Effectiveness of Messenger Plants (*Peperomia Pellucida L.*) and *Tai Chi* Exercise as Modality Therapy for Elderly with Hyperuricemia: A Literature review

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ABSTRACT

Hyperuricemia correlated with an increase in monosodium urate crystals, which was a precipitating factor for gout arthritis. In Indonesia, this disease was more common in individuals over 34 years of age. The elderly who had gout can be associated with metabolic syndrome and increase the risk of cardiovascular disease. Pharmacological therapy of gout, such as allopurinol, had a greater potential to cause side effects in patients. This article aimed to determine the effectiveness of *Peperomia pellucida* and *Tai Chi* exercise to overcome hyperuricemia in elderly. The writing method used was a literature review. Articles were obtained through search sites: Google Scholar, Science-Direct, and NCBI, and the criteria used were Indonesian and English with a range of publication years 2014-2020. Eight main journals were used as references; from these journals, it was concluded that *Peperomia pellucida* could inhibit the formation of uric acid with xanthine oxidase activity through its quercetin-type flavonoid content in elderly, while *Tai Chi* exercise could increase joint flexibility and blood circulation in patients with gout. Therefore, it is hoped that the nurse can consider this modality therapy to be given to elderly with hyperuricemia. So, elderly with hyperuricemia can control their uric acid levels by implementing this combination therapy.

Keywords: Gout; *Peperomia Pellucida*; *Tai Chi*

INTRODUCTION

Gout is a degenerative disease caused by the build-up of monosodium urate crystals in the body, especially in the joint area. A person is said to have gouty arthritis if the uric acid level in the blood is above normal limits (hyperuricemia), and the results of microscopic examination of joint fluid reveal monosodium urate crystals. This build-up of crystals can occur anywhere but is most common in the joints. Crystal deposition that occurs repeatedly will cause inflammation (Susianti et al., 2015). The typical clinical manifestation of gouty arthritis is joint enlargement which affects more than one joint for a longer duration. Hyperuricemia in the elderly is associated with metabolic syndrome and increases the risk of comorbid diseases such as cardiovascular disease, and it is related to the enzyme xanthine oxidase and endothelial damage to blood vessels (Puspasari et al., 2017).

Patients with gout in the world continued to experience a two-fold increase between 1990-2010. The prevalence of hyperuricemia and gout in Asia in the last decade has reached 13%-25%. It is estimated that gout occurs in 840 people out of every 100,000 people. In Indonesia, gout at the age of under 34 years is 32%, and age above 34 is 68% (Hastuti et al., 2018). According to the World Health Organization (WHO), 24% of gout sufferers in Indonesia go to doctors or other health workers, while 71% tend only to take over-the-counter pain relievers. Gout arthritis in Indonesia occupies the second position as a joint disease after osteoporosis (Jaliana et al., 2018).

Recommendations for gout management are pharmacological therapy and management of diet and lifestyle patterns. The recommended diet to reduce uric acid levels is to avoid consuming foods that contain lots of purines, such as organ meats and foods or fruit juices that contain high-fructose corn syrup (Harding, 2016). Pharmacological therapy such as allopurinol can cause more significant side effects than non-pharmacological therapy using herbal plants and physical exercise. In addition, non-pharmacological therapies tend to be in demand because they are relatively cheap and easy to find. However, not much is known about its benefits as a therapy for hyperuricemia.

Suruhan plants (*Peperomia pellucida L.*) have been used in traditional medicine for thousands of years to treat various diseases, such as gout (Prhastuti et al., 2017). Several studies have shown that *Peperomia Pellucida* has analgesic, anti-inflammatory, antipyretic, antioxidant, antihyperglycemic, anti-hyperuricemic, cytotoxic, antimicrobial,
lipase inhibitor, and anti-osteoporosis effects (Farida et al. 2016). In addition, Peperomia Pellucida contains many alkaloids, tannins, saponins, quinines, and flavonoids that can be used to treat gout and skin inflammation. The optimal level of flavonoid in Peperomia Pellucida extract to inhibit xanthine oxidase activity is 300 gr. (Kartika et al., 2016). Based on Mutee’s research (2010), the dose used is 1000mg/kg body weight per day orally in male rats with oedema in the hind legs. The results show that herbal messengers can reduce uric acid levels not much different from allopurinol as a uric acid synthesis drug in mice (Bakar, A et al., 2018).

In addition to consuming ordered herbal plants, gout sufferers can also do Tai Chi Gymnastics. Tai Chi is an exercise regimen rooted in eastern philosophy and Chinese medicine (Holly Black 2015). This gymnastics is an exercise that combines physical movements, breathing, feelings, and thoughts in a single unit that aims to produce inner peace that has physical and psychological therapeutic value (Blake and Hawley, 2016). Thus, the supply of food and oxygen in the muscle tissue becomes better so that muscle flexibility increases. In addition, according to Komang (2017), tai chi exercise interventions can reduce joint pain in the elderly with hyperuricemia, rheumatism, and other joint pain (Kisner, 2014, Komang 2017).

Based on the above phenomenon, the authors are interested in conducting a literature review that aims to determine the Peperomia pellucida and tai chi exercise as therapy in older adults with gout by minimizing the side effects of therapy. Furthermore, it is hoped that this study can provide an overview of alternative therapies for nurse and the broader community in utilizing messenger plants that are very easy to get in the tropical environment of Indonesia and doing tai chi exercises.

**METHOD**

The method used in writing this article is a literature review. The literature used is articles obtained from electronic media obtained from Google Scholar, Science Direct, and NCBI through “Peperomia Pellucida and Gout”, “Tai Chi Exercise and Gout”, and “Peperomia Pellucida, Tai Chi Exercise and Gout”. The criteria used in the search for articles were articles in Indonesian and English, published in 2015-2020, and articles reviewing the effectiveness of Peperomia pellucida and tai chi exercises for reducing hyperuricemia levels or related to joint disease, which has characteristics similar to gout. The literature collected according to the criteria will be analyzed and synthesized by categorizing it into problem analysis and potential problem solving so that an idea emerges based on scientific evidence and supported by previous research. The guidelines in writing this literature review use the PRISMA (Preferred Reporting Items for Systematic Reviews and Meta-analyses) method. To assess the feasibility of articles to be included in this study, the authors use the Critical Appraisal Skills Program (CASP).

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Chart 1. Article Selection Using PRISMA Diagram
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Potential and relevant articles based on identification from electronic databases (n=1,048) That are, Science Direct (n=87), NCBI (n=118), and Google Scholar (n=843)

Records were rejected because the topic was not relevant (n =824), duplicated records remove (n=187)

Full-text articles excluded according to inclusion and exclusion criteria (n=17)

Record screened (n=37)

Full-text articles assessed for eligibility (n=20)

Studies focus included in the literature review (n=8)
RESULT

Table 1. Journal analysis of 8 main articles included in this study

<table>
<thead>
<tr>
<th>Author</th>
<th>Year</th>
<th>Title of Articles</th>
<th>Method</th>
<th>Result &amp; Discussion</th>
<th>Conclusion</th>
</tr>
</thead>
<tbody>
<tr>
<td>Asiyah Dwiyaningsih</td>
<td>2017</td>
<td>The Effect of Ergonomic Gymnastics and Tai Chi Exercises on Reducing Uric Acid Levels in the Elderly</td>
<td>Quasi-experiment with pre-test and post-test two group design</td>
<td>Based on the results of research from 60 elderly respondents, it was found that there was an effect of decreasing uric acid levels before and after giving Tai Chi exercises</td>
<td>There are differences in ergonomic exercise and tai chi exercise in reducing uric acid levels in the elderly.</td>
</tr>
<tr>
<td>Riska Yunda B</td>
<td>2018</td>
<td>The effect of Tai Chi exercise on joint pain and joint flexibility in the elderly at the Elderly Posyandu</td>
<td>Quasi-experiment with One Group Pretest Posttest design</td>
<td>After the six Tai Chi interventions, the elderly experienced a decrease in the level of moderate joint pain from the number of respondents 24 to 8 respondents. This study showed a significant difference after and before the intervention.</td>
<td>There is an effect of giving Tai Chi exercise intervention on joint pain with a p-value of 0.000, and there is an effect on joint flexibility with a p-value of 0.001</td>
</tr>
<tr>
<td>Komang Tri Adi, Made Muliarta, Muhammad Irfan</td>
<td>2017</td>
<td>Tai Chi Gymnastics Is More Effective In Increasing Flexibility And Balance Than Elderly Fitness Gymnastics In The Elderly In Denpasar City</td>
<td>Quasi-experiment with pre-test and post-test two group design</td>
<td>The results showed that there were differences in the mean flexibility and balance between the two groups (Group 1 Tai Chi and other fitness exercises) using the independent T-test, which showed a significant difference with the average flexibility value of 3.15 cm and -3.15/sec</td>
<td>Tai Chi Gymnastics and Elderly Fitness Gymnastics are effective in increasing flexibility and balance in the elderly in Denpasar City, but Tai Chi Gymnastics is more effective in increasing flexibility and balance in the elderly in Denpasar City.</td>
</tr>
<tr>
<td>Suci Indah P, Diyan Indriyani and Yeni Suryaningsih</td>
<td>2015</td>
<td>The Effect of Tai Chi Exercise on Joint Pain in the Elderly at Nursing Home Bondowoso</td>
<td>Pra-Eksperimental with one-group pretest and postest design</td>
<td>After the Tai Chi exercise intervention in the elderly, the results showed a difference in the pain value of 4.5758 and after being given gymnastics tai chi 1.1818</td>
<td>This research is recommended to health agencies to provide tai chi exercises regularly, especially at the posyandu for the elderly.</td>
</tr>
<tr>
<td>Victor Manuel Mendozañez, Taide laurita Arista-Ugalde, Juana rosado-Pérez, Mirma ruiz-ramos, and edelmiro santiago-Osorio</td>
<td>2018</td>
<td>hypoglycemic and antioxidant effect of Tai chi exercise training in older adults with metabolic syndrome</td>
<td>Quasi-experiment with pre-test and post-test two group design</td>
<td>A very statistically significant decrease in the concentration of HbA1c was observed in the Tai Chi exercise group compared to the control group, and the intervention group also showed a statistically significant increase in TAS and a decrease in oxidative scores.</td>
<td>This study found that the intervention of Tai Chi exercise in the elderly has antioxidant and hypoglycemic effects. Therefore, it can be recommended to make Tai Chi exercise for the elderly with gout.</td>
</tr>
<tr>
<td>Yunahara Farida and Rifaldi Agustian Firmansyah</td>
<td>2016</td>
<td>Xanthine Oxidase Inhibitory Activity Ethanol and Water Extract from Herbs (Peperomia pellucida L.)</td>
<td>Experimental studies with comparison</td>
<td>messenger plant (Peperomia pellucida L.) is a plant that has activity in inhibiting xanthine oxidase. The test results showed that the thick extract of the ordered herbs in ethanol solvent and water solvent had a total flavonoid content of 4.23% and IC50 40.79 bpj, and the thick extract of ordered herbs in water solvent had a flavonoid content of 1.43% and IC50 43.11 bpj. Ethanol extract and herbal water have an inhibitory effect on xanthine oxidase.</td>
<td>Peperomia pellucida contains a class of flavonoid compounds, alkaloids, saponins, quinones and steroids. The ethanol extract of the messenger herbs had a better xanthine oxidase inhibitory activity (IC50 40.79 bpj) than the aqueous extract of the herbal extracts (IC50 43.11 bpj).</td>
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</tbody>
</table>
Table 1. Journal analysis of 8 main articles included in this study (Cont.)

<table>
<thead>
<tr>
<th>Author</th>
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<th>Result &amp; Discussion</th>
<th>Conclusion</th>
</tr>
</thead>
<tbody>
<tr>
<td>Ema P. Yunita</td>
<td>2018</td>
<td>Relationship Between Obesity, High Purine Consumption, and Treatment of Gout With Allopurinol.</td>
<td>Quasi-experiment with pre-test and post-test design</td>
<td>The study’s findings revealed that individuals who consumed high-purine foods without restriction and took aspirin, although taking the medicine allopurinol, had uncontrolled uric acid levels.</td>
<td>Although the patient took allopurinol, this study found no link between obesity, high purine consumption, and medicine consumption that affects uric acid levels.</td>
</tr>
<tr>
<td>Angelina Ajeng Prihastuti, Sumi Wijaya, Lanny Hartanti</td>
<td>2017</td>
<td>Xanthine Oxidase Inhibitor Activity Test from The Fraction of Peperomia pellucida Ethanol Extract</td>
<td>Experimental</td>
<td>The result showed that ethyl acetate-ethanol fraction potentially inhibited xanthine oxidase with an IC50 value of 5.00 ± 0.06 ppm, while ethanolic extract of Peperomia pellucida (L.) Kunth herb and allopurinol have IC50 values 0.33± 0.07 ppm and 0.84 ± 0.02 ppm, respectively.</td>
<td>The ethyl acetate-ethanol fraction of the ethanol extract of the messenger herb (Peperomia pellucida I) had an inhibitory effect on the xanthine oxidase enzyme with an IC50 value of 5.00 + 0.06 ppm. Thus, the ethyl acetate-ethanol fraction of ordered herbs containing flavonoids has a potency of 5 times less than allopurinol and 15 times less than its extra ethanol.</td>
</tr>
</tbody>
</table>

Based on those who have met the inclusion criteria, an increase in results related to an increase in the range of motion of the elderly, a decrease in pain, flexibility of researchers, the balance of journals that took part in tai chi exercises was obtained. In addition, an ethanolic extract was discovered based on the components in the Peperomia pellucida L. that serves as an inhibitor of xanthine oxidase. This enzyme functions as a catalyst in the oxidation of hypoxanthine to xanthine and then to uric acid.

**DISCUSSION**

**Risk of Gout Arthritis and Hyperuricemia**

Hyperuricemia is caused by high uric acid levels in the body due to excessive uric acid synthesis and decreased excretion of uric acid in the distal tubule of the kidney. Hyperuricemia accompanied by synovial fluid or soft tissue saturation can increase the risk of formation of sharp monosodium urate crystals and trigger inflammation. Monocytes and macrophages will eliminate crystals through phagocytosis accompanied by the release of inflammatory mediators into the surrounding area and trigger inflammation and tissue damage (Muqorrobin, 2017). Diet and the breakdown of ribonucleic acid from cells produce purines as the end product of metabolism. Through the normal pathway, the purines will be metabolized by hypoxanthine and converted to xanthine. Xanthine through the enzyme xanthine oxidase will be overhauled into uric acid. The kidneys will excrete uric acid through urine—a mechanism of inhibition of uric acid formation through xanthine oxidase inhibitors.

**Messenger plant (Peperomia pellucida) in reducing the amount of uric acid**

This plant (Peperomia pellucida I) has been used in traditional medicine for thousands of years to treat various diseases. Chinese betel contains many alkaloids, tannins, saponins, quinine, and flavonoids, which can treat acne, gout, skin inflammation, or ulcers. Herbs have properties as an analgesic and antipyretic, anti-inflammatory, antioxidant, antimicrobial and antitumor, antihyperglycemic, and antihyperuricemia (Farida et al., 2016). In addition, it has a spicy and cool aroma and has anti-fungal properties. Orders can be consumed directly as fresh vegetables, used as a concoction by drinking boiled water or grinding, and then affixed to the affected area.

Based on Mutee (2010) research, the dose that can be used is 1000mg/kg body weight per day orally in male rats with oedema in the hind legs. The results show that herbal messengers can reduce uric acid levels not much different
from allopurinol as a uric acid synthesis drug in mice (Bakar, A et al., 2018). Several studies have shown many pharmacological effects contained in peperomia pellucida, including analgesic, anti-inflammatory, antipyretic, antioxidant, antihyperglycemic, antihyperuricemia, cytotoxic, antimicrobial, cell anisikling, a lipase inhibitor, fibrinolytic, thrombolytic, antiasthmatic, and antosteoporosis. The optimal level of flavonoids in the Chinese extract to inhibit xanthine oxidase activity is 300 g. (Kartika et al., 2016).

Common flavonoid compounds in inhibiting xanthine oxidase are apigenin, luteolin, camphorol, quercetin and myricetin. The content of flavonoids in the herbs has the effect of lowering uric acid levels by inhibiting the activity of the enzyme xanthine oxidase on purine bases so that the production of uric acid can be reduced. The content of flavonoids can inhibit the activity of the xanthine oxidase enzyme through competitive inhibition. Quercetin type flavonoid has a hydroxyl group that is competitive with xanthine (substrate). Therefore, flavonoids (quercetin) can react more with the xanthine oxidase enzyme so that the inhibitory effect of uric acid formation can last as long as the flavonoids are still in the plasma. The optimal level of flavonoids in the extract of the Chinese order to inhibit the activity of xanthine oxidase is 200 gr.

Yunahira (2016) explained that the inhibitory effect of xanthine oxidase inhibitory water extract of 50 mL china was 51.77% with IC50, or the concentration that could inhibit xanthine oxidase enzyme activity 50% of the Chinese extract was 43.11%. Therefore, the IC50 of the alkaloid extract was lower than that of the ethanolic extract, indicating that the alkaloidal extract was more active than the ethanolic extract. This follows the statement that active alkaloid compounds inhibit the xanthine oxidase enzyme (Farida and Firmansyah, 2016). This is supported by the results of research conducted by Imbar et al. (2019), namely the ethanol extract of the host plant can have the effect of reducing uric acid levels with a dose of 200 mg of the ethanolic extract of the messenger plant, which is the optimum dose in lowering uric acid levels. Therefore, the ethanol extract of the messenger plant can be recommended as an alternative medicine to reduce uric acid. The ability of the ethanol extract of the messenger plant to reduce uric acid levels in the blood is thought to be caused by the flavonoid compounds contained in the plant extract (Imbar et al., 2019).

In addition, research by Kartika et al. (2016) showed that the anti-inflammatory effect of the herbal plant was able to inhibit oedema at a dose of 400mg/kg BW close to indomethacin 10mg/kg BW. Oral administration of ordered herbal plant extracts at a dose of 1500-2500mg/kg WB significantly affects inflammation. The antihyperuricemia effect of the suruhan herb showed that the ethanol extract at a dose of 50 mg/kg could reduce uric acid levels by 24.35%, while a dose of 200 mg/kg reduced uric acid levels by 31.52%, and a dose of 100 mg/kg reduced uric acid levels. uric acid 32.20% (Kartika et al., 2016).

**Tai Chi Exercise**

The advantages of Tai Chi on Fitness, Flexibility and Joint Pain

Tai Chi gymnastics consists of functional movements that are carried out daily, such as walking, standing, squatting, sitting to standing, and going up and down stairs which are still not found in fitness gymnastics and other physical exercises that are often done by the elderly such as yoga, swimming, cycling. Relaxing, ergonomic exercise. Research conducted by Lam and Horstman in 2013 showed that Tai Chi exercise interventions with several modifications could improve muscle strength, balance, flexibility or flexibility, and increase pain threshold and improve the overall quality of life (Suci, 2017).

Pharmacological therapy of hyperuricemia has a greater potential to cause side effects than non-pharmacological therapy using herbal plants and physical exercise. In addition, non-pharmacological therapies tend to be in demand because they are relatively cheap and easy to find. China's messenger plants are not widely known for their benefits as a therapy for hyperuricemia. The principle of giving Chinese tea is that individuals are encouraged to fast for 2 hours before drinking the tea. This therapy can be continued until uric acid levels return to normal. Based on several studies, giving this tea can be done for 14 days to achieve optimal results.

The target of this research intervention, especially Tai Chi exercise, is the elderly aged 60 years and over who are advised to do light exercise that is not too burdensome to the bones (Suci, 2015). In Tai Chi gymnastics, there are dynamic active stretching movements, semi squats, closed-chain-kinetic, and core stability exercises from the beginning of the exercise to the end, which is useful for increasing flexibility and balance in the elderly due to changes in muscle morphology, sensory systems and the occurrence of cartilage protein depletion from elderly joints. Semi squat movement with closed-chain-kinetic will activate the quadriceps muscle group, lumbar vertebrae muscles and pelvic floor muscles. Performing semi-squat movements with closed-chain-kinetic will stimulate the quadriceps muscle group, especially the vastus medial oblique and vastus lateral (Kisner, 2014, Komang 2017). Closed-Chain-Kinetic provides a large proprioceptive and kinesthetic stimulus through the resulting joint approximation. Multiple contractions accompanied by weight-bearing elements (axial loading) during the closed-kinetic-chain process cause joint
approximation; this stimulates mechanoreceptors in muscles and receptors around the joint to increase sensory input in the movement control process. Closed-kinetic-chain involves a strengthening component between the agonist and antagonist knee muscles simultaneously related to the physiological movements of the lower limbs; besides that, eccentric muscle work occurs during closed-kinetic-chain administration, resulting in greater tension in the muscles so that it further increases the ability functional (Komang, 2017).

Then the Core Stability movement is the coactivation of the inner muscles of the lower trunk to control weight transfer and stepping during the walking process. Tai Chi has core stability exercises that develop the work of the dynamic muscular corset muscles. Coordinated and concurrent contractions of these muscles will provide calendar rigidity to support the trunk, resulting in reduced intradiscal pressure. In addition, they will reduce the workload of the lumbar muscles so that the tissue is not easily injured; abnormal lumbar muscle tension will be reduced so that there is an improvement in the muscle pump, which results in increased blood circulation in the back muscle tissue (Komang, 2017).

The supply of food and oxygen in muscle tissue becomes better, so that muscle flexibility increases. The activation of the core muscles that function as spinal stabilizing muscles will relax the global muscles that were previously spasming, thereby obtaining good vertebral stability and the spine's position in a state of functional activity. Reduced intradiscal pressure will make it easier to perform functional activities. It can be concluded that the tai chi exercises intervention affects joint pain in the elderly with hyperuricemia, rheumatism and other joint pain (Kisner, 2014, Komang 2017).

Older adults with gout or at risk for hyperuricemia will be given Tai Chi exercise therapy in the morning around 06.00-07.00 WIB (Riska, 2018, Aisyah, 2017, Suci, 2015) and combined with herbal therapy Peperomia pellucida for approximately 15 days or when the elderly uric acid levels are normal. The techniques used in Tai Chi exercises include dynamic active stretching movements, semi squats, closed-chain-kinetic, and core stability exercises that function to decrease muscle tension, reduce joint pain, and increase muscle flexibility, which is carried out for approximately 20 minutes. -30 minutes every day for 15 days (Riska, 2018). The technique to consume tea made from peperomia pellucida drunk tea water boiled with hot water at a temperature of around 100C is as much as one in a day with the dose in a teabag that has been measured (Sagita, 2014). The result expected from the combination of Tai Chi exercise therapy with tea orders is to help the elderly who experience gout to easily accelerate the decrease in uric acid levels and as a modality and complementary therapy to reduce the side effects of pharmacological drugs.

CONCLUSION

From the eight main journals analyzed by the authors, it was concluded that the messenger herbal plant (Peperomia pellucida I), based on several studies in vitro and in vivo, could inhibit xanthine oxidase through a mechanism that occurs due to the flavonoid content in its 1000 mg/kg body weight. In addition, tai chi exercise can be used for muscle strength, balance, flexibility or flexibility, and can lower the pain threshold. It is hoped that the nurse can consider the effectiveness of this combination therapy to be implemented to the elderly with hyperuricemia. We believe that consuming peperomia pellucida and doing tai chi exercise can control the elderly uric acid level. This therapy is an appropriate and complementary approach which is easily carried out by older adults with gout, because this therapy is simple and consists of slow movements with lower joint impacts and loading, safe, and low-cost exercise.

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REFERENCES


