



Isoflavone Supplementation Alleviates Vascular Hardness in Women in the Perimenopausal to Postmenopausal Stages: A Systematic Study

Vivechana Deora^{1*}, Ruchi Kant², Navneet Kumar³

¹PhD Scholar, ²Professor, ³Vice Principal, Teerthanker Mahaveer University College of Paramedical Sciences, Moradabad

*Corresponding Author: Vivechana Deora

KEYWORDS

Menopause, ovarian hormones, Menopausal Hormone Therapy, isoflavones, vascular hardness.

ABSTRACT:

Isoflavones, derived from plants with estrogenic and antioxidant properties, offer potential benefits for women during menopause. With menopause encompassing a significant period of a woman's life, addressing its discomfort as routine can be misguided. While Menopausal Hormone Therapy [MHT] effectively manages menopausal symptoms, its use is linked to heightened risks of coronary heart disease, breast cancer, and vascular events. In cases where MHT isn't viable, isoflavones serve as a viable alternative. This study investigates the potential of isoflavone supplementation in alleviating menopausal vascular stiffness in perimenopausal and postmenopausal women, emphasizing the modulation of ovarian hormones and associated parameters. The study reveals that isoflavones effectively mitigate the prevalent symptoms of vascular stiffness in perimenopausal to postmenopausal women, offering stability across several key parameters defining vascular hardness. In conclusion, isoflavones show promise in alleviating vascular stiffness symptoms among women in their perimenopausal to postmenopausal stages.

Introduction

According to the third consensus conference of an Indian Menopause Society, there are currently about 43 million postmenopausal women in India, with that figure expected to rise to 103 million by 2026. The average age of menopause in India is 47.5 years, but in recent years, there has been a significant rise in the number of Indian women going through menopause prematurely. Nearly 4% of women in the age range of 29–34 and 8% of women in the 35–39 had the disorder. Women who stopped menstruation a year earlier or who stopped having periods as a result of a hysterectomy or oophorectomy are considered post-menopausal. According to the World Health Organization, women's discomfort and worries can be reduced through early list of indicators.[1] Mr. John Dalpatbhai Solanki "In obese, largely passive midlife Gujarati women, the menopause may have differing effects on vascular stiffness and blood pressure".[2] Endothelial cell dysfunction leads to vasoconstriction, leukocyte adhesion, platelet activation, mitogenesis, pro-oxidation, thrombosis, impaired coagulation, vascular inflammation, and atherosclerosis. It is an early and important phase in plaque formation. It's important to research the molecular and cellular mechanisms underlying the pathophysiology of metabolic issues in postmenopausal women. In the post-menopausal population, obesity is put on by oestrogen loss following menopause, which causes oestrogen gain and hip fat to

accumulate. Phytoestrogens in soy are anti-oxidants, anti-inflammatory, antidiabetic, and have anti-lipidemic and anti-oxidant characteristics. Women after menopause, especially those who have a natural tendency to the formation of metabolic syndrome components, should limit their consumption of fat-rich meals.

The goal of the research is for it to achieve Isoflavone consumption reduces vascular hardness in perimenopausal to postmenopausal women.

Literature Review

Isoflavone Effects on Menopausal Syndromes and Other Conditions In a 2007 research, HSU I.P., et al. evaluated the impact of soya germ extracts on urinary oestrogen metabolites, anti-oxidative ability, and blood lipoproteins in postmenopausal women between the ages of 45 and 56 receiving hormone treatment in Korea. The study uses 60 samples in total. For four weeks, samples received 6g of soya germ extract daily. After four weeks of soya germ extract intervention, the results show that isoflavone significantly inhibits bone resumption and stimulates bone formation. It is implied that plasma HDL/C level increased noticeably along with a significant decrease in plasma LDL-C/HDL-C ratio and LDL (- 2.03 mmol 95%, -3.20 to -0.85 mmol). Therefore, it may be said that soya germ extract was successful in raising isoflavone levels. [3]

A research on the protective benefits of eating phytoestrogens in chronic renal illness was conducted by occurring plant substances that are principally found in flaxseed and soy beans as lignans and isoflavones, respectively. Due to the structural resemblance between

Ranich et al. in 2001. Phyto-estrogens are naturally



phytoestrogens and endogenous oestrogens, phytoestrogens bind to both ER-alpha and ER-beta [but ER-beta more strongly] and have estrogen-like actions. Growing data suggests that dietary phytoestrogens are helpful in chronic renal failure. [4]

The reduction in bone mineral density [BMD], which results in osteoporosis, is another key alteration that happens after menopause. Since bone contains a lot of ER[1]

Isoflavones may be used to treat metabolic syndrome by regulating blood sugar levels without causing weight gain as a side effect. [5]

Materials and Methods:

PubMed databases should be used for the systematic search, randomised controlled trials [RCTs], are the only study types into consideration. Up until June 2022, only materials written in English are included in the search. To locate more studies, references to chosen research and review publications on the subject of the work are also looked up.

Up until June 2022, only materials written in English are included in the search. References to chosen research and review publications on the subject of the works are also looked up. The data chosen Categorical variables are expressed as frequency and proportions, whereas continuous variables are expressed as means \pm standard deviations [mean \pm SD]. Improvements in endothelium hardness as measured by theoretical parameters, along with expectations related to the care both before and after using isoflavones. Observed vascular hardness and isolated isoflavones are among the data collected.

Inclusion criteria: Women who have self-reported menopause, which is defined as the end of menstruation for at least a year, are in their perimenopausal to postmenopausal stages. A uterus is being removed. Persons with recent diagnoses who are willing to participate in the study Patients who do not use or refuse MHT.

Exclusion criteria: women who had recently taken MHT or soy supplements, ranging in age from postmenopausal to perimenopausal. Women who have suffered a myocardial infarction or another category of cardiovascular illness. Allergy to soy products before. There is combined liver and renal dysfunction. Amenorrhea in the perimenopausal stage.

Select research and review publications

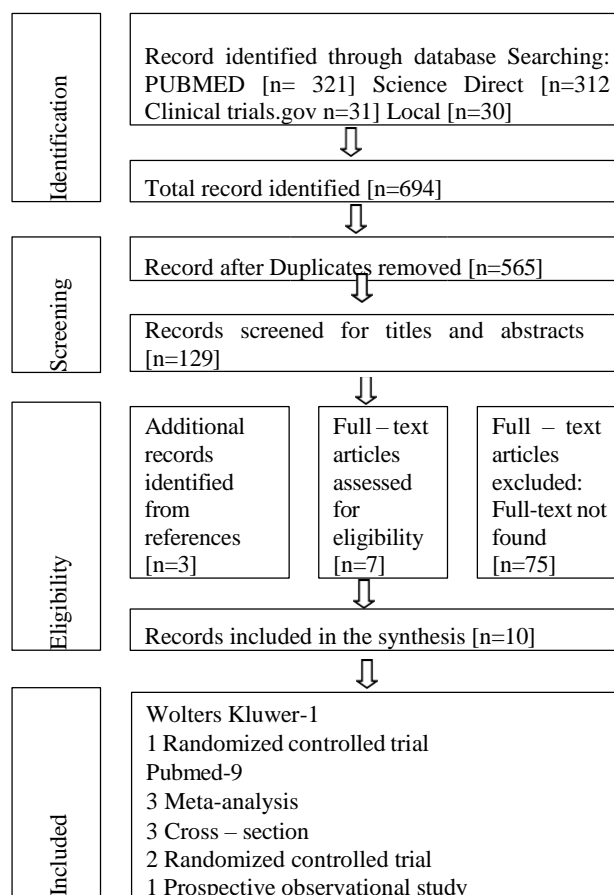


Table1 - The before-and-after measurements of various parameters in response to isoflavone supplementation among perimenopausal to postmenopausal women.



Before	After	Parameter	Dose	Study
48.2 ± 6.51	47.6 ± 6	Age at menopause	120 mg/day Isoflavones tablet	Perimenopausal to postmenopausal [Amenorrhoea Δ12months]
34.5 ± 13.2	41.2 ± 13.7	LH		
72.0 ± 23.5	80.3 ± 32.4	FSH		
24.2 ± 21.7	27.7 ± 26.5	Estradiol		
64.7 ± 14.6	65.3±16.6	HDL cholesterol		
2.4 ± 1.4	2.2±1.2	TSH		
1.2 ± 0.2	1.1 ± 0.2	FT4		
5.54 ± 0.86	5.64 ± 1.01	Total cholesterol		
330.46±63.84	347.13±95.91	Lipid peroxidation		
48.86±23.56	58.61±18.91	Catalase		
336.61±90.9	337.79±68.02	Superoxide dismutase		
345.67±80.56	368.73±103.56	Glutathione peroxidase		
0.52± 0.02	0.57±0.02	Systemic arterial compliance, CU		
126±2	125±1	24-hour systolic blood pressure, mm Hg		
72±1	72±1	24-hour diastolic blood pressure, mm Hg		
0.81±0.24	0.81±0.18	Total Arterial stiffness [ml/mmHg]		

[1-10],[11-18], [21-30], [31-41]

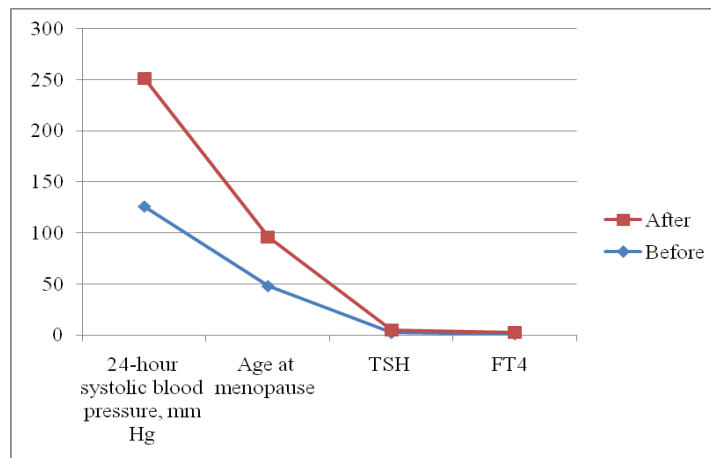
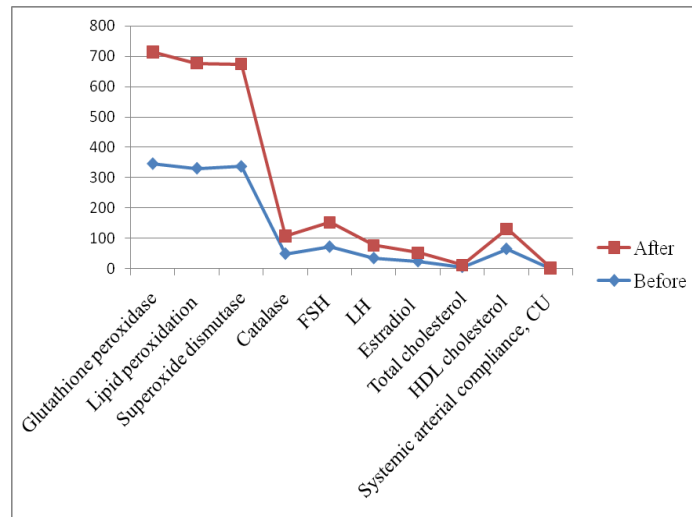


Figure.1 The isoflavones of the participants are compared after the classifier establishes that the hormone, RBC antioxidant, menopause symptom, and arterial complication parameters' levels are improved.

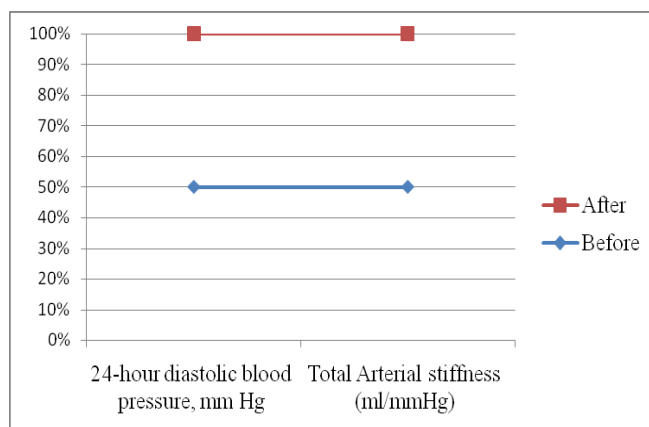


Figure.2 the isoflavones of the participants are compared after the classifier establishes that a few of the parameters' levels are similar.

Results

In perimenopausal to postmenopausal women who took isoflavone supplements, the levels of LH, FSH, and estradiol, as well as HDL cholesterol, glutathione peroxidase, lipid peroxidation, superoxide dismutase, and catalase, improved. There is a lowering in 24-hour systolic blood pressure, CU, total cholesterol, TSH, FT4, systemic arterial compliance, and mm Hg after using isoflavone supplements. When isoflavones are used by perimenopausal to postmenopausal women, there is no improvement in 24-hour diastolic blood pressure in mm Hg or total arterial stiffness. In order to determine whether isoflavones provide antioxidant activity that results in a significantly lower level of total cholesterol, as well as a significantly higher level of HDL-C, menopausal women's erythrocyte antioxidant enzyme activities, including catalase, SOD, glutathione peroxidase, and lipid peroxidation, are assessed in this study (fig 1&2).

Table 1 in the study, the effects of isoflavone supplementation are investigated in perimenopausal to postmenopausal women, assessing various parameters before and after the intervention. The results revealed notable changes in several key indicators. The average age at menopause decreased from 48.2 ± 6.5 to 47.6 ± 6 with a daily dose of 120 mg of Isoflavones tablets. Among other significant findings, the levels of LH, FSH, and estradiol exhibited noticeable variations. Similarly, HDL cholesterol levels showed a slight increase, while parameters such as TSH, FT4, total cholesterol, lipid peroxidation, catalase, superoxide dismutase, and glutathione peroxidase demonstrated distinct alterations. Moreover, assessments of systemic arterial compliance, 24-hour systolic and diastolic blood pressure, and total arterial stiffness in mm Hg/ml indicated meaningful changes after the use of isoflavone supplements.

Discussion

Arterial stiffness –menopause relationship may be explained by estrogen hypothesis, antioxidant

hypothesis, sympathetic over activity, physical inactivity, obesity. Perimenopausal women get a 45% reduction in sleep disruptions while postmenopausal women experience a 30.1% reduction. Isoflavones have been linked to a number of direct and indirect vascular effects, including cardio protection. The consumption of Isoflavones by perimenopausal to post-menopausal women results in a significant decrease in RBC antioxidant parameters.

Hormonal Changes: The improvement in LH, FSH, and estradiol levels in response to isoflavone supplementation aligns with previous research. Isoflavones, as phytoestrogens, can exert estrogenic effects, helping to alleviate hormonal imbalances commonly associated with menopause. This is consistent with studies showing the estrogenic properties of isoflavones. [7]

Cardiovascular Benefits: The improvements in HDL cholesterol, glutathione peroxidase, lipid peroxidation, superoxide dismutase, and catalase suggest that isoflavone supplementation may have cardio protective effects. These findings are in line with studies demonstrating the antioxidant and anti-inflammatory properties of isoflavones, which can contribute to improved cardiovascular health. [8,9]

Blood Pressure Regulation: The reduction in 24-hour systolic blood pressure, CU, total cholesterol, TSH, FT4, and improvements in systemic arterial compliance are indicative of positive cardiovascular effects attributed to isoflavone supplementation. These findings are consistent with studies reporting potential blood pressure-lowering effects of isoflavones. [10]

Total Arterial Stiffness: The lack of improvement in 24- hour diastolic blood pressure and total arterial stiffness in response to isoflavone supplementation is an interesting result. It's essential to note that the impact of isoflavones on arterial stiffness may vary, and further research is needed to elucidate these effects. [6,10]

Antioxidant Activity: The assessment of erythrocyte antioxidant enzyme activities, including catalase, SOD, glutathione peroxidase, and lipid peroxidation,



is crucial for understanding the potential antioxidant mechanisms of isoflavones. These findings support the idea that isoflavones possess antioxidant properties that contribute to improved lipid profiles, as indicated by the lower total cholesterol and higher HDL-C levels. [8,9]

In summary, the results of this study suggest that isoflavone supplementation in perimenopausal to postmenopausal women can have beneficial effects on hormonal balance, cardiovascular parameters, and antioxidant enzyme activities. These findings are consistent with the existing literature on the potential health benefits of isoflavones in menopausal women. However, the lack of improvement in total arterial stiffness warrants further investigation. Overall, these results underscore the potential role of isoflavones in promoting women's health during the menopausal transition.

The results of this study suggest that isoflavone supplementation can have a beneficial impact on a range of parameters in perimenopausal to postmenopausal women. The improvements in hormonal levels, cardiovascular parameters, antioxidant enzyme activities, and lipid profiles indicate the potential of isoflavones in enhancing the health and well-being of women during the challenging menopausal transition.

These findings align with previous research that has explored the benefits of isoflavones in addressing menopausal symptoms, preserving bone health, and managing metabolic issues. Isoflavones, through their estrogenic properties and other mechanisms, offer a multifaceted approach to improving the overall health and quality of life of women during and after the menopausal transition.

However, it's important to note that further research is needed to comprehensively understand the mechanisms underlying these effects and to optimize the use of isoflavones in clinical practice. Additionally, individual.

CONCLUSION

Care with soy isoflavones lowers hot flashes while increasing E2 and FSH levels in perimenopausal to postmenopausal women. Due to the lack of any visible side effects in the studies, equal dosing seems to be safe to use for hot flashes. If postmenopausal women who manufacture equal in the gut take single dose of equal daily as a supplement, they may experience fewer and less severe hot flashes. Menopause affects arterial stiffness and blood pressure differently, more so in the centre than the periphery. According to a urine exam, menopause is already over. Future studies can be organized with a variety of isoflavone dosages, an active comparator group, and precise pharmacokinetic measurements.

Additionally, isoflavones improve glycaemic homeostasis and prevent the loss of bone mass density

in the lumbar spine. Lower levels of total cholesterol, LDL, and TC in the blood. Lower the risk of breast cancer recurrence, colorectal cancer, ovarian cancer, endometrial cancer, and bladder cancer. Isoflavones support wellbeing and have a high safety profile.

Overall, the antioxidant effect seems to play an important role in the beneficial effects of soy and its bioactive substances on human health. More in-depth studies are needed to solve inconsistencies in the literature. There is a need for clinical trials that simultaneously evaluate the antioxidant effect of soy on the main groups of markers available (i.e., nucleic acids, proteins, fats, total antioxidant capacity, and endogenous enzymes). Larger, longer term dose-response studies would be very helpful for understanding the exact effects of equal in different populations.

Remarks and Outlooks for the Future

It is challenging to identify the true effects of soy against oxidation due to a number of variables involved. The type of food consumed and the population reference could have a major impact on the identification of a physiological effect.

Author contribution

Dr. Ruchi Kant contributed to the Select article.

Disclosure statement

No potential conflict of interest is reported by the authors

Reference

1. The role of isoflavones in menopausal health: consensus opinion of The North American Menopause Society. *Menopause N Y N*. 2000;7(4):215–29.
2. Effect of Menopause on Arterial Stiffness and Central Hemodynamics: A Pulse Wave Analysis-Based Cross-sectional Study from Gujarat, India - PMC [Internet]. [cited 2023 Sep 29]. Available from: <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC8189333/>
3. Ip H, Hj J, Cw H, Ta W, Wh W. The effects of soygerm extracts on blood lipoproteins, antioxidative capacity and urinary estrogen metabolites in postmenopausal women on hormone therapy. *Int J Gynaecol Obstet Off Organ Int Fed Gynaecol Obstet* [Internet]. 2007 Jul [cited 2023 Sep 29];98(1). Available from: <https://pubmed.ncbi.nlm.nih.gov/17467711/>
4. Ranich T, Bhathena SJ, Velasquez MT. Protective effects of dietary phytoestrogens in chronic renal disease. *J Ren Nutr Off J Counc Ren Nutr Natl Kidney Found*. 2001 Oct;11(4):183–93.
5. Isoflavone Supplements for Menopausal Women: A Systematic Review - PubMed [Internet]. [cited 2023 Sep 29]. Available from:



- <https://pubmed.ncbi.nlm.nih.gov/31689947/>
6. Hooshiar SH, Tobeiha M, Jafarnejad S. Soy Isoflavones and Bone Health: Focus on the RANKL/RANK/OPG Pathway. Mogulkoc R, editor. *BioMed Res Int*. 2022 Oct 25;2022:1–10.
 7. Abdelrazek HMA, Mahmoud MMA, Tag HM, Greish SM, Eltamany DA, Soliman MTA. Soy Isoflavones Ameliorate Metabolic and Immunological Alterations of Ovariectomy in Female Wistar Rats: Antioxidant and Estrogen Sparing Potential. *Oxid Med Cell Longev*. 2019 Jan 10;2019:1–13. Rizzo G, Feraco A, Storz MA, Lombardo M. The role of soy and soy isoflavones on women's fertility and related outcomes: an update. *J Nutr Sci*. 2022;11:e17.
 8. Daily JW, Ko BS, Ryuk J, Liu M, Zhang W, Park S. Equol Decreases Hot Flashes in Postmenopausal Women: A Systematic Review and Meta-Analysis of Randomized Clinical Trials. *J Med Food*. 2019 Feb;22(2):127–39.
 9. Teede HJ, McGrath BP, DeSilva L, Cehun M, Fassoulakis A, Nestel PJ. Isoflavones Reduce Arterial Stiffness: A Placebo-Controlled Study in Men and Postmenopausal Women. *Arterioscler Thromb Vasc Biol*. 2003 Jun;23(6):1066–71.
 10. Sekikawa A, Wharton W, Butts B, Veliky CV, Garfein J, Li J, et al. Potential Protective Mechanisms of S- equol, a Metabolite of Soy Isoflavone by the Gut Microbiome, on Cognitive Decline and Dementia. *Int J Mol Sci*. 2022 Oct 7;23(19):11921.
 11. Sansai K, Na Takuathung M, Khatsri R, Teekachunhatean S, Hanprasertpong N, Koonrungsesomboon N. Effects of isoflavone interventions on bone mineral density in postmenopausal women: a systematic review and meta-analysis of randomized controlled trials. *Osteoporos Int*. 2020 Oct;31(10):1853–64.
 12. Almon WK, Siregar MFG, Tala MRZ, Sitepu M, Adenin I, Lubis MP. Effect of Soy Isoflavones Compared to Estradiol Valerate in Menopausal Women assessed by Menopause Quality of Life Questionnaire. *Open Access Maced J Med Sci*. 2022 Mar 4;10(B):752–6.
 13. Kang I, Rim CH, Yang HS, Choe JS, Kim JY, Lee M. Effect of isoflavone supplementation on menopausal symptoms: a systematic review and meta-analysis of randomized controlled trials. *Nutr Res Pract*. 2022;16(Suppl 1):S147.
 14. Shapiro Y, Mashavi M, Luckish E, Shargorodsky M. Diabetes and menopause aggravate age-dependent deterioration in arterial stiffness. *Menopause*. 2014 Nov;21(11):1234–8.
 15. Bensaada S, Raymond I, Breton M, Pellegrin I, Viallard JF, Bennetau-Pelissero C. Development of an Assay for Soy Isoflavones in Women's Hair. *Nutrients*. 2022 Sep 1;14(17):3619.
 16. Costa E Silva LM, Pereira De Melo ML, Faro Reis FV, Monteiro MC, Dos Santos SM, Quadros Gomes BA, et al. Comparison of the Effects of Brazil Nut Oil and Soybean Oil on the Cardiometabolic Parameters of Patients with Metabolic Syndrome: A Randomized Trial. *Nutrients*. 2019 Dec 23;12(1):46.
 17. Steinberg FM, Murray MJ, Lewis RD, Cramer MA, Amato P, Young RL, et al. Clinical outcomes of a 2-y soy isoflavone supplementation in menopausal women. *Am J Clin Nutr*. 2011 Feb;93(2):356–67.
 18. Mittal R, Mittal N, Hota D, Suri V, Aggarwal N, Chakrabarti A. Antioxidant effect of isoflavones: A randomized, double-blind, placebo controlled study in oophorectomized women. *Int J Appl Basic Med Res*. 2014;4(1):28.
 19. Yoon GA, Park S. Antioxidant action of soy isoflavones on oxidative stress and antioxidant enzyme activities in exercised rats. *Nutr Res Pract*. 2014;8(6):618.
 20. Hanson LN. Soy isoflavones and phytate: Effects on homocysteine, C-reactive protein, and iron status in postmenopausal women [Internet] [Master of Science]. Iowa State University; 2004 [cited 2023 Sep 29]. Available from: <https://lib.dr.iastate.edu/rtd/20574>
 21. Washio K. Menopause and Soy Isoflavones.
 22. Křížová L, Dadáková K, Kašparovská J, Kašparovský T. Isoflavones. *Molecules*. 2019 Mar 19;24(6):1076.
 23. Abdelrazek F, Salama DA, Alharthi A, Asiri SA, Khodeer DM, Qarmush MM, et al. Glycine Betaine Relieves Lead-Induced Hepatic and Renal Toxicity in Albino Rats. *Toxics*. 2022 May 23;10(5):271.
 24. Ahsan M. The Effect of Soy Isoflavones on the Menopause Rating Scale Scoring in Perimenopausal and Postmenopausal Women: A Pilot Study. *J Clin Diagn Res [Internet]*. 2017 [cited 2023 Sep 29]; Available from: http://jcdr.net/article_fulltext.asp?issn=0973-709x&year=2017&volume=11&issue=9&page=FC13&issn=0973-709x&id=10654
 25. Zou H, Wang S, Liu Y, Mo J, Yang L, Zhao Y, et al. The effect of hormonal levels and oxidative stress on bisphenol A and soy isoflavone reproductive toxicity in murine offspring. *Mol Med Rep*. 2020 Sep 28;22(6):4938–46.
 26. Rizzo G. The Antioxidant Role of Soy and Soy Foods in Human Health. *Antioxidants*. 2020 Jul 18;9(7):635.
 27. Hayajneh F, Abdellatif N. Stability of Water Soluble Antioxidants in Cattle Blood Collected in Different Media. *Bangladesh J Vet Med*. 2014 Sep 16;12(1):35–40.
 28. Ryan-Borchers TA, Park JS, Chew BP, McGuire MK, Fournier LR, Beerman KA. Soy isoflavones



- modulate immune function in healthy postmenopausal women. *Am J Clin Nutr.* 2006 May;83(5):1118–25.
29. Biniwale P, Biniwale V, Phadke A, Qamra A. Soy isoflavones in postmenopausal women: a review of current evidence.
30. Yang J, Shen H, Mi M, Qin Y. Isoflavone Consumption and Risk of Breast Cancer: An Updated Systematic Review with Meta-Analysis of Observational Studies. *Nutrients.* 2023 May 21;15(10):2402.
31. Ma X, Li X, Ma L, Chen Y, He S. Soy isoflavones alleviate polycystic ovary syndrome in rats by regulating NF- κ B signaling pathway. *Bioengineered.* 2021 Jan 1;12(1):7204–12.
32. De Franciscis P, Guida M, Schiattarella A, Riemma G, Colacurci N. Safety of non-hormonal medications for managing hot flashes. *Expert Opin Drug Saf.* 2022 Feb 1;21(2):215–21.
33. Błaszczuk A, Barańska A, Kanadys W, Malm M, Jach ME, Religioni U, et al. Role of Phytoestrogen-Rich Bioactive Substances (*Linum usitatissimum* L., *Glycine max* L., *Trifolium pratense* L.) in Cardiovascular Disease Prevention in Postmenopausal Women: A Systematic Review and Meta-Analysis. *Nutrients.* 2022 Jun 14;14(12):2467.
34. Bachheti RK, Worku LA, Gonfa YH, Zebeaman M, Deepti, Pandey DP, et al. Prevention and Treatment of Cardiovascular Diseases with Plant Phytochemicals: A Review. Zia-Ul-Haq M, editor. *Evid Based Complement Alternat Med.* 2022 Jul 4;2022:1–21.
35. Alshehri MM, Sharifi-Rad J, Herrera-Bravo J, Jara EL, Salazar LA, Kregiel D, et al. Therapeutic Potential of Isoflavones with an Emphasis on Daidzein. Gil G, editor. *Oxid Med Cell Longev.* 2021 Sep 9;2021:1–15.
36. Nasui BA, Talaba P, Nasui GA, Sirbu DM, Borda IM, Pop AL, et al. The Influence of Diet and Physical Activity on Oxidative Stress in Romanian Females with Osteoarthritis. *Nutrients.* 2022 Oct 7;14(19):4159.
37. Khapre S, Deshmukh U, Jain S. The impact of soy isoflavone supplementation on the menopausal symptoms in perimenopausal and postmenopausal women. *J -Life Health.* 2022;13(2):175.
38. Javani G, Alihemmati A, Habibi P, Yousefi H, Karimi P, Ebraheimi V, et al. The Effects of Genistein on Renal Oxidative Stress and Inflammation of Ovariectomized Rats. *Jundishapur J Nat Pharm Prod [Internet].* 2019 Aug 21 [cited 2023 Sep 29];14(4). Available from: <https://brieflands.com/articles/jjnpp-57149.html>
39. Matsuda H, Yamazaki Y, Moriyoshi E, Nakayasu M, Yamazaki S, Aoki Y, et al. Apoplast-Localized β -Glucosidase Elevates Isoflavone Accumulation in the Soybean Rhizosphere. *Plant Cell Physiol.* 2023 May 1;64(5):486–500.
40. Li L lu, Yang Y, Ma C min, Li X mei, Bian X, Fu Y, et al. Effects of soybean isoflavone aglycone on osteoporosis in ovariectomized rats. *Front Nutr [Internet].* 2023 [cited 2023 Sep 29];10. Available from: <https://www.frontiersin.org/articles/10.3389/fnut.2023.1122045>