Literatur Review

Anti-Dyslipidemia Effects Of Psidium guajava L.

Amelia¹, Sumihar MR Pasaribu²

- 1. Student of Masters Program in Biomedical Sciences, Faculty of Medicine, Universitas Methodist Indonesia, Medan
- 2. Department of Bio Molecular, Masters Program of Biomedical Sciences, Faculty of Medicine, Universitas Methodist Indonesia, Medan
- * Correspondence: e-mail: <u>ameli-</u> <u>amilala69@gmail.com</u> Phone.: +62821-6944-2433

Backgrounds:Dyslipidemia is a clinical condition characterized by an increase in the plasma concentration of triglycerides and/or total cholesterol or its fraction relative to a reference value that is considered normal. These changes include hypertriglyceridemia due to increased synthesis of very low-density lipoprotein triglyceride (VLDL), reduced hydrolysis and/or hypercholesterolemia, due to accumulation of cholesterol-rich lipoproteins, such as low-density lipoprotein (LDL). One of the treatments for dyslipidemia besides using conventional therapy can also be done with traditional medicine or herbal medicine such as guava (Psidium guajava L.). Psidium guajava L. contains lycopene and 9-oxo-ODA which has potential as an antidyslipidemic.

Abstract

Methods: This paper uses a literature study or literature review of research articles. Search articles through Google Scholar and Pubmed with keywords used Psidium guajava L. and Dyslipidemia or Hypercholesterolemia. The criteria for articles used in the last 5 years are from 2017 to 2021.

Results: Based on the literature search, there were5studies related to the effect of Psidium guajava L. on dyslipidemia in the obesity rat model, where all studies were tested on experimental animals.

Conclusion: Based on the literature study, it was found thatPsidium guajava L.is an alternative in the treatment of dyslipidemia which has a fairly effective effect of lowering blood cholesterol (LDL and Triglycerides) levels.

Keywords: Psidium guajava, Dyslipidemia, Hypercholesterolemia

INTRODUCTION

Excess body fat or obesity is a health problem that often occurs in modern times. This happens because of an imbalance between energy intake and expenditure, resulting in excessive fat accumulation in the body[1] [2].In Indonesia, there are three diseases that cause the most death, namely stroke, coronary heart disease, and diabetes mellitus and their complications. Obesity is one of the risk factors for these diseases. In 2014, more than 1.9 billion (39%) adults aged 18 years and over worldwide were overweight and about 671 million (13%) were obese [1]. More than 50% of the 671 million obese people worldwide are in the following 10 countries based on rankings from one to ten, including: America, China, India, Russia, Brazil, Mexico, Egypt, Germany, Pakistan and Indonesia are ranked 10th in this world[3].According to the 2013 Basic Health Research (RISKESDAS) data, the prevalence of obesity in Indonesia reached 32.9%, where the Special Region of Yogyakarta (DIY) was included in sixteen provinces with obesity prevalence above the national obesity prevalence.[4].

Excess body fat (obesity) is currently an epidemic that appears throughout the world, including in developing countries[5].Excessive fat accumulation in obese patients with dyslipidemia results in an increase in the amount of free fatty acids hydrolyzed by endothelial LPL. This increase triggers the production of oxidants that have a negative effect on the endoplasmic reticulum and mitochondria. Free Fatty Acid (FFA) which is released due to excessive fat accumulation also inhibits lipogenesis, thereby inhibiting serum triacylglycerol clearance, resulting in an increase in blood TG levels and hypertriglyceridemia.[6].

Dyslipidemia conditions can affect changes in the synthesis of Very Low Density Lipoprotein (VLDL) in liver the and disturbances in Lipoprotein Lipase (LPL) resulting in an increase in TG and total cholesterol levels in the body3. In dyslipidemia there is an increase in total cholesterol levels caused by an increase in cholesterol found in VLDL and LDL due to a large increase in circulating TG. resulting in excessive fat accumulation in the body and hypercholesterolemia.[7].

Drugs such as statins, fibrates, nicotinic acid and cholesterol absorption inhibitors are often used for the treatment of dyslipidemia and its complications, but the side effects caused by long-term use of synthetic drugs and the increasing need for drugs encourage people to look for ingredients natural or traditional ingredients (herbal plants) which have low prices and are easy to obtain as an alternative treatment.One of the natural ingredients known to reduce cholesterol levels in guava leaves and fruit (*Psidium guajava* L).

Research conducted by Wurdianing et al. (2014), showed that administration of soursop leaf extract (Annona muricata) in experimental rats induced by a high-fat diet showed changes in lipid profiles which were marked by a decrease in cholesterol levels.[8].

The results of research conducted by Adeneye and Olagunju (2009) proved that the content of saponins in papaya seeds (Carica papaya L) has an effect in lowering total cholesterol by binding to bile acids in the intestine so that the enterohepatic process does not occur.[9].

Research conducted by Diarti et al. (2018) with effect titleMelon seed flour (Cucumis Melo I.) on total cholesterol levels of male white rats (Rattus Norvegicus) wistar strain, that administration of Melon Seed Flour (Cucumis melo L.) for 14 days can reduce total cholesterol levels in male white rats. (Rattus norvegicus) Wistar . strain[10].

Based on the above background, the researchers are interested in proving the effect of guava extract (*Psidium guajava* L.) which has an effect as an antidyslipidemic (hypolipidemic) blood. *Psidium guajava* L. which has the potential to regulate lipid profiles, by reducing blood cholesterol (LDL) and triglyceride levels.

METHODS

1. Study Design

This research is a research using literature study method or literature review. A literature review is a comprehensive overview of the research that has been done on a specific topic to show the reader what is already known about the topic and what is not known, to seek rationale from research that has been done or for further research ideas.

2. Population and Sample

The data used in this study comes from the results of research that has been done (literature studies) and published in national and international online journals. In conducting this research, the researchers searched for research journals published on the internet using the Google Schoolar and PubMed search engines (table 1).

3. Study Variable

The dependent variable in this study was dyslipidemia while the

independent variable was *Psidium guajava* L. extract (*Aloe vera* L).

4. Operational Definition of Variables

Dyslipidemiarefers to unhealthy levels of one or more kinds of lipid (fat) in your blood. Blood contains 3 main types of lipids: highdensity lipoprotein (HDL), lowdensity lipoprotein (LDL) and triglycerides

Psidium guajava L.is a plant thatalso called guava stone, guava siki and guava klutuk is a tropical plant originating from Brazil, distributed toIndonesiathroughThailand. Guava has green fruit with white or red flesh and has a sweet-sour taste. Guava fruit is known to contain a lot ofvitamin C

5. Data analysis

This research uses literature review method design with identification, evaluation, and interpretation of all research results related to certain topics. Methodliterature review, summarizes the results of primary research in a more comprehensive presentation of comprehensive and balanced facts.

6. Research Ethics

There is no ethical clearance in this study because it only uses literature study.

| Criteria | inclusion | Criteria | Exclusion |
|-----------------------------|---|--------------|---|
| Period of time | Publication date for the last 5 years starting from 2017 to 2021 | Article Type | The research method is not de- scriptive because researchers need to identify the effects of herbal plants as hypolipidemic not just a description. |
| Article type | International | Results | Research results that have been published and must have a p value or must be read by statis- tics because researchers need to see whether or not there is a re- lationship effect of <i>Psidium</i> <i>guajava</i> L. as a hypolipidemic or antidyslipidemic |
| Article content theme | Effect of <i>Psidium guajava</i> L. on Hy- percholesterolemia OR dyslipidem- ia. <i>Psidium guajava</i> L. and Hypercho- lesterolemia/dyslipidemia | | |

Table 1. Inclusion and Exclusion Criteria in *Literature Review*

Table 2. Study Description Literature Review

OPEN ACCESS

| Name / Year | Title | Destination | Design | Sample | Results |
|---|--|--|---|--|---|
| Siti Pan- danwangi, Dian Oktavi- an, (2018) | The Effective- ness Test of the Combination of Purple Eggplant Peel Extract (<i>So- lanum melongena</i> L) and Guava Leaves (<i>Psidium</i> <i>guajava</i> L) as Cholesterol Lowering in White Rats (<i>Rat- tus novergicus</i>) | To Deter- mine The Effectiveness And At What Dosage The Combi- nation Ex- tract Suspen- sion Of Pur- ple Eggplant Peel (Sola- num melongena L) And Guava Leaves (Psid- ium guajava) Can Lower Cholesterol Levels In Male White Rats (Rattus Norvegicus) Induced With Propylthiou- racil And Domestic Chicken Egg Yolk. | Research Design Us- ing Experi- mental La- boratory With Pre And Post Test Method With Control Group De- sign | White Rat (Rattus novergicus) | That Suspension Combination of Purple Eggplant Peel Extract Suspension And Guava Leaves Can Lower Cho- lesterol Levels In The Blood. |
| Muhammad Aulia Rah- man, Irfan Hamdani, Isra Thristy, Muhammad Jalaluddin Assuyuthi Chalil (2019) | Comparison of the Effective- ness of Tomato (Lycopersicum Esculantum Mill.) Juice with Red Guava (<i>Psidium guajava</i> L.) Juice on Re- duction of Total Cholesterol | The Research Aims at the Effectiveness of Giving Tomato Fruit Juice With Red Guava Juice To De- crease Total Cholesterol In White | This Study Was Experi- mental Using Pretest Post- test With Control Group De- sign | White Rat (Rattus Norvegicus L.) Male 24 Wistar Strain Induced Egg Yolk 6.25gr / Kgbw, Grouped Into 4, Nega- tive Controls | Giving Tomato Fruit Juice And Guava Fruit Juice For 2 Weeks Can Lower Total Cholesterol Lev- els In The Blood Of Male White Rats Of Wistar Strain Previous- |

| | Levels In Male White Rats In- duced by Egg Yolk | Rats Induced By Egg Yolk. | | Were Given Distilled Wa- ter, Positive Control Was Given Egg Yolk, One Treatment Was Given Tomato Juice 3 Cc And Two Treat- ment Was Adminis- tered 3 Cc Red Guava Fruit Juice For 2 Weeks Then Blood Was Taking For 3 Times, Pretest, In- tervention And Posttest. | ly Induced By Egg Yolk. And Also There Is A Significant Dif- ference In Giv- ing Tomato Fruit Juice And Guava Juice To Reduction In Total Cholester- ol Levels Of Wistar Male White Rats In- duced Egg Yolk. Giving guava fruit juice has a higher level of effectiveness in reducing total cholesterol lev- els in male white rats of the Wistar strain compared to |
|--|---|---|---|---|---|
| Maria Evane Navy Caha- ya Putri, Nita Pranita- sari (2018) | Effect of Ad- ministration of Guava Extract (<i>Psidium guaja-</i> <i>va</i>) on Blood Triglyceride Levels of Male White Rats (Rat- tus Norvegicus) Wistar Strain Induced by Dexamethasone | To Investi- gate Guava (<i>Psidium</i> guajava) Con- tains Flavo- noid Com- ponents (Quercetin) Which Has Inhibitory Activity Against En- zymes In- volved In Triglyceride Synthesis So | Samples of Experimental Animals Used A total of 24 animals were divided into 3 groups | The experi- mental ani- mal samples used were 24 animals di- vided into 3 groups: ex- perimental animal group fed standard feed, exper- imental ani- mal group induced with dexame- thasone at a | giving tomato fruit juice Administration of Guava Extract (<i>Psidium guaja-</i> <i>va</i>) Significantly Reduced Blood Triglyceride Levels of Exper- imental Animals Induced by Dexamethasone Because Guavas Contain Flavo- noid Compo- nents (Querce- tin). |

| | | It Can Lower | | dose of 0.13 | |
|-------------|-----------------|---------------|--------------|----------------|----------------------------------|
| | | Blood Tri- | | mg/kg for 14 | |
| | | glyceride | | days starting | |
| | | Levels. | | on day 8, and | |
| | | | | experimental | |
| | | | | group in- | |
| | | | | duced Dex- | |
| | | | | amethasone | |
| | | | | with a dose | |
| | | | | of 0.13 mg/kg | |
| | | | | for 14 days | |
| | | | | starting on | |
| | | | | the 8th day | |
| | | | | and given | |
| | | | | guava extract | |
| | | | | with a dose | |
| | | | | of 3 gr/kg for | |
| | | | | 21 days. On | |
| | | | | Day 22, | |
| | | | | Blood Tri- | |
| | | | | glyceride | |
| | | | | Levels were | |
| | | | | Checked Us- | |
| | | | | ing the Glyc- | |
| | | | | erol Blanking | |
| | | | | Method. | |
| | | | | inculou. | Guava Leaf |
| Md. Abdul- | | | | | Powder Sup- |
| lah Al | | To Study The | | | plementation |
| Mamun, Md. | High Carbohy- | Effect Of | | | Prevents Obesi- |
| Faruk, Md. | drate High Fat | Psidium | | | ty, Improves |
| Mizanur | Diet Induced | guajava Leaf | Experimental | | Glucose Intoler- |
| | Hepatic Steato- | Powder | rats with a | | |
| Rahman, | sis And | | total of 28 | Experimental | ance, And Re- duces Inflamma- |
| Kamrun Na- | Dyslipidemia | Supplemen- | tails were | Design | |
| har, Fariha | Were Amelio- | tation On | divided into | | tion And Oxida- |
| Kabir, Md | rated By Psidi- | Obesity And | 4 groups | | tive Stress In |
| Ashraful | um guajava | Liver Status | _ | | The Liver Of |
| Alam, And | Leaf Powder | Using Exper- | | | Rats Given A |
| Nusrat Sub- | Supplementa- | imental Rats. | | | High Carbohy- |
| han | tion In Rats | | | | drate High Fat |
| | | | | | Diet. |

| Brito, A., | | | | | |
|--------------|--|---------------|--|-----------------------------------|--|
| Lima, GM, | | Assessing | | | |
| Farias, LM, | | Effects Of 28 | | | |
| | Lycopene-Rich Extract From Red Guava (<i>Psidium guaja-</i> <i>va</i> L.) Decreases Plasma Triglyc- erides And Im- proves Oxida- tive Stress Bi- omarkers On Experimentally- Induced Dyslipidemia | e | Male Ham- sters (116.5 \pm 2.16 G) were given 93g Ain Feed Containing Casein (20%), Coco- nut Fat (13.5%) and Cholesterol (0.1%). | Experimental Model De- sign | The Lycopene- Rich Extract From Red Gua- va Fruit (<i>Psidium</i> <i>guajava</i> L.) Pro- moted Hypotri- glyceridemic Effect Only At 25 Mg/Kg In An Experimental Model Of Dyslipidemia In Hamsters. |
| Arcanjo, D., | In Hamsters | Dyslipidem- | | | |
| & Martins, | | ia | | | |
| M (2019) | | | | | |

RESULT

Based on the literature search, there were 5 studies related to the effect of *Psidium guajava* L. on lipid profile levels, of which there were 5 studies on experimental animals (Table2).

DISCUSSION

Based on the literature search, it was found that there were 5 studies related to the effect of *Psidium guajava* L. extract on blood cholesterol, where all the experimental studies on experimental animals varied the dosage of *Psidium guajava* L. extract used. Each test obtained from this literature study has a different dose and duration of research - different from one another.

Research conducted by Pandanwangi and Oktaviani (2018), aims toThe purpose of this study was to determine the effectiveness and at what dose the combination of purple (Solanum eggplant peel extract melongena leaves L) and guava (Psidium guajava) can reduce cholesterol levels in male white rats (Rattus norvegicus) induced with propylthiouracil and domestic chicken egg yolk. The results showed that the results of the study of reducing

cholesterol levels in male white rats showed that the combination suspension of purple eggplant skin extract (Solanum melongena L) and guava leaves (Psidium guajava) was the most effective for reducing cholesterol levels in male white rats (Rattus norvegicus). with dose а of 50mg/200gBW/day: 40mg/200gBW/day[11].

Research conducted by Rahman et al. (2019), regarding the comparison of the effectiveness of giving tomato juice with red guava fruit juice on reducing total cholesterol in white rats induced by egg yolk. The results of this study indicate that guava fruit juice has a higher level of effectiveness in reducing total cholesterol levels in Wistar male white rats compared to giving tomato juice.[12].

Research conducted by Putri and Pranitasari (2018), regardingextract effect Guava (Psidium guajava) contains а flavonoid component (quercetin) which has inhibitory activity against enzymes involved in triglyceride synthesis so that it can reduce blood triglyceride levels. This study showed that blood triglyceride levels in the treatment group induced by dexamethasone and given guava extract (Psidium guajava) decreased not significantly compared blood to triglyceride levels in the positive

control group induced by dexamethasone.[13].

In the research of Mamun et al. (2019), Psidium guajava leaves are reported to contain many bioactive polyphenols that play an important role in the prevention and treatment of various diseases. The aim of this study was to examine the effect of Psidium guajava leaf powder supplementation on obesity and liver status using experimental rats. The results showed that guava leaf powder supplementation showed a significant reduction in fat accumulation in obese rats. In addition, liver enzyme function was significantly increased in mice fed a high-fat diet compared to control mice which was further improved by guava leaf powder supplementation in mice fed a high-fat diet. Administration of a highfat diet also decreased the function of antioxidant enzymes and increased lipid peroxidation products compared to control rats. Supplementation of guava leaf powder in rats fed a highfat diet reduced markers of oxidative stress and reestablished antioxidant enzyme systems in experimental animals. Guava leaf powder supplementation in rats fed a high-fat diet also showed a relative decrease in inflammatory cell infiltration and collagen deposition in the liver compared to rats fed a high-fat diet. This study

showed that guava leaf powder supplementation prevented obesity, improved glucose intolerance, and reduced inflammation and oxidative stress in the liver of rats fed a highcarbohydrate, high-fat diet. Supplementation of guava leaf powder in rats fed a high-fat diet reduced markers of oxidative stress and reestablished antioxidant enzyme systems in experimental animals. Guava leaf powder supplementation in rats fed a high-fat diet also showed a relative decrease in inflammatory cell infiltration and collagen deposition in the liver compared to rats fed a high-fat diet. This study showed that guava leaf powder supplementation prevented obesity, improved glucose intolerance, and reduced inflammation and oxidative stress in the liver of rats fed a highcarbohydrate, high-fat diet. Supplementation of guava leaf powder in rats fed a high-fat diet reduced markers of oxidative stress and reestablished antioxidant enzyme systems in experimental animals. Guava leaf powder supplementation in rats fed a high-fat diet also showed a relative decrease in inflammatory cell infiltration and collagen deposition in the liver compared to rats fed a high-fat diet. This study showed that guava leaf powder supplementation prevented obesity, improved glucose intolerance, and reduced inflammation and oxidative stress in the liver of rats fed a highcarbohydrate, high-fat diet. Guava leaf powder supplementation in rats fed a high-fat diet also showed a relative decrease in inflammatory cell infiltration and collagen deposition in the liver compared to rats fed a high-fat diet. This study showed that guava leaf powder supplementation prevented obesity, improved glucose intolerance, and reduced inflammation and oxidative stress in the liver of rats fed a high-carbohydrate, high-fat diet. Guava leaf powder supplementation in rats fed a high-fat diet also showed a relative decrease in inflammatory cell infiltration and collagen deposition in the liver compared to rats fed a highfat diet. This study showed that guava leaf powder supplementation prevented obesity, improved glucose intolerance, and reduced inflammation and oxidative stress in the liver of rats fed a high-carbohydrate, high-fat diet.[14].

The results of research by Brito et al. (2019), on lycopene-rich extract (LRE) from red guava fruit (*Psidium guajava* L.) on lipid profile and oxidative stress in an experimental model of dyslipidemia. Lycopene-rich extract from red guava fruit (*Psidium guajava* L.) promoted a hypotriglyceridemic effect at only 25 mg/kg in an experimental model of dyslipidemia in hamsters. In addition, both doses of 25 and 50 mg/kg decreased plasma levels of lipid peroxidation biomarkers, as evidenced by decreased plasma concentrations of Malondialdehyde (MDA) and Myeloperoxidase (MPO).[15].

The content of red guava fruit is vitamin C and beta carotene. So that red guava fruit can increase endurance because of the antioxidant content in it. Red guava fruit also contains fiber that contains pectin, which makes it hypocholesterolemic and hypoglycemic. This can prevent blockages in blood vessels because it can lower cholesterol levels in the blood [16].

The content of red guava fruit is vitamin C and beta carotene. So that red guava fruit can increase endurance because of the antioxidant content in it. Red guava fruit (Psidium guajava L) also contains fiber that contains pectin, which makes it hypocholesterolemic and hypoglycemic. This can prevent blockages in blood vessels because it can lower cholesterol levels in the blood.Giving red guava fruit juice (Psidium guajava L.) can reduce total cholesterol levels in rats because both fruits contain high antioxidants such as lycopene. Lycopene inhibits the action of the HMG-CoA reductase enzyme which plays a role in

cholesterol synthesis in the liver so that it has a hypocholesterolemic effect. activate LDL receptors, and can increase LDL degradation. In addition, red guava juice contains 9-oxo-ODA is an agonist of peroxisome Proliferator-Activated Receptor (PPARa). PPARa is a receptor that functions in fat oxidation. When this receptor is activated, fatty acid oxidation will occur in the tissue so that it will reduce the accumulation of triglycerides in the tissue. This receptor will also induce the expression of lipoprotein lipase which will increase lipolysis of lipoprotein so that it will reduce LDL levels and total cholesterol levels in plasma. [17].

CONCLUSION

There is an effect of guava extract (*Psidium guajava* L.) on blood cholesterol levels in experimental animal models of hypercholesterolemia

CONFLICT OF INTEREST

The author declares that there is no conflict of interest in this study.

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