chemical transformations based on a chemical profiling method using UPLC-Q-TOFMS/MS techniques. Using an *in vitro* test, Qi *et al.* [13] attempted to describe the cytotoxic activity of *P. quinquefolium* after it had been processed into red ginseng. Zhao *et al.* [14] established and validated a quantitative analytical procedure for detecting the levels of the ginsenoside Rg₃ in plasma and urine that can be used in comparative pharmacokinetic studies. Yan *et al.* [15] elucidated the biotransformation pathway of the ginsenoside Rb1 to compound K in the presence of the glucosidase from *Paecilomyces Brainnii* sp. 229. Xue *et al.* [16] developed an X-ray phase contrast microscopy technique to identify wild ginseng by comparing the microstructures of ginseng roots. Sahashi *et al.* [17] analyzed the bioactive compounds in *P. ginseng* using nanoparticle-assisted laser desorption/ionization (nano-PALDI) mass spectrometry by preparing manganese oxide nanoparticles and developing a nano-PALDI MS method to analyze the ginsenosides in ginseng extracts. Li *et al.* [18] found that *P. quinquefolium* suppresses oxidative stress and oxidative stress-induced cell death in cardiomyocytes via the activation of the Nrf2 pathway in the murine heart. Baek *et al.* [19] demonstrated the antidiarrheal effect of ginseng using an *in vitro* model of rotavirus infection, which is the leading cause of severe diarrhea, and identified two pectic polysaccharides that are important contributors to this effect. Liu *et al.* [20] suggested that the administration of *P. notoginseng* saponins can attenuate atherogenesis through their anti-inflammatory action and their regulation of the blood lipid profile. A Brazilian scientist studied the sensory acceptance of a *P. ginseng* extract supplemented with nectar and determined that the optimal effect was observed when the extract was administered at a concentration of 20 mg per 100 mL of nectar [21]. Chan *et al.* [22] identified the α -glucosidase inhibitors contained in *P. japonicas*, including four polyacetyles, five phenolic compounds, one sesquiterpenoid, and one sterol glucoside. Zhao *et al.* [23] of Yunnan Ag-

ricultural University in China reviewed the biosynthetic pathways of triterpenoid saponins. Son *et al.* [24] of the College of Pharmacy at Chungnam National University in Korea determined the glycemic index of Korean red ginseng in human subjects and found that ginseng is beneficial for patients with metabolic disorders. Lei *et al.* [25] studied the allelopathic effects of ginsenosides on the *in vitro* growth and antioxidant activity of the ginseng callus and found that all of the ginsenosides examined inhibited the growth of the ginseng callus; the addition of the total ginsenosides mixture increased the enzymatic antioxidant activity of the ginseng callus. Shin *et al.* [26] studied the effect of Korean red ginseng on neonatal hypoxia-induced hyperactivity in rats and found that Korean red ginseng improved the hyperactivity phenotype without an incremental decrease in the locomotor activity in normal animals. Lee *et al.* [27] suggested that black ginseng improves the memory of mice with scopolamine-induced amnesia. Ma *et al.* [28] found that the ginsenoside Rg₁ promotes peripheral nerve regeneration in a crush-injury rat model. Ha *et al.* [29] found that the ginsenoside Rg₂ prevents UVB-induced cytotoxicity in a concentration- and time-dependent manner by increasing the activity of the DNA repair system, possibly by modulating the levels of the proteins involved in the p53 signaling pathway. Zhang *et al.* [30] proposed a new method for grading the quality of ginseng with the aid of 2D-IR correlation spectroscopy. Kim and Lee [31] suggested that pretreatment with panaxatriol ameliorates ischemia/reperfusion-induced myocardial damage and that this healing property is caused by a reduction in oxidative stress. Khalid *et al.* [32] of the Department of Anatomy at the University of Health Science in Pakistan studied the histological changes in the fetal brain of albino mice after maternal treatment with *P. ginseng*. Dai and Orsat [33] proposed a microwave-assisted extraction method to improve the efficiency of ginsenoside extraction from fresh American ginseng root. Wang *et al.* [34] presented data suggesting that administration of the ginsenoside Rb3 from *P. ginseng* protects against isoproterenol-induced myocardial injury and heart function impairment in the rat and that the mechanism of pharmacological action is related to the antioxidant activity of this Rb3. Sharma *et al.* [35] proposed that a synaptic mutation in the herbaceous perennial *P. sikkimensis* was responsible for male sterility in this species. Shen *et al.* [36] proposed that *P. notoginseng* saponins might be an alternative medicine for the prevention and treatment of postmenopausal osteoporosis. YoKang *et al.* [37] of the Institute of Natural Medicine at Toyama University in Japan provid-

ed evidence that the 20(*S*)-ginsenoside Rg₃ prevents the progression of renal damage and dysfunction in type 2 diabetic rats by inhibiting oxidative stress and the formation of advanced glycation end products. Dewir *et al.* [38] of the Department of Horticulture at Kafrelsheikh University in Egypt suggested that linolenic acid is an elicitor of ginsenoside accumulation in ginseng adventitious root cultures. Kim *et al.* [39] investigated the association between the pattern of dietary supplement use and the sociodemographic and lifestyle characteristics of Korean consumers and found that women use dietary supplements more regularly than men, but males preferred ginseng to other supplements. Jager *et al.* [40] of the Department of Medicinal Chemistry at the University of Copenhagen in Denmark found that *P. ginseng* added to dairy products can be pasteurized without significant changes in the ginsenoside patterns. Lee *et al.* [41] suggested that an American ginseng extract can protect lymphocytes obtained from healthy individuals against radiation injury. Huang *et al.* [42] proposed that the compounds K and Rg₁ can stimulate glucose uptake in 3T3-L1 cells, suggesting a potential application of ginseng for diabetic individuals because of its hypoglycemic properties. Xu *et al.* [43] investigated the effectiveness of a nutritional mixture containing a *P. ginseng* extract against aging. Their report suggested that the administration of the mixture in the early-midlife years significantly decreases age-related mitochondrial functional decline and preserves physical performance. Kim *et al.* [44] discovered the efficacy of the ginsenoside Rg₃ for the treatment of restraint-stressed animals. Li *et al.* [45] found that the ginsenoside Rb1 from *P. ginseng* relieves cerebral vasospasms and potentially protects against stroke and subarachnoid hemorrhaging and that the underlying mechanism may be partly related to the inhibition of the p53- and Bax-dependent proapoptosis pathways by Rb1. Jovanovski *et al.* [46] demonstrated that Korean red ginseng can improve arterial stiffness and that the principal pharmacological active agent might be the ginsenosides. Leonti *et al.* [47] of the Institute of Biochemistry and Molecular Medicine at the University of Bern in Switzerland studied the feasibility of reducing dermatitis by applying falcarinol (panaxynol, carotatoxin), which can be found in carrots, parsley, celery and *P. ginseng*. Chai *et al.* [48] of the Department of Surgery at Stanford University in the USA demonstrated that the ginsenoside Rb1 attenuates homocysteine-augmented guide wire injury-induced intimal hyperplasia in mice. These authors also suggested the potential of using ginseng to control homocysteine-related vascular injuries. Dickman *et al.* [49] of

the University of Wisconsin in the USA warned that the chronic intake of American ginseng can cause oxidative damage in postmenopausal women, based on an increase in oxidative damage markers (plasma malondialdehyde and urine 8-hydroxydeoxyguanosine) and erythrocyte antioxidant enzyme activities (superoxide dismutase and GSH reductase). Oh *et al.* [50] of the Department of Urology in the School of Medicine at Chunnam National University in Korea demonstrated that the administration of Korean red ginseng to menopausal women can improve sexual arousal. Chang *et al.* [51] of the College of Medicine at Taiwan National University in Taiwan suggested that the ginsenoside Rg₁ from *P. ginseng* possesses a remarkable potential for treating osteonecrosis of the femoral head. These authors further confirmed the angiogenic effect of Rg₁ by demonstrating tube formation in human umbilical vein endothelial cells. Zhang *et al.* [52] demonstrated the usefulness of ginsenosides against glycerol-induced acute renal failure and the activation of catecholaminergic neurons in the locus coeruleus of rats. Rosner [53] of the Department of Analytical Chemistry at the BAM Federal Institute of Material Research and Testing in Germany developed a method for validating the provenance of ginseng by analyzing strontium isotope ratios (Sr₈₇/SR₈₆) using ICP-MS. Yoo *et al.* [54] discovered the effectiveness of Sun-ginseng, a heat processed form of ginseng consisting primarily of the Rg₃, Rk₁ and Rg₅ ginsenosides, for the treatment of prostate cancer in an animal model by inhibiting cyclin and regulating the expression of the TNFRSF25 and ADRA2A genes. Wahid *et al.* [55] suggested that Korean red ginseng is useful for improving the optical process in the eyes of bullfrogs, which suggests the potential for the use of Korean red ginseng as a treatment for certain ophthalmic diseases in humans. Sagar [56] of the Juravinski Cancer Center in Canada studied the efficacy of Asian botanicals, including *P. ginseng*, for radiotherapy on tumor tissues. These authors showed an increase in the efficacy of the radiotherapy, which allowed for a reduction in the radiation doses applied to normal tissues. Ginseng selectivity protected normal tissues or increased tissue repair following radiation therapy. These results strongly suggest that ginseng agents are promising candidates for clinical trials. Kim *et al.* [57] suggested the usefulness of Korean red ginseng that is rich in the ginsenosides Rg₃, Rk₁, and Rg₅ for the treatment of collagen-induced arthritis. Thomson [58] examined the science-based medical evidence supporting the use of traditional Asian medicinal plants, including *P. ginseng*, and suggested that ginseng is an important medicinal plant based on the strong

support from medical efficacy trials. Ock *et al.* [59] of the College of Medicine at the Catholic University in Korea found that the most commonly used dietary supplement in Korea was *P. ginseng*. Abdel-Wahhab *et al.* [60] of the National Research Center in Egypt suggested that the consumption of *P. ginseng* extract can protect against aflatoxin B-1- and fumonisin-induced hepatic precancerous lesions in rats. Shin *et al.* [61] evaluated the toxicity of Korean red ginseng against embryonic implantation and mortality and fetal body weight gain at doses of up to 2,000 mg/kg/d, which is approximately 200 fold greater than the clinical dose recommended by the Korean Food and Drug Administration. The authors concluded that the consumption of ginseng did not cause embryo-fetal death or abnormalities. Wen *et al.* [62] attempted to degrade the residual fungicides in American ginseng by gamma irradiation. Quan *et al.* [63] tried to eliminate organochlorine pesticide residues from ginseng using a supercritical fluid extraction method. Lin *et al.* [64] developed a method for determining the ages of cultivations of *P. ginseng* with the aid of H₁ NMR fingerprint analysis. Kim *et al.* [65] of the Korea Research Institute of Standards and Science succeeded in creating a pure certified reference material for the ginsenoside Rg₁ [65]. Gravas *et al.* [66] of the School of Medicine at the University of Thessalia in Greece recommended the discontinuation of herbal medication for up to 15 days prior to surgery. Li *et al.* [67] demonstrated that the ginsenoside Rd from *P. ginseng* can prevent glutamate-induced apoptosis in rat cortical neurons. These results provide further evidence for the potential of voltage-independent Ca²⁺ channel blockers as novel neuroprotective drugs for the prevention of neuronal apoptosis and death by cerebral ischemia. Kim *et al.* [68] determined the efficacy of *P. ginseng* against the intestinal damage induced by gamma irradiation in mice.

CONCLUSION

A total of 470 papers (399 research articles, 30 reviews, 30 meeting abstracts, 7 proceedings and 4 letters) were published in 2010 by scholars from 29 nations. The majority of these publications were contributed by scientists from Korea, China, and the USA. *P. ginseng* was the predominant species examined in the pharmacodynamic studies.

REFERENCES

1. Scholey A, Ossoukhova A, Owen L, Ibarra A, Pipin-

- gas A, He K, Roller M, Stough C. Effects of American ginseng (*Panax quinquefolius*) on neurocognitive function: an acute, randomised, double-blind, placebo-controlled, crossover study. *Psychopharmacology (Berl)* 2010;212:345-356.
- Wang W, Liao QP, Quan LH, Liu CY, Chang Q, Liu XM, Liao YH. The effect of *Acorus gramineus* on the bioavailabilities and brain concentrations of ginsenosides Rg₁, Re and Rb₁ after oral administration of Kai-Xin-San preparations in rats. *J Ethnopharmacol* 2010;131:313-320.
 - Hao H, Lai L, Zheng C, Wang Q, Yu G, Zhou X, Wu L, Gong P, Wang G. Microsomal cytochrome p450-mediated metabolism of protopanaxatriol ginsenosides: metabolite profile, reaction phenotyping, and structure-metabolism relationship. *Drug Metab Dispos* 2010;38:1731-1739.
 - Cui X, Jin Y, Poudyal D, Chumanovich AA, Davis T, Windust A, Hofseth A, Wu W, Habiger J, Pena E *et al.* Mechanistic insight into the ability of American ginseng to suppress colon cancer associated with colitis. *Carcinogenesis* 2010;31:1734-1741.
 - Cheng JT. Merit of ginseng in the improvement of insulin resistance. *J Ginseng Res* 2010;34:155-159.
 - Park JW, Lee BJ, Bu Y, Yeo I, Kim J, Ryu B. Effects of Korean red ginseng on dry mouth: a randomized, double-blind, placebo-controlled trial. *J Ginseng Res* 2010;34:183-191.
 - Chung IM, Lim JW, Pyun WB, Kim H. Korean red ginseng improves vascular stiffness in patients with coronary artery disease. *J Ginseng Res* 2010;34:212-218.
 - Musende AG, Eberding A, Jia W, Ramsay E, Bally MB, Guns ET. Rh₂ or its aglycone aPPD in combination with docetaxel for treatment of prostate cancer. *Prostate* 2010;70:1437-1447.
 - Sakamoto S, Pongkitwittoon B, Nakamura S, Maenaka K, Tanaka H, Morimoto S. Efficient silkworm expression of single-chain variable fragment antibody against ginsenoside Re using *Bombyx mori* nucleopolyhedrovirus bacmid DNA system and its application in enzyme-linked immunosorbent assay for quality control of total ginsenosides. *J Biochem* 2010;148:335-340.
 - Reay JL, Scholey AB, Kennedy DO. *Panax ginseng* (G115) improves aspects of working memory performance and subjective ratings of calmness in healthy young adults. *Hum Psychopharmacol* 2010;25:462-471.
 - Kim Y, Choi EH, Doo M, Kim JY, Kim CJ, Kim CT, Kim IH. Anti-stress effects of ginseng via down-regulation of tyrosine hydroxylase (TH) and dopamine β-hydroxylase (DBH) gene expression in immobilization-stressed rats and PC12 cells. *Nutr Res Pract* 2010;4:270-275.
 - Li SL, Lai SF, Song JZ, Qiao CF, Liu X, Zhou Y, Cai H, Cai BC, Xu HX. Decocting-induced chemical transformations and global quality of Du-Shen-Tang, the decoction of ginseng evaluated by UPLC-Q-TOF-MS/MS based chemical profiling approach. *J Pharm Biomed Anal* 2010;53:946-957.
 - Qi LW, Wang CZ, Yuan CS. American ginseng: potential structure-function relationship in cancer chemoprevention. *Biochem Pharmacol* 2010;80:947-954.
 - Zhao Q, Zheng X, Jiang J, Zhou H, Hu P. Determination of ginsenoside Rg₃ in human plasma and urine by high performance liquid chromatography-tandem mass spectrometry. *J Chromatogr B Analyt Technol Biomed Life Sci* 2010;878:2266-2273.
 - Yan Q, Zhou W, Shi XL, Zhou P, Ju DW, Feng MQ. Bio-transformation pathways of ginsenoside Rb₁ to compound K by beta-glucosidases in fungus *Paecilomyces* Bainier sp. 229. *Process Biochem* 2010;45:1550-1556.
 - Xue YL, Xiao TQ, Wu LH, Chen C, Guo RY, Du GH, Xie HL, Deng B, Ren YQ, Xu HJ. Investigation of characteristic microstructures of wild ginseng by X-ray phase contrast microscopy. *Acta Phys Sinica* 2010;59:5496-5507.
 - Sahashi Y, Osaka I, Taira S. Nutrition analysis by nanoparticle-assisted laser desorption/ionisation mass spectrometry. *Food Chem* 2010;123:865-871.
 - Li J, Ichikawa T, Jin Y, Hofseth LJ, Nagarkatti P, Nagarkatti M, Windust A, Cui T. An essential role of Nrf2 in American ginseng-mediated anti-oxidative actions in cardiomyocytes. *J Ethnopharmacol* 2010;130:222-230.
 - Baek SH, Lee JG, Park SY, Bae ON, Kim DH, Park JH. Pectic polysaccharides from *Panax ginseng* as the antirotavirus principals in ginseng. *Biomacromolecules* 2010;11:2044-2052.
 - Liu Y, Zhang HG, Jia Y, Li XH. *Panax notoginseng* saponins attenuate atherogenesis accelerated by zymosan in rabbits. *Biol Pharm Bull* 2010;33:1324-1330.
 - Sousa PH, Ramos AM, Maia GA, de Brito ES, Garruti DS, da Fonseca AV. Addition of *Ginkgo biloba* and *Panax ginseng* extracts to mixed tropical fruit nectars. *Cienc Technol Aliment* 2010;30:463-470.
 - Chan HH, Sun HD, Reddy MV, Wu TS. Potent alpha-glucosidase inhibitors from the roots of *Panax japonicus* C. A. Meyer var. major. *Phytochemistry* 2010;71:1360-1364.
 - Zhao CL, Cui XM, Chen YP, Liang Q. Key enzymes of triterpenoid saponin biosynthesis and the induction of their activities and gene expressions in plants. *Nat Prod Commun* 2010;5:1147-1158.
 - Son D, Lee JW, Lee P, Bae KH. Glycemic index of Insu 100 (R) herbal preparation containing Korean red ginseng, carob, mulberry, and banaba. *J Ginseng Res*

- 2010;34:89-92.
25. Lei FJ, Zhang AH, Xu YH, Zhang LX. Allelopathic effects of ginsenosides on *in vitro* growth and antioxidant enzymes activity of ginseng callus. *J Allelopath* 2010;26:13-22.
 26. Kim HJ, Joo SH, Choi I, Kim P, Kim MK, Park SH, Cheong JH, Shin CY. Effects of red ginseng on neonatal hypoxia-induced hyperactivity phenotype in rats. *J Ginseng Res* 2010;34:8-16.
 27. Lee MR, Yun BS, Liu L, Zhang DL, Wang Z, Wang CL, Gu LJ, Wang CY, Mo EK, Sung CK. Effect of black ginseng on memory improvement in the amnesic mice induced by scopolamine. *J Ginseng Res* 2010;34:51-58.
 28. Ma J, Li W, Tian R, Lei W. Ginsenoside Rg₁ promotes peripheral nerve regeneration in rat model of nerve crush injury. *Neurosci Lett* 2010;478:66-71.
 29. Ha SE, Shin DH, Kim HD, Shim SM, Kim HS, Kim BH, Lee JS, Park JK. Effects of ginsenoside Rg₂ on the ultraviolet B-induced DNA damage responses in HaCaT cells. *Naunyn Schmiedebergs Arch Pharmacol* 2010;382:89-101.
 30. Zhang YL, Chen JB, Lei Y, Zhou Q, Sun SQ, Noda I. Evaluation of different grades of ginseng using Fourier-transform infrared and two-dimensional infrared correlation spectroscopy. *J Mol Struct* 2010;974:94-102.
 31. Kim TH, Lee SM. The effects of ginseng total saponin, panaxadiol and panaxatriol on ischemia/reperfusion injury in isolated rat heart. *Food Chem Toxicol* 2010;48:1516-1520.
 32. Khalid S, Tahir M, Lone KP. Histological changes in the fetal brain of albino mouse after maternal treatment with *Panax ginseng*. *Pakistan J Zool* 2010;42:645-650.
 33. Dai J, Orsat V. Extraction of ginsenosides from American ginseng (*Panax quinquefolium* L.) root. *Int J Food Eng* 2010;6:article 3.
 34. Wang T, Yu X, Qu S, Xu H, Han B, Sui D. Effect of ginsenoside Rb₃ on myocardial injury and heart function impairment induced by isoproterenol in rats. *Eur J Pharmacol* 2010;636:121-125.
 35. Sharma SK, Bisht MS, Pandit MK. Synaptic mutation-driven male sterility in *Panax sikkimensis* Ban. (Araliaceae) from Eastern Himalaya, India. *Plant Syst Evol* 2010;287:29-36.
 36. Shen Y, Li YQ, Li SP, Ma L, Ding LJ, Ji H. Alleviation of ovariectomy-induced osteoporosis in rats by *Panax notoginseng* saponins. *J Nat Med* 2010;64:336-345.
 37. Kang KS, Yamabe N, Kim HY, Park JH, Yokozawa T. Effects of heat-processed ginseng and its active component ginsenoside 20(S)-Rg₃ on the progression of renal damage and dysfunction in type 2 diabetic Otsuka Long-Evans Tokushima Fatty rats. *Biol Pharm Bull* 2010;33:1077-1081.
 38. Dewir YH, Chakrabarty D, Wu CH, Hahn EJ, Jeon WK, Paek KY. Influences of polyunsaturated fatty acids (PUFAs) on growth and secondary metabolite accumulation in *Panax ginseng* C.A. Meyer adventitious roots cultured in air-lift bioreactors. *S Afr J Bot* 2010;76:354-358.
 39. Kim J, Lee JS, Shin A, Kang MH, Shin DS, Chung HR, Kim WK. Sociodemographic and lifestyle factors are associated with the use of dietary supplements in a Korean population. *J Epidemiol* 2010;20:197-203.
 40. Jager AK, Saaby L, Kudsk DS, Witt KC, Molgaard P. Short communication: influence of pasteurization on the active compounds in medicinal plants to be used in dairy products. *J Dairy Sci* 2010;93:2351-2353.
 41. Lee TK, O'Brien KF, Wang W, Johnke RM, Sheng C, Benhabib SM, Wang T, Allison RR. Radioprotective effect of American ginseng on human lymphocytes at 90 minutes postirradiation: a study of 40 cases. *J Altern Complement Med* 2010;16:561-567.
 42. Huang YC, Lin CY, Huang SF, Lin HC, Chang WL, Chang TC. Effect and mechanism of ginsenosides CK and Rg₁ on stimulation of glucose uptake in 3T3-L1 adipocytes. *J Agric Food Chem* 2010;58:6039-6047.
 43. Xu J, Seo AY, Vorobyeva DA, Carter CS, Anton SD, Lezza AM, Leeuwenburgh C. Beneficial effects of a Q-ter based nutritional mixture on functional performance, mitochondrial function, and oxidative stress in rats. *PLoS One* 2010;5:e10572.
 44. Kim CS, Jo YJ, Park SH, Kim HJ, Han JY, Hong JT, Cheong JH, Oh KW. Anti-stress effects of ginsenoside Rg(3)-standardized ginseng extract in restraint stressed animals. *Biomol Ther* 2010;18:219-225.
 45. Li Y, Tang J, Khatibi NH, Zhu M, Chen D, Zheng W, Wang S. Ginsenoside Rbeta1 reduces neurologic damage, is anti-apoptotic, and down-regulates p53 and BAX in subarachnoid hemorrhage. *Curr Neurovasc Res* 2010;7:85-94.
 46. Jovanovski E, Jenkins A, Dias AG, Peeva V, Sievenpiper J, Arnason JT, Rahelic D, Josse RG, Vuksan V. Effects of Korean red ginseng (*Panax ginseng* C.A. Mayer) and its isolated ginsenosides and polysaccharides on arterial stiffness in healthy individuals. *Am J Hypertens* 2010;23:469-472.
 47. Leonti M, Casu L, Raduner S, Cottiglia F, Floris C, Altmann KH, Gertsch J. Falcarinol is a covalent cannabinoid CB1 receptor antagonist and induces pro-allergic effects in skin. *Biochem Pharmacol* 2010;79:1815-1826.
 48. Chai H, Dong Y, Wang X, Zhou W. Ginsenoside Rb₁ attenuates homocysteine-augmented guidewire injury-

- induced intimal hyperplasia in mice. *J Surg Res* 2009;157:193-198.
49. Dickman JR, Koenig RT, Ji LL. American ginseng supplementation induces an oxidative stress in postmenopausal women. *J Am Coll Nutr* 2009;28:219-228.
50. Oh KJ, Chae MJ, Lee HS, Hong HD, Park K. Effects of Korean red ginseng on sexual arousal in menopausal women: placebo-controlled, double-blind crossover clinical study. *J Sex Med* 2010;7(4 Pt 1):1469-1477.
51. Chang CH, Liao TC, Hsu YM, Fang HW, Chen CC, Lin FH. A poly(propylene fumarate)--calcium phosphate based angiogenic injectable bone cement for femoral head osteonecrosis. *Biomaterials* 2010;31:4048-4055.
52. Zhang HA, Wang M, Zhou J, Yao QY, Ma JM, Jiang CL. Protective effect of ginsenoside against acute renal failure and expression of tyrosine hydroxylase in the locus coeruleus. *Physiol Res* 2010;59:61-70.
53. Rosner M. Geochemical and instrumental fundamentals for accurate and precise strontium isotope data of food samples: comment on 'determination of the strontium isotope ratio by ICP-MS ginseng as a tracer of regional origin.' *Food Chem* 2010;121:918-921.
54. Yoo JH, Kwon HC, Kim YJ, Park JH, Yang HO. KG-135, enriched with selected ginsenosides, inhibits the proliferation of human prostate cancer cells in culture and inhibits xenograft growth in athymic mice. *Cancer Lett* 2010;289:99-110.
55. Wahid F, Jung H, Khan T, Hwang KH, Kim YY. Effects of red ginseng extract on visual sensitivity and ERG b-wave of bullfrog's eye. *Planta Med* 2010;76:426-432.
56. Sagar SM. Can the therapeutic gain of radiotherapy be increased by concurrent administration of Asian botanicals? *Integr Cancer Ther* 2010;9:5-13.
57. Kim KR, Chung TY, Shin H, Son SH, Park KK, Choi JH, Chung WY. Red ginseng saponin extract attenuates murine collagen-induced arthritis by reducing pro-inflammatory responses and matrix metalloproteinase-3 expression. *Biol Pharm Bull* 2010;33:604-610.
58. Thomson GE. Further consideration of Asian medicinal plants in treating common chronic diseases in the West. *J Med Plant Res* 2010;4:125-130.
59. Ock SM, Hwang SS, Lee JS, Song CH, Ock CM. Dietary supplement use by South Korean adults: data from the national complementary and alternative medicine use survey (NCAMUS) in 2006. *Nutr Res Pract* 2010;4:69-74.
60. Abdel-Wahhab MA, Hassan NS, El-Kady AA, Khadrawy YA, El-Nekeety AA, Mohamed SR, Sharaf HA, Mannaa FA. Red ginseng extract protects against aflatoxin B1 and fumonisins-induced hepatic pre-cancerous lesions in rats. *Food Chem Toxicol* 2010;48:733-742.
61. Shin S, Jang JY, Park D, Yon JM, Baek IJ, Hwang BY, Nam SY, Yun YW, Kim KY, Joo SS *et al.* Korean red ginseng extract does not cause embryo-fetal death or abnormalities in mice. *Birth Defects Res B Dev Reprod Toxicol* 2010;89:78-85.
62. Wen HW, Hsieh MF, Wang YT, Chung HP, Hsieh PC, Lin IH, Chou FI. Application of gamma irradiation in ginseng for both photodegradation of pesticide pentachloronitrobenzene and microbial decontamination. *J Hazard Mater* 2010;176:280-287.
63. Quan C, Shang YG, Li SF, Tang SK, Huang T, Fang X. Kinetic study of supercritical fluid extraction of organochlorine pesticides from ginseng by Simulink (R) simulation. *J Taiwan Inst Chem Eng* 2010;41:44-48.
64. Lin WN, Lu HY, Lee MS, Yang SY, Chen HJ, Chang YS, Chang WT. Evaluation of the cultivation age of dried ginseng radix and its commercial products by using (1)H-NMR fingerprint analysis. *Am J Chin Med* 2010;38:205-218.
65. Kim DH, Chang JK, Shon HJ, Cho BG, Ko SR, Nho KB, Jang DS, Lee SM. Certification of a pure reference material for the ginsenoside Rg (1). *Accred Qual Assur* 2010;15:81-87.
66. Gravas S, Tzortzis V, Rountas C, Melekos MD. Extracorporeal shock-wave lithotripsy and garlic consumption: a lesson to learn. *Urol Res* 2010;38:61-63.
67. Li XY, Liang J, Tang YB, Zhou JG, Guan YY. Ginsenoside Rd prevents glutamate-induced apoptosis in rat cortical neurons. *Clin Exp Pharmacol Physiol* 2010;37:199-204.
68. Kim SH, Lee HJ, Kim JS, Moon C, Kim JC, Park HR, Jung U, Jang JS, Jo SK. Protective effect of an herbal preparation (HemoHIM) on radiation-induced intestinal injury in mice. *J Med Food* 2009;12:1353-1358.