Effects of Pineapple Juice and Ginger Drink for Relieving Primary Dysmenorrhea Pain among Adolescents

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ABSTRACT

Background: Primary dysmenorrhea is characterized by low abdominal pain occurring during menstruation. It is negatively impact on all aspects of life of adolescents such as increased school absenteeism, emotional disorders, impaired social interaction and productivity. The herbal drink which they can self-prepared is common choice as home remedies to relieve the pain. This study aimed to examine the effect of pineapple juice and ginger drink in providing pain relief to the adolescent with primary dysmenorrhea.

Subjects and Method: This was a randomized trial study conducted at three high schools in Surakarta, Central Java, Indonesia, from January to April 2018. A sample of 39 female students aged 15-17 years was selected for this study, divided into three groups. The treatment was given for 3 days from first day of menstruation for 3 menstruation cycles. The pain score were measured by numeric rating scale (NRS). The data were analyzed by Mann-Whitney.

Results: Pain score in pineapple juice group (mean= 2.25; SD= 1.79) and ginger drink group (mean= 3.63; SD= 2.02) were lower than control group (mean= 5.64; SD= 1.25) and it was statistically significant.

Conclusion: Pineapple juice and ginger drink effective to reduce pain of primary dysmenorrhea among female adolescent.

Keywords: Dysmenorrhea, pain, pineapple juice, ginger drink, female adolescents

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BACKGROUND

Adolescence is the transitional phase between childhood and adulthood and characterized by physical growth, psychological changes, and secondary sexual changes. The most striking change is the onset of menstruation (Santoro and Neal-Perry, 2010). After menarche, the most common gynaecological disorders that may encounter is dysmenorrhea (Jenabi, 2013). Dysmenorrhea is a common gynaecological disorder with painful cramps of uterin origin. The term dysmenorrhea is derived from the Greek words dys (difficult, pain, abnormal), mena (month), and reha (flow) (Madhubala and Jyoti, 2012). It can be either primary (absence of an identifiable pathological condition) or secondary (due to pathology condition such as endometriosis). Primary dysmenorrhea is highly prevalent that estimated occurs in 45-90% of productive women (Bernardi et al., 2017). It usually begins before or at the start of menstruation, and continue for 2-3 days (Wallace et al., 2010). The pain felt mainly in the lower abdomen, but it may radiate to the back and along the thighs. In a severe case, it usually accompanied by some other symptoms such as nausea, vomiting, diarrhea, headache, fatigue, dizziness, decreased concentration, loss of appetite, insomnia, to loss of consciousness (Fatima et al., 2016). The consequences not only affect reduces productivity such as school or works absen-
teeism, but also effects of quality of life because it is decreased academic performance, lead depressed mood, lethargy, and disruption of social (Al-Asadi and Abdul Quadir, 2013).

Primary dysmenorrhea well-recognized by increasing of prostaglandin (PGs). Prostaglandins have a role inducing uterine contraction, decreasing uterine blood flow, increasing peripheral nerve hypersensitivity resulting in pain and cramps. Treatment for primary dysmenorrhea could be a self-medication such as non-pharmacological home remedies (warm beverages, heat application, back massage, low-fat diet, lifestyle modification, and herbal treatment), taking over-the-counter drugs or seeking medical advice such as take prescription analgesics and hormonal contraceptives. Non-steroidal anti-inflammation drugs (NSAIDs) and antispasmodics are commonly used drugs. NSAIDs such as ibuprofen, mefenamic acid, naproxen, ketoprofen, celecoxib, and diclofenac are proven to be effective by inhibiting cyclooxygenase leading to the reduction of prostaglandins secretion. Hormonal therapy in the form of oral contraceptive usually used for patients with lack of response to NSAIDs. Despite it is considerable, but the failure rate is still high, the pain relief inadequate for some women, and the side effects not be well tolerated (Ramya et al., 2014). Therefore, over the last 15 to 20 years, there has been an increase in the use of herbal medicine which they can self-prepared as home remedies (Wong et al., 2009).

The herbal products used for the treatment of primary dysmenorrhea throughout the world vary. In Indonesia, ginger root (Zingibers officinale) is well-known herbal products common used as traditional medicine, spices and drink to warm the body. Study by Chen et al. (2016) found that ginger has been proven as anti-inflammatory agent and has therapeutic effect to relieves the pain of dysmenorrhea. This benefits comes from gingerol and 6-shogaol as active component of ginger. Gingerol works to inhibit the formation of cyclooxygenase (COX) enzymes and prostaglandin synthesis. This mechanism is similar with action of non-steroidal anti-inflammation drugs (NSAIDs) such as ibuprofen and naproxen which also work by inhibits the activity of COX-1 and COX-2 enzymes (Ozgoli et al., 2009).

Besides ginger, pineapple fruit (Ananas comosus (L.)) is also well-known fruit used as a meat tenderizer and traditional remedy for several ailments related inflammation. Therapeutic benefits from pineapple fruits are attributed to bromelain. Bromelain is an extract obtained from pineapple that belongs to a group of protein-digesting enzymes and a protease inhibitor. A wide range of therapeutic benefit have been claimed for bromelain, such as accelerates post-traumatic wound healing (Hirche et al., 2017), relieves joint pain due to arthritis and rheumatic (Jayachandran and Kobre, 2017), immune-modulator with anti-tumor and anti-cancer activity (Mohammed et al., 2017), anti-plaque and anti-gingivitis (Tadikonda et al., 2017), antioxidant (Rodrigues et al., 2017), anti-ulcer (Sowjanya et al., 2016), anti-biotics in cases of pneumonia, bronchitis, and sinusitis (Dighe et al., 2010) and potentially as pain reliever and analgesic in several inflammatory reactions (Singh et al., 2016). The anti-inflammatory and analgesic activity of bromelain are proven to be effective leading reduction of prostaglandin production by inhibits cyclooxygenase pathway (Conteras et al., 2008). With this mechanism, bromelain could be considered as potential phytomedical or herbal remedies for relieves the pain of primary dysmenorrhea. This present study was to
assess the effectiveness of pineapple juice and ginger drink in providing pain relief to the adolescent with primary dysmenorrhea.

SUBJECTS AND METHOD

1. Study Design
This was a randomized control trial conducted in Senior High Schools in Surakarta, Central Java.

2. Study Sample
A sample of 39 female students aged 15-17 years was selected for this study by simple random sampling, divided into three groups. The treatment was given for 3 days from first day of menstruation for 3 menstruation cycles. The exclusion criteria included students who had mild dysmenorrhea, taking either home remedies or oral analgesics for relieves the pain, smoking, drink alcohol, have a history gynaecological disorders or under medical treatment.

3. Study Variables
The dependent variable was pain of dysmenorrhea. The independent variables were the administration of pineapple juice and ginger drink.

4. Study Instruments
Pain was measured by Numeric Rating Scale (NRS). Data about socio-demographics and menstrual pattern were collected by questionnaire. BMI was calculated using height (meter) / weight² (kg).

The pineapple juice and ginger drink was prepared freshly. The pineapple juice made by certain dose, 3.75g/kgBW of fresh pineapple fruits (Ananas comosus (L.) var. Cayenne, 10 grams of sugar, and 200ml water then it blends until smooth, and the ginger drink made by boiling 15 grams of ginger (Zingibers officinale) var. amarum and 10 grams palm sugar with 500 ml water during 15 minutes until the water volume become 200 ml.

5. Data Analysis
The socio-demographic and menstrual characteristic described with descriptive analysis. Test the normality of data using Saphiro-Wilk. Differences in mean pain score before and after intervention in each group using Wilcoxon test. Mann-Whitney (Post-hoc) test was used to determine the difference in mean pain score between groups and see which treatment groups were different.

6. Research Ethics
This study has been approved by the Research Ethics Committee of Dr. Moejadi Hospital and Faculty of Medicine Universitas Sebelas Maret Surakarta with ethical number 1.160/XII/HREC/2017. We obtained written informed consent from the study subjects.

RESULTS
The study involved a total of 39 subjects. Two subjects from pineapple juice group discontinued the trial due to had their menstruation when they went to Bali to study tour, two subjects from ginger drink group discontinued due to took oral analgesics for relieved their pain, and two subjects from control group discontinued the trial because rejected to continued this research project any longer, so the subjects analyzed were 33 subjects. No subjects in the trial had any complication after had been treated.

The mean age of the subjects was 15.67±0.65 years. Most of them (57.6%) had a normal nutritional status, and 42.4% had malnutrition with BMI mean was 20.57±0.74 kg/m². Most of them (90.9%) hailed from the upper middle class of socio-economic status. The study subject had attained menarche ranging from 9-15 years with mean 12.18±0.26 years. Most of them (93.9%) had a family history of dysmenorrhea. Around 25 subjects (75.8%) had
the length of the cycle of 25-32 days and 8 subjects (24.2%) had irregular cycles. Majority of the subjects (90.9%), had the menstrual flow of 4-7 days duration. About half of the subject (51.5%) suffered a moderate dysmenorrhea, and the other half suffered a severe dysmenorrhea. There were no significant differences between the two groups concerning baseline characteristics including the socio-demographic or menstrual pattern (Table 1).

Table 1. Socio-Demographic and Menstrual Characteristics Subjects

| Characteristics                        | Pineapple juice group | Ginger drink group | Control group | Total |
|----------------------------------------|-----------------------|-------------------|---------------|-------|
|                                        | n  | %  | n  | %  | n  | %  | n  | %  |     |
| Age                                    |    |    |    |    |     |     |     |     |     |
| 15 years                               | 4  | 36.4 | 5  | 45.5 | 5  | 45.5 | 14 | 42.4 |     |
| 16 years                               | 6  | 54.5 | 5  | 45.5 | 5  | 45.5 | 16 | 48.5 |     |
| 17 years                               | 1  | 9.1  | 1  | 9.1  | 1  | 9.1  | 3  | 9.1  |     |
| Grade of school                        |    |    |    |    |     |     |     |     |     |
| X                                      | 4  | 36.4 | 5  | 45.5 | 5  | 45.5 | 14 | 42.4 |     |
| XI                                     | 7  | 63.6 | 6  | 54.5 | 6  | 54.5 | 19 | 57.6 |     |
| BMI (kg/m²)                            |    |    |    |    |     |     |     |     |     |
| Underweight                            | 3  | 27.3 | 3  | 27.3 | 3  | 27.3 | 9  | 27.3 |     |
| Normal                                 | 6  | 54.5 | 7  | 63.6 | 6  | 54.5 | 19 | 57.6 |     |
| Overweight                             | 2  | 18.2 | 1  | 9.1  | 2  | 18.2 | 5  | 15.1 |     |
| Family income                          |    |    |    |    |     |     |     |     |     |
| High                                   | 10 | 81.8 | 9  | 81.8 | 11 | 100  | 30 | 90.9 |     |
| Low                                    | 1  | 18.2 | 2  | 18.2 | 0  | 0    | 3  | 9.1  |     |
| Age at Menarche                        |    |    |    |    |     |     |     |     |     |
| <12 years                               | 5  | 45.5 | 5  | 45.5 | 5  | 45.5 | 15 | 45.5 |     |
| 12 - 14 years                          | 6  | 54.5 | 5  | 45.5 | 5  | 45.5 | 16 | 48.5 |     |
| >14 years                              | -  | -    | 1  | 9.0  | 1  | 9.0  | 2  | 6.0  |     |
| Family history of dysmenorrhea        |    |    |    |    |     |     |     |     |     |
| Yes                                    | 11 | 100  | 10 | 90.9 | 10 | 90.9 | 31 | 93.9 |     |
| No                                     | -  | -    | 1  | 9.1  | 1  | 9.1  | 2  | 6.1  |     |
| Menstrual cycle duration               |    |    |    |    |     |     |     |     |     |
| <25 days                               | 2  | 18.2 | 3  | 23.7 | 2  | 18.2 | 7  | 21.2 |     |
| 25-32 days                             | 9  | 81.8 | 7  | 63.6 | 9  | 81.8 | 25 | 78.8 |     |
| >32 days                               | -  | -    | 1  | 9.1  | -  | -    | 1  | 3.0  |     |
| Menstrual bleeding duration            |    |    |    |    |     |     |     |     |     |
| 4-7 days                               | 10 | 90.9 | 10 | 90.9 | 10 | 90.9 | 30 | 90.9 |     |
| >7 days                                | 1  | 9.1  | 1  | 9.1  | 1  | 9.1  | 3  | 9.1  |     |
| Severity of Dysmenorrhea               |    |    |    |    |     |     |     |     |     |
| Moderate Pain                          | 5  | 45.5 | 6  | 45.5 | 6  | 45.5 | 17 | 51.5 |     |
| Severe Pain                            | 6  | 54.5 | 5  | 45.5 | 5  | 45.5 | 16 | 48.5 |     |

Table 2 shows that the greatest decreased of before-after treatment score was in the pineapple juice group. The value of \( p<0.001 \) and \( p=0.008 \) (\( p<0.05 \)) means that there were a significant difference in pain score in the three groups, but in the control group, the pain score after treatment was higher than before treatment score. It can be concluded that in the control group, the pain score was increased.

Post hoc test showed that there were significant differences in pain score between pineapple juice group with control group (\( p<0.001 \)), between ginger drink group and control group (\( p=0.001 \)), and between pineapple juice group with ginger drink group (\( p<0.008 \)). This means admi-
administration of both pineapple juice and ginger drink are effective to providing pain relief, but pineapple juice was found had a greater effect to relieving pain of primary dysmenorrhea among adolescent (Table 2).

**Table 2. Pain Scores Between Groups**

| Type of treatment      | Before treatment (Mean±SD) | After treatment (Mean±SD) | Δ After Before (Mean±SD) | p<sup>a</sup> Post hoc | Δ After Treatment between groups (Mean±SD) | p<sup>b</sup> |
|------------------------|---------------------------|---------------------------|-------------------------|-------------------------|--------------------------------------------|-------------|
| Pineapple juice (n=11) | 5.24±1.95                 | 2.25±1.79                 | 2.99±1.87               | <0.001                  | Pineapple juice vs control                 | 3.39±0.54   | <0.001 |
| Ginger drink (n=11)    | 5.55±1.68                 | 3.63±2.02                 | 1.92±1.85               | <0.001                  | Ginger drink vs control                    | 2.01±0.77   | <0.001 |
| Control (n=11)         | 5.44±1.07                 | 5.64±1.25                 | 0.2±1.16                | 0.008                   | Pineapple juice vs ginger drink            | 1.38±0.23   | <0.001 |

Notes:

<sup>a</sup>p-value for Wilcoxon’s test

<sup>b</sup>p-value for Mann-Whitney’s (post hoc) test

**DISCUSSION**

The administration of pineapple juice and ginger drink in the 3 days after the onset of menstrual during 3 menstruation cycle can decrease the pain score of dysmenorrhea among adolescents. The dependent analysis showed that pineapple juice and ginger drink could decrease the pain score for about 2.99±1.87, 1.92±1.85, respectively and statistically significant (p=0.001). The result of the present study supported by the study by Hidayat (2010) which conducted in wistar mice that found bromelain on pineapple juice has 56.82% anti-inflammatory effect and 58.90% analgesic effects. Another study conducted a study in vivo and in vitro to endometriosis cases related to dysmenorrhea and pelvic pain. This study concluded that bromelain was an effective treatment for endometriosis that may have potential therapeutic uses in the prevention and treatment of the patient (Agostinis et al., 2014).

As long as author’s findings, there was no specific research that assesses the potential of bromelain in pineapple juice for relieving pain of primary dysmenorrhea, but study about bromelain commonly concluded have potential therapeutic effects in either acute or chronic inflammatory disease (Muller et al., 2013; Bolten et al., 2015; Singh et al., 2016; Rathnavele et al., 2016; Zhou et al., 2017; Sahbaz et al., 2015). The effect of pineapple juice on relieving the pain of dysmenorrhea may be contribute to bromelain content on pineapple fruit. Onken et al. (2008) showed that bromelain contain on pineapple fruits has an anti-inflammatory effect and analgesic activity because acts as inhibitor that against Phospholipase A<sub>2</sub> (Pla2) activity. Pla2 has been identified contribute to the initiation of inflammation which subsequently induces the production of arachidonic acid (Onken et al., 2008). Arachidonic acid is metabolized from either the cyclooxygenase or lipoxygenase enzymatic pathway for the production of prostaglandins and leukotrienes, which are primarily in the inflammatory pathway (Tap, 2018).
Muhammad and Ahmad (2017) showed that bromelain acts lowering plasmakinin (bradykinin) at inflammatory site, lowering prostaglandin E2 (PGE2) levels, increases platelet cyclic adenosine monophosphate (cAMP) levels thus increasing prostaglandin I2 (PGI2) which acts as vasodilator at inflammation site. However, bromelain inhibits the production of prostaglandin F2α (PGF2α) and Thromboxane A2 (TXA2) by inhibits cyclooxygenase-2 (COX-2) pathway (Bhui et al., 2009). Bromelain was considerable to activated immune system due to it leads to the formation of inflammation mediator such as (IL)-1β, IL-6, interferon (INF)-γ to accelerated homeostasis and control the inflammation (Pavan et al., 2012).

Consumption of ginger drink can decreased the pain score of primary dysmenorrhea (Latha and Venketesan, 2016; Leelavathi and Hemavaty, 2015; Rahnama et al., 2012). Ginger has been proven as an herbal drug for treatment of dysmenorrhea due to have anti-inflammatory properties and acts as COX-2 inhibitors similar to NSAIDs such as mfenamic acid (Shirvani et al., 2014). To explain the effects of ginger drink on reducing pain score of dysmenorrhea, it has been reported that ginger root contains an active compound such as gingerol and 6-shogaol which considered as anti-inflammatory and analgesics agent (Ali et al., 2008). Pre-clinical research explains that gingerol and 6-shogaol inhibits the metabolism of arachidonic acid and prostaglandin through the COX-2 pathway. In addition, gingerol and 6-shogaol also inhibits lipopolysaccharide (LPS) which induces PGE2 (Dugasani et al., 2010) and NO synthesis (Jung et al., 2009). Genetically, gingerol and 6-shogaol also has the ability to inhibit the coding of genes that induces pro-inflammatory cytokines synthesis and secreted in the inflammatory area (Pan et al., 2008).

Pineapple juice and ginger drink also contains ascorbic acid which acts as anti-oxidant and anti-inflammatory agent. It has been widely studied that ascorbic acid was a potent free radical scavenger, and protecting cells against oxidative damage caused by ROS. It is closely associated because in several chronic and acute inflammation and oxidative stress would lead to its depletion and worse the inflammation (Sorice et al., 2014). However, the inhibition of prostaglandin levels during menstruation was the key mechanism for pineapple juice and ginger drink’s effect on pain of dysmenorrhea in this study.

Limitation of this study was did not analyze the intake of other nutrient which probably affect the decreased of dysmenorrhea pain. Moreover, the pain was a personal, subjective experience, influenced by learning, and other psychological variables, it could be biased in this study. Authors declared that there is no conflict of interest regarding of this article.

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