- 2 Aromatherapy massage versus reflexology on female elderlies with the acute coronary syndrome
- 3 4

5 Abstract

Aim: The present study compared the effects of aromatherapy massage and reflexology on fatigue and
 cardiovascular parameters in older female patients with acute coronary syndrome.

Background: Fatigue and abnormalities in cardiovascular parameters are recognized as major problems
for patients with acute coronary syndrome. Non-pharmacological nursing interventions are useful for
controlling this fatigue and reducing patients' suffering during hospitalization.

11 *Design:* A randomized clinical trial.

Method: The study was conducted with 135 older female patients with acute coronary syndrome who were hospitalized on a cardiac care unit in 2014. They were invited to participate in the study and then were randomly divided into three groups: 'aromatherapy massage', 'reflexology', and 'control'. The fatigue severity and cardiovascular parameters were assessed through the Rhoten fatigue scale and a checklist. Measurements in the groups were performed before and immediately after the intervention.

17 Data analysis was performed using descriptive and analytical statistics via the SPSS software.

18 **Results:** Aromatherapy massage significantly decreased fatigue, systolic blood pressure, mean arterial 19 pressure and O_2 saturation more than the reflexology intervention. However, reflexology reduced 20 patients' heart rates more than an aromatherapy massage (P < 0.05). Moreover, no significant changes 21 were observed in patients' diastolic blood pressures when compared to the control group (P = 0.37).

Conclusions: Implementation of both aromatherapy massage and reflexology has positive effects on the
 fatigue and cardiovascular parameters of patients with acute coronary syndrome. However, aromatherapy
 massage can be more beneficial to apply as a supportive approach in coronary diseases.

25

Relevance to clinical practice: The need for reducing fatigue in ACS patients on a cardiac care unit is
evident. The implementation of aromatherapy massage and reflexology had positive effects on patients'
fatigue as related to both physical and mental health.

29

Key words: acute coronary syndrome, aromatherapy, alternative therapies, cardiovascular parameters,
 fatigue, reflexology,

- 32
- 33
- 34 INTRODUCTION

Fatigue is one of the most common and distressing symptoms in patients with acute coronary syndrome (ACS), often impeding patients' active lifestyle (Pelletier et al., 2015; Shlomai et al., 2015). The progression of fatigue may negatively affect the ACS and exaggerate this disease (Franssen et al., 2003; Eckhardt et al., 2014). Moreover, fatigue alters patients' autonomic nervous activities and may lead to abnormalities in cardiovascular parameters (Silverman et al., 2010; Fagundes et al., 2011). In addition, fatigue is a poorly understood phenomena in healthcare settings that often requires a multidisciplinary approach for management (Ozdemir et al., 2013).

8 Despite the use of various medications for symptom relief in patients with ACS, pharmacologic 9 modalities may not be able to comprehensively address these symptoms, including fatigue 10 (Peuckmann-Post et al., 2010; Fake et al., 2016). Complementary and alternative medicine (CAM) 11 services may be the most powerful therapeutic interventions in patients with cardiovascular diseases (Yeh 12 et al., 2006).

Aromatherapy is a holistic intervention applied through inhalation or massage (Bikmoradiet al., 2015; Karadag et al., 2015; Seyyed-Rasooli et al., 2016). Aromatherapy massage consists of the use of essential oil, extracted from the vapor of the volatile essence of plants, through massage. Lavender (Lavandula angustifolia) is widely used in different contexts (Field et al., 2007; Hosseini et al., 2016). Previous studies have confirmed the sedative effects of Lavender on the parasympathetic system. Also, it has been found that lavender promotes the heart function and coronary blood flow (Shiina et al., 2008; Lytle et al., 2014; Bikmoradi et al., 2015). Tisserand (1988) suggested lavender may have a similar function to the

- benzodiazepines, and can promote the effects of gamma-amino-butyric acid in the amygdala (Tisserand.,
 1988).
- Several studies reported that lavender resulted in reduced fatigue in different groups (Kang and Kim.,
 2002; Bahraini et al., 2011). For instance, in randomized controlled studies, Balouchi et al. (2016) and
 Chen et al. (2015) found that aromatherapy with lavender significantly decreased fatigue (Balouchi et al.,
 2016; Chen and Chen 2015). Tahmasebi et al. (2014) in a quasi-experimental clinical trial also stated that
- 26 lavender alleviated cardiovascular parameters in cardiovascular patients (Tahmasebi., 2015).

27 Another complementary therapy approach with a potential beneficial effect for reducing fatigue is 28 reflexology (Özdemiret al., 2013). Reflexology is an ancient practice using the thumb and fingers on 29 hands, feet and ears via stimulating some areas called reflex zones. A reflex zone corresponds with an organ, gland or body parts and is proposed to help with the body's self-improvement (Pitman and 30 31 Mackenzie., 2002; Wang et al., 2008; Ozdemiret al., 2013). Reflexologists claim that massage pressure on the reflex zone increases blood supply to the corresponding organ associated with the area being 32 massaged (Jones et al., 2013). Reflexology is useful for managing the adverse physical and psychological 33 symptoms of a disease and its treatment process, and reducing fatigue (Yang., 2005; Wang et al., 2008). 34 A randomized controlled trial reported that reflexology significantly decreased patients' fatigue after 35 eight reflexology sessions (Unal and Balci Akpinar., 2016). However, another randomized controlled trial 36 study, with healthy volunteers, demonstrated that reflexology did not affect cardiovascular parameters 37 38 (Jones et al., 2012).

patients with ACS. The aim of this study was to compare the effects of aromatherapy massage and
reflexology on fatigue and cardiovascular parameters in female older patients with ACS.

4 MATERIALS AND METHODS

5 Design

This randomized controlled trial involved 135 female older patients with an acute coronary syndrome
who were assigned to one of three groups: 1) reflexology (n=45), 2) aromatherapy massage (n=45) and 3)
a control group (n=45). The primary outcome measures were the effects of reflexology and aromatherapy
massage on patient-reported fatigue and their cardiovascular parameters before treatment and after the

- 10 treatments (Figure 1).
- 11 Setting
- The participants were recruited from high turnover coronary care units (CCU) of a large tertiary referral teaching hospital in an urban area of Iran, between from July 2014 to December 2015.
- 14 Sample

All patients with ACS, hospitalized in the CCU, were invited to take part in the study on the first day oftheir admission.

17 Sample size

In order to demonstrate a significant difference between the aromatherapy massage and reflexology groups, the sample size was determined through the following process. given the sample size in a previous study, $\alpha = 0.05 \beta = 90\%$ p=Time correlation in this study was intended= 0.5, $\sigma = 8.73$, n= the number of group's=3, s2/x= variance, the time intervention=0.06 hours, d= Three estimated mean difference before and after the intervention=6.01. N= 2*(1.96+1.28)^2*((1-0.5)*8.73^2)/(3*0.06*6.01^2) = 133.05~135. Therefore, the number of patients in each group was 45 patients (Mahmoudirad et al., 2014).

25 Randomization

26 Following receipt of a university's ethical committee approval, and obtaining the healthcare organization's permission to conduct the study on the CCU, the nurse manager was informed of the 27 study's purpose and inclusion criteria to help with the identification of eligible participants. A 28 convenience sample of older female patients, meeting the inclusion criteria, was chosen with no patient 29 30 declining to participate. The allocation of the patients to the groups was done randomly through a system 31 of sealed envelopes with each envelope assigned to a specific group. The sampling process continued 32 until the required number of the participants were recruited into each group (Figure 1). In order to avoid 33 bias, one researcher generated the random allocation process and a different researcher enrolled the 34 patients and assigned the patients to groups.

- 1 It was noted that the second author generated the random allocation sequence, the first author enrolled the
- 2 patients and assigned participants to interventions.

3 Blinding

It was impossible to control the patients or nurses' awareness of the group assignment due to the nature of the intervention (aromatherapy massage) and lavender smell. Nevertheless, the data analyst was unaware of the group assignment. In addition, the randomization code was available only to a research fellow who was not connected to this study. The code was disclosed to the researchers after the statistical data analysis was completed.

9 Eligibility criteria

10 Inclusion criteria

11 The inclusion criteria were patients who: were female; were diagnosed with ACS; were 60 years or older

12 (Zaninotto et al. 2016); had received no anxiolytics or sedative drugs in the last four hours; had received

13 no alternative or complementary healthcare services in the last 48 hours; did not have foot ulcers; had no

- 14 history of drug addiction, asthma, eczema or allergy; and had passed the olfactory test and the abbreviated
- 15 mental test (score \geq 7).
- 16 Exclusion criteria

Patients who had severe hemodynamic instability (according to physicians' orders) during theintervention or unwillingness to continue with the study were excluded from the study.

- 19 Interventions and control groups
- 20 Control group
- 21 The patients in the control group received the usual care in the CCU.
- 22 Reflexology group

23 Reflexology, the stimulation of specific zones on the feet associated with different organs, based on the Ingham method of reflexology, was administered to the reflexology intervention group (Byers., 2004). 24 25 The researcher, qualified to administer reflexology, performed the intervention in the morning shift once the patients were comfortable in a bed in the supine position. Nursing and medical professionals were 26 27 instructed not to disturb the patient during the intervention or the rest period unless it was necessary. 28 Furthermore, the patients were asked to refrain from speaking during the intervention unless it was 29 necessary, and focus on feeling the sensations of their body and expressing those feelings as this process could help the intervention. A pillow was placed under the patients' knees to facilitate the massage. The 30 31 investigator washed her hands with warm water and performed general foot massage with six drops of 32 almond oil dropped in each foot. Relaxation techniques used for loosening the foot and preparing it for 33 reflexology included effleurage movements (ten times), stretching toes by holding them between thumbs

1 and other fingers (five times in both directions) and moderate rotational movements around the ankle

2 (five times).

3 Afterward, systematic reflexology focusing on the reflex zones corresponding to the patients' solar plexus 4 (14 seconds), pituitary gland (40 seconds), brain (5 seconds), heart (10 seconds), intestines (5 seconds), 5 vertebral column (5 times), adrenal gland and kidney (5 times) were performed. These actions were 6 performed using a firm downward pressure of the thumb and rubbing movements for stimulating the 7 intestines and kidneys. Prior to, and after, the reflexology intervention, the patient's levels of anxiety, 8 depression, and cardiovascular parameters were assessed. The total duration of the reflexology 9 intervention was 20 minutes. The timing of the intervention was flexible. However, the intervention was 10 conducted during the morning as the patients collaborated more easily and the visit did not appear to 11 disturb them.

12 Aromatherapy massage group

13 The aromatherapy massage, using lavender essential oil, was consisted of Linalool (27.11 %) and Linalyl 14 acetate (23.33%) acetate. Essential oil was formulated in the ratio of 3:3:2:2 ml. in 100 ml. of coconut 15 carrier oil. Lavender essential oil was chosen through consultation with a Department of Pharmacognosy. 16 Reflexology, with ten drops of the essential lavender oil, formed the aromatherapy massage. The massage 17 was applied to each foot (total 20 drops) on the same reflex zones as the reflexology group. The total 18 duration of the aromatherapy massage lasted for 20 minutes. It should be noted that no identified side 19 effects or risk factors with regard to the intervention and lavender oil were reported (Molavi Vardanjani 20 et al., 2013, Hashemi et al., 2015).

21 Measures

22 The demographic and medical information form

The demographic questionnaire included items related to age, marital status, employment status,educational level, living status and history of hospitalization.

- 25 *Cardiovascular parameters*
- 26 The systolic blood pressure (SBP), diastolic blood pressure (DBP), mean arterial pressure (MBP), heart
- rate (HR), respiratory rate (RR) and O_2 saturation (SpO₂) data were measured using a standard and calibrated monitoring machine.
- 29 The Rohten fatigue scale (RFS).

30 The RFS consisted of a 10 cm. line with extremely positive statements on one end and extremely negative

statements on the other end. The most positive and negative fatigue statements were scored 0 and 10,

32 respectively. Based on the use of a visual analogue scale (VAS), the ratings of this line included 0 (lack of

fatigue), 1-3 (low fatigue), 4-6 (moderate fatigue), 7-9 (high fatigue) and 10 (severe fatigue) (Adaryani et

- al., 2007).
- 35 *Abbreviated mental test (AMT).*

- The AMT enabled the researchers to rapidly identify patients with cognitive disorders. Patients with a
 score of greater than seven out of ten were considered normal. Patients with a score lower than seven
 were identified as having a cognitive impairment (Faraji et al., 2013). In terms of reliability, the
- 4 Cronbach's alpha coefficient of the AMT was reported as 0.76 (Bakhtiyari et al., 2014).
- 5 Data collection
- 6 After patients were allocated to one of the three research groups (aromatherapy massage, reflexology or
- control), the RFS and the cardiovascular parameters checklist were completed by a nurse before andimmediately after the intervention for all groups.
- 9 Data analysis
- 10 *Statistical procedures*

11 Data analysis used descriptive statistics (frequency, percentage, mean and standard deviation) and inferential statistics (one-way ANOVA, Chi-squared test, Tukey's, Cramer's and Phi test and Cohen's d 12 13 test). A chi square test was used to assess whether there were any significant differences between the fatigue reported by the patients in each group. Cardiovascular parameters, using one-way ANOVA and 14 15 LSD, were assessed for determining the between the group effects of the interventions. The Eta 16 correlation ratio was used to examine the relationship between the rating scale and the interventions, but 17 the pair wise comparisons were performed by using the Cohen's d test. The Cramer's and phi correlation ratio examined the relationship between the interventions and fatigue. In addition, the Kolmogorov-18 19 Smirnov test examined the normal distribution of the data while the Leven's test was used to report the homogeneity of the variance. The alpha was set at p < 0.05. Data was analyzed using the SPSS software 20 21 (SPSS Inc., Chicago, IL).

22 RESULTS

23 The demographic characteristics of the participants

24 One hundred thirty five patients met the inclusion criteria and were randomized into the three groups

25 (n=45 per group). The study population had a mean age of 72.78 ± 7.65 years. The ANOVA and Chi-

square test showed no statistically significant differences between the groups in terms of age, marital

status, employment status, educational level, living status and history of hospitalization (SupplementaryTable 1).

- 29 Fatigue
- 30 Significant differences were reported in the levels of fatigue between the patients in the control and the
- intervention groups ($x^2 = 51.262$, p = 0.001). Most of the patients (63%, n=85) reported severe levels of

32 fatigue. However, after the interventions, 42.2% (n=57) reported only moderate fatigue. The pair wise

33 comparison of the groups revealed that both intervention groups showed a reduction of the levels of

- fatigue (p = 0.001). However, the Crammer's and Phi test indicated a larger effect of aromatherapy
- 35 massage on the reduction of the level of fatigue as compared with the reflexology intervention ($\Phi = 0.67$)
- 36 (Table 1).

1 Cardiovascular parameters

- 2 Systolic blood pressure
- 3 The results of the ANOVA test showed a significant reduction of the SBP (p= 0.01). There was a direct
- 4 correlation between the SBP and the interventions ($\dot{\eta}_p^2 = 0.065$). The result of Tukey's test revealed that
- aromatherapy massage was responsible for this SBP reduction (p = 0.005) and Cohen's d identified a
- 6 medium effect for this intervention (d=0.58).
- 7 Diastolic blood pressure
- 8 The results of the ANOVA test showed that there was no significant reduction in DBP (p < 0.98).
- 9 *Heart rate*
- 10 An ANOVA test identified a significant reduction in patients' HRs (p = 0.01). Also, there was a direct
- 11 correlation between the patients' HRs and the interventions ($\eta^2 p = 0.062$). A Tukey's test revealed that
- 12 the reflexology intervention was responsible for the HR reduction (P = 0.01), with Cohen's d identifying
- 13 a medium effect for this intervention (d = 0.65). (Table 2)
- 14 Respiration rate
- 15 Based on an ANOVA test, a significant reduction was identified in the patients' RRs (p = 0.04).
- 16 Moreover, there was a direct correlation between the patients' RRs and the interventions ($\eta^2 p = 0.079$). A
- 17 Tukey's test revealed that both the aromatherapy massage and the reflexology intervention decreased the
- 18 patients' RRs (p < 0.05), with Cohen's d identifying a medium effect of this reduction (d = 0.62, d =
- 19 0.50). (Table 2)
- 20 Oxygen saturation
- A significant reduction in the patients' SpO_2s (p = 0. 001) was identified through an ANOVA test. Also,
- 22 there was a direct correlation? between the patients' RRs and the interventions ($\eta^2 p=0.107$). A Tukey's
- 23 test identified that the aromatherapy massage intervention could decrease the patients' SpO_{2s} (p = 0.01),
- 24 with Cohen's d identified the associated large effect (d = 1.04). (Table 2)
- 25 Mean arterial pressure
- 26 An ANOVA identified a significant reduction in the patients' MAPs (p = 0.04). There was also a direct
- 27 correlation between the patients' MAPs and the interventions ($\eta^2 p = 0.080$). Based on the results of a
- 28 Tukey's test, the aromatherapy massage intervention was responsible for the patients' MBPs reduction (P
- 29 = 0.01), with a Cohen's d identifying a medium effect (d = 0.58). (Table 2)

30 DISCUSSION

- This study was conducted with the aim of comparing the effects of aromatherapy massage and reflexology on the fatigue and cardiovascular parameters in female older patients with the acute coronary
- 33 syndrome. The results of this study demonstrated that aromatherapy massage and reflexology had more

- 1 effects on fatigue and cardiovascular parameters, with the exception of DBPs, as compared with a control
- 2 group. Moreover, both interventions decreased fatigue severity in patients with ACS, but this reduction
- 3 was greater with the aromatherapy massage group. The combination of aromatherapy massage and
- lavender essential oil appeared to accelerate the removal of catabolites and fortified the heart (Nunes et 4
- 5 al., 2016).
- 6 Similar to the current study, a previous study also identified that aromatherapy massage, administered to
- 7 middle-aged female patients, also reduced their fatigue (Kim and Kim., 2012). In addition, Fellowes et al.
- 8 (2008) stated that aromatherapy massage reduced fatigue in patients with cancer (Fellowes et al., 2004).
- 9 In contrast, findings of a long-term randomized controlled trial by Metin and et al. (2016) demonstrated
- 10 that reflexology reduced fatigue earlier than aromatherapy massage. This postponement may be due to the
- 11 time required for essential oil absorption in inflamed joints (Gorji et al., 2015; Gok Metin., 2016 and
- 12 Ozdemir., 2013).
- 13 The comparison of the patients' cardiovascular parameters before and after the intervention demonstrated
- that aromatherapy massage reduced patients' RRs and had a greater effect on controlling patients' SpO₂s. 14
- However, patients' DBPs showed no changes with either of the intervention groups. Reflexology seemed 15
- to have a greater effect on reducing patients' HR, while aromatherapy massage better controlled patients' 16
- 17 SBP, MAP and SpO₂ rather than reflexology.
- 18 Eguchi et al. (2016) reported that aromatherapy foot massage decreased patients' SBPs and DBPs.
- 19 Furthermore, Hur et al. (2007) achieved the same results among climacteric women (Hur et al., 2007;
- 20 Eguchi, et al., 2016). Yi et al. (2002) reported that patients' SBPs, DBPs, and HRs differed significantly
- 21 between the aromatherapy massage and control groups in pre-surgical patients.
- 22 Kaur et al. (2012) suggested that three reflexology sessions could significantly decrease patients' SBPs, 23 increase DBPs, reduce HRs and improve SpO₂s. However, no statistically significant differences were 24 found in patients' BPs and HRs (Yi 2002).
- 25
 - Such cardiovascular contradictions may be due to differential response times among individuals and the 26 reflexology locations used, making it difficult for the methods and findings to be reproduced. Possibly, 27 nonspecific effects such as the sympathy, verbal and psychological content of the communications between the researcher and the patients, during a massage, could also have affected the hemodynamic 28
 - 29 status of an individual organ (Joneset al., 2013).
 - 30 This study supported the distinction between the effect of aromatherapy massage with the use of essential oil from only massage by using reflexology for both intervention groups. As a result, a more useful
 - 31
 - intervention has been identified. 32

33 Limitation

- The difficulty in blinding data collector and patients' to the type of intervention being used could be a 34
- potential bias. Also, the homogeneous nature of the samples makes the generalizability of the findings 35
- difficult to all patients with ACS. 36

37 **Recommendation for further research**

- 1 Few studies have compared the effects of aromatherapy massage and reflexology interventions on 2 patients' fatigue and cardiovascular parameters. Therefore, further studies are needed to identify
- 3 additional effective non-pharmacologic interventions.
- Replication research, using the same outcome measures and scales, could lend support to the current
 findings. Moreover, longitudinal studies could further describe the full impact of the interventions.
- 6 Further research could explore the effects of aromatherapy massage and reflexology interventions on the
- 7 physiologic and psychological symptoms (anxiety, depression and sleep disturbance), following
- 8 hospitalization for patients with ACS, when providing holistic nursing care.

9 CONCLUSION

- 10 The findings of this study confirmed that aromatherapy massage and reflexology interventions could be
- used as non-pharmacologic interventions for managing the fatigue and cardiovascular parameters in older
- 12 female patients with ACS. Along with routine care, critical care nurses can independently use these non-
- 13 pharmacologic alternative interventions for improving the quality of care delivered to patients.

14 ETHICAL CONSIDERATIONS

This study was approved by the Ethical Commission affiliated with a University, Tehran, Iran. In 15 addition, the study was registered in the Iranian Registry of Clinical Trials (code: 16 IRCT201512027529N8). The study was conducted in accordance with the ethical principles provided by 17 18 the Declaration of Helsinki and the guidelines of the Iranian Ministry of Health and Medical Education. The aim and the method of the study were explained to each patient and an informed verbal consent form 19 20 was signed by those who agreed to take part in this study. The confidentiality and anonymity of the participants were ensured by using code numbers instead of names. Patients could also withdraw from the 21 22 study at any time without penalization.

WHAT IS KNOWN ABOUT THIS TOPIC

• Coronary diseases are considered the main cause for death across the world.

• Acute coronary syndrome (ACS) is a stressful condition, and patients may suffer from fatigue, stress and tension, resulting in a decreased quality of life.

WHAT THIS PAPER ADDS

1

• Implementation of both aromatherapy massage and reflexology interventions have positive effects on fatigue and cardiovascular parameters in patients with ACS.

•Aromatherapy massage can be more beneficial than reflexology when applied as a supportive approach in patients with ACS.

• Critical care nurses can use aromatherapy massage and reflexology interventions as alternative therapies for relieving ACS patients' fatigue and improving their physiologic parameters.

1 References

- Adaryani M. R, Ahmadi F, Fatehi A, Mohammadi E, Faghih Zadeh S. F. (2007). The effect of changing position on patients' Fatigue and satisfaction after coronary angiography. Iran Journal of Nursing; 19(48):
- 4 25-35.
- 5 Bahraini S, Naji S, Mannani R. (2011). Effect of aromatherapy and its application. Journal of Urmia
 6 Nursing and Midwifery Faculty; 9(1).
- 7 Bakhtiyari F, Foroughan M, Fakhrzadeh H, Nazari N, Najafi B, Alizadeh M, Arzaghi M, Sharifi F .S,
- 8 Shoaee Q. M. (2014). Validation of the persian version of Abbreviated Mental Test (AMT) in elderly
- 9 residents of Kahrizak charity foundation. Iranian Journal of Diabetes and Metabolism; 13(6): 487-494.
- Balouchi A, Masinaeinezhad N, Abdollahimohammad A, Firouzkouhi M, Sepehri Z. (2016). Comparison
 of Effects of Orange and Lavender Extract on Fatigue in Hemodialysis Patients., 8(7):148-52. Available
 fromhttp http://scholarsresearchlibrary.com/archive.html
- 13 Bikmoradi A, Seifi Z, Poorolajal J, Araghchian M, Safiaryan R Oshvandi K. (2015). Effect of inhalation
- 14 aromatherapy with lavender essential oil on stress and vital signs in patients undergoing coronary artery
- bypass surgery: A single-blinded randomized clinical trial. Complementry Therapy in Medicine; 23(3):
 331-338.
- ByersD. C. (2004). Better health with foot reflexology: The Ingham method of reflexology, Recording for
 the Blind & Dyslexic;2004.
- Chen S. L. Chen C. H. (2015). Effects of Lavender Tea on Fatigue, Depression, and Maternal-Infant
 Attachment in Sleep-Disturbed Postnatal Women. Worldviews Evidence Based Nursing; 12(6): 370-379.
- Eckhardt A. L, Devon H. A., Piano M. R., Ryan C. J. Zerwic J. J. (2014). "Fatigue in the presence of coronary heart disease. Nursing Research, 63(2): 83-93.
- Eguchi E, Funakubo N, Tomooka K, Ohira T, Ogino K, Tanigawa T. (2016). The effects of aroma foot
 massage on blood pressure and anxiety in japanese community-dwelling men and women: a crossover
 randomized controlled trial. PloS one; 11(3): e0151712.
- 26 Fagundes C. P, Murray D. M, Hwang B. S, Gouin J. P, Thayer J. F Sollers J. J, Shapiro C. L, Malarkey W.
- B, Kiecolt-Glaser J. K. (2011). Sympathetic and parasympathetic activity in cancer-related fatigue: more
 evidence for a physiological substrate in cancer survivors. Psychoneuroendocrinology; 36(8): 1137-1147.
- Fake A. L, Harding S. A, Matsis P. P, Larsen P. D. (2016). Pharmacological therapy following acute
 coronary syndromes in patients with atrial fibrillation: how do we balance ischaemic risk with bleeding
- risk? New Zeland Medicine Journal; 129(1435): 39-49.
- Falk K, Patel H, Swedberg K, Ekman I. (2009). Fatigue in patients with chronic heart failure—a burden
 associated with emotional and symptom distress. European Journal of Cardiovascular Nursing; 8(2): 9196.
- Faraji J, Fallahi K. M, Khankeh H. (2013). The Effect of poetry therapy on the cognitive statuse in elderly
 residents of a nursing home. Complementary Medicine Journal of faculty of Nursing; 2(4):33-44.
- Fellowes D, Barnes K, Wilkinson S. (2004). Aromatherapy and massage for symptom relief in patients with
 cancer. Cochrane Database Systematic Review;8(4).
- Field T, Diego M, Hernandez-Reif M. (2007). Massage therapy research. Developmental Review Int
 Journal of Neuroscience; 27(1): 75-89.
- 41 Franssen P. M, Bultmann U, Kant I, van Amelsvoort L. G. (2003). The association between chronic
 42 diseases and fatigue in the working population. Journal of Psychosomatic Research; 54(4): 339-344.
- 43 Gok Metin Z, Ozdemir L. (2016). The Effects of Aromatherapy Massage and Reflexology on Pain and
- Fatigue in Patients with Rheumatoid Arthritis: A Randomized Controlled Trial. Pain ManagementNursing; 17(2): 140-149.
- 46 Gorji M. A. H, Ashrastaghi O. G, Habibi V, Charati J. Y, Ebrahimzadeh M. AAyasi M. (2015). The
- 47 effectiveness of lavender essence on stremotomy related pain intensity after coronary artery bypass
- 48 grafting. Advanced biomedical research; 4;4:127.

- 1 Hosseini S, Heydari A, Vakili M, Moghadam S, Tazyky S. (2016). Effect of lavender essence inhalation on
- the level of anxiety and blood cortisol in candidates for open-heart surgery. Iranian Journal of Nursing
 and Midwifery Research; 21(4): 397-401.
- 4 Hashemi S, Hajbagheri H, Aghajani A. M. (2015). The Effect of Massage With Lavender Oil on Restless
 5 Leg Syndrome in Hemodialysis Patients: A Randomized Controlled Trial. Nursing and Midwifery
- 6 Studies; 4(4): e29617. http://doi.org/10.17795/nmsjournal29617
- 7 Hur M. H, Oh H, Lee M. S., Kim C, Choi A.N, Shin G.R. (2007). Effects of aromatherapy massage on
 blood pressure and lipid profile in korean climacteric women. International Journal of Neuroscience;
 9 117(9): 1281-1287.
- 10 Jones J, Thomson P, Irvine K, Leslie S. J. (2013). Is there a specific hemodynamic effect in reflexology? A
- systematic review of randomized controlled trials. The Journal of Alternative and Complementary
 Medicine; 19(4): 319-328.
- 13 Jones J, Thomson P, Lauder W, Howie K, Leslie S. J. (2012). Reflexology has an acute (immediate)
- haemodynamic effect in healthy volunteers: a double-blind randomised controlled trial. Complementary
 Therapies in clinical practice; 18(4): 204-211.
- Kang J. Y, Kim K. S. (2002). Effect of aromatherapy on anxiety and fatigue in students nurses experiencing
 their first clinical practice. Journal of Korean Academy of Fundamentals of Nursing; 9(2): 226-236.
- 18 Karadag E, Samancioglu S, Ozden D, Bakir E. (2015). Effects of aromatherapy on sleep quality and anxiety
 19 of patients. Nursing in Critical Care; 27.
- 20 Kim J. O, Kim, I. S. (2012). Effects of Aroma Self-Foot Reflexology Massage on Stress and Immune
 21 Responses and Fatigue in Middle-Aged Women in Rural Areas. Journal of Korean Academy of Nursing
 22 42(5).
- Lytle J, Mwatha C, Davis K. K. (2014). Effect of lavender aromatherapy on vital signs and perceived
 quality of sleep in the intermediate care unit: a pilot study. American Journal Of Critical Care; 23(1): 2429.
- Mahmoudirad G, Ghaedi Mosolo M, Bahrami H. (2014). Effect of foot reflexology on anxiety of patients
 undergoing coronary angiography. Iranian Journal of Critical Care Nursing. 6(4) 235-242.
- 28 Molavi Vardanjani M, Masoudi Alavi N, Razavi N. S, Aghajani M., Azizi-Fini E., Vaghefi S. M (2013). A
- Randomized-Controlled Trial Examining the Effects of Reflexology on Anxiety of Patients Undergoing
 Coronary Angiography. Nursing and Midwifery Studies; 2(3): 3–9.
- 31 Nunes G, Bender S. P. U, Menezes F. S , Yamashitafuji I, Vargas V. Z, Wageck B. (2016). Massage
- therapy decreases pain and perceived fatigue after long-distance Ironman triathlon: a randomised trial.
 Journal of physiotherapy; 62(2): 83-87.
- Ozdemir G, Ovayolu N, Ovayolu O. (2013). The effect of reflexology applied on haemodialysis patients
 with fatigue, pain and cramps. International Journal Of Nursing Practice; 19(3): 265-273.
- Pelletier R, Bacon S. L, Arsenault A, Dupuis J, Laurin C, Blais L. and Lavoie K. L. (2015). "Relative associations between depression and anxiety on adverse cardiovascular events: does a history of coronary
- artery disease matter? A prospective observational study. BMJ open; 5(12): e006582.
- 39 Peuckmann-Post V, Elsner F, Krumm N,, Trottenberg P, Radbruch L. (2010). Pharmacological treatments
- 40 for fatigue associated with palliative care. Cochrane Database Systematic Review. 10(11):CD006788.
- 41 Pitman, V. and K. MacKenzie (2002). Reflexology: a practical approach, Nelson Thornes.
- 42 Seyyed-Rasooli A, Salehi F, Mohammadpoorasl A, Goljaryan S, Seyyedi Z, Thomson B. (2016).
 43 Comparing the effects of aromatherapy massage and inhalation aromatherapy on anxiety and pain in burn 44 patients: A single-blind randomized clinical trial. Burns;42(8):1774-1780
- 45 Shiina Y, Funabashi N, Lee K, Toyoda T, Sekine S, Honjo R, Hasegawa T, Kawata, Y, Wakatsuki Hayashi
- 46 S. (2008). Relaxation effects of lavender aromatherapy improve coronary flow velocity reserve in healthy
- 47 men evaluated by transthoracic Doppler echocardiography. International journal of cardiology; 129(2):
 48 193-197.
- 49 Shlomai, G., E. Kopel, I. Goldenberg and E. Grossman (2015). The association between elevated admission
- 50 systolic blood pressure in patients with acute coronary syndrome and favorable early and late outcomes.
- 51 Journal of the American Society of Hypertension; 9(2): 97-103.

- Silverman, M. N., C. M. Heim, U. M. Nater, A. H. Marques and E. M. Sternberg (2010). Neuroendocrine
 and immune contributors to fatigue. Physical Medicine and Rehabilitation; 2(5): 338-346.
- Tahmasebi H, Abdi H, Abbasi A, Asghari N. The effect of benson relaxation and aromatherapy on anxiety
 and phisiological indicators in patients undrgoing Coronary angiography (2015). Journal of Nursing and
 Michael S, Journal O, Michael S, Journal O
- 5 Midwifery Urmia University of Medical Sciences; 12(12): 1094-1103.
- 6 Tisserand, R. (1988). Lavender beats benzodiazepines. International Journal of Aromatherapy; 1(2): 102.
- 7 Unal, K. S. and. Balci Akpinar, R (2016). The effect of foot reflexology and back massage on hemodialysis
 8 patients' fatigue and sleep quality. Complementry Therapy in Clininical Practice; 24: 139-144.
- 9 Wang, M. Y., Tsai, P. S., P. H. Lee, W. Y. Chang and C. M. Yang (2008). The efficacy of reflexology:
 10 systematic review. Journal of advanced nursing; 62(5): 512-520.
- 11 Yang, J. H. (2005). The effects of foot reflexology on nausea, vomiting and fatigue of breast cancer patients
- 12 undergoing chemotherapy. Taehan Kanho Hakhoe Chi; 35(1): 177-185.
- 13 Yeh, G. Y., R. B. Davis and R. S. Phillips (2006). Use of complementary therapies in patients with
- 14 cardiovascular disease. The American journal of cardiology; 98(5): 673-680.
- Yi Y. (2002). The effects of aromatherapy on the preoperative anxiety of surgical patients. Unpublishedmaster's thesis, Kyunghee University, Seoul.
- Zaninotto, P., A. Sacker, E. Breeze, A. McMunn and A. Steptoe (2016). "Gender-specific changes in well being in older people with coronary heart disease: evidence from the English Longitudinal Study of
- 19 Ageing." Aging Ment Health 20(4): 432-440.
- 20
- 21
- 22
- 23
- 24
- 25
- 26
- _5
- 27
- 28
- 29
- ~~
- 30
- 31
- 32
- 33
- 34
- 35
- 55
- 36



1 Figure 1. The process of the study according to the Consort flow diagram (2010)

Demographic	Groups	Mean ±SD			df		F	Sig
Age	1	72.62±7.93 2					U	
	2	72.86±7.98		132			0.379	0.685
	3	73.97±7.69			134			
		Illiterate	Elementary	Diploma	Total	X^2	df	Sig
	1	30 (66.7%)	12 (26.7%)	3(6.7%)	45 (100%)		4	0.29
Education level	2	38 (84.4%)	5 (11.1%)	2 (4.4%)	45 (100%)	4.00		
	3	36 (80%)	8 (17.8%)	1(2.2%)	45 (100%)	4.90		
	Total	104 (77%)	25 (18.5%)	6 (4.5%)	135 (100%)	_		
		Married	Single	Widow	Total	X^2	df	Sig
	1	15 (33.3%)	1 (2.2%)	29 (64.4%)	45 (100%)		4	0.34
Manital status	2	12 (26.7%)	0 (0%)	33 (73.3%)	45 (100%)	-		
Marital status	3	11(24.4%)	3 (6.7%)	31 (68.9%)	45 (100%)	- 4.44		
	Total	38(28.1%)	4 (3%)	9 (68.9%)	135 (100%)	_		
		Housewife	Retired	Out of Work	Total	X^2	df	Sig
	1	35 (77.8%)	2 (4.4%)	8(17.8%)	45 (100%)		4	0.27
Occupation	2	32 (71.1%)	7 (15.6%)	6(13.3%)	45 (100%)	5.08		
Occupation	3	35 (77.8%)	2 (4.4%)	8(17.8%)	45 (100%)	-		
	Total	102 (75.6%)	11(8.1%)	22(16.3%)	135 (100%)	_		
		Alone	Spouse	With child	Total	X^2	df	Sig
	1	19 (42.2%)	15 (33.3%)	11 (24.4%)	45 (100%)		4	0.689
T	2	23 (51.1%)	11 (24.4%)	11 (24.4%)	45 (100%)	- 2.25		
Living condition	3	19 (42.2%)	11 (24.4%)	15 (33.3%)	45 (100%)	2.25		
	Total	61 (45.2%)	37 (27.4%)	37 (27.4%)	135 (100%)	-		
		Rental	Personal		Total	X^2	df	Sig
	1	4 (8.9%)	41 (91.1%)		45 (100%)		2	0.62
Housing	2	7 (15.6%)	38 (84.4%)		45 (100%)	- 0.04		
condition	3	6 (13.3%)	39 (86.7%)		45 (100%)	- 0.94		
	Total	17 (12.6%)	118 (87.4%)		135 (100%)	_		
		Yes	No		Total	X^2	df	Sig
	1	32 (71.1%)	13 (28.9%)		45 (100%)		2	0.10
Hospital stay	2	37 (82.2%)	8 (17.8%)		45 (100%)	1 16		
	3	28 (62.2%)	17 (37.8%)		45 (100%)	4.40		
	Total	97 (71.9%)	38 (28.1%)		135 (100%)	-		

1	Supplementary table 1.	The demographic characteristics of the samples

4 using the one-way ANOVA (quantitative variable) given the equality of variance.

6 The Chi-square test was used to evaluate the qualitative parameters.7

8 Control group = 1; Reflexology intervention group = 2; Aromatherapy massage intervention
9 group = 3

- 9 g

³ P-values indicated significance of the differences between the intervention and control groups

Cramer's \mathbf{X}^2 df Sig Group Moderate High Total Parameters Low Severe phi 1 2 (4.4%) 10 (22.2%) 8 (17.8%) 25 (55.6%) 45 (100%) Fatigue 9 (<u>20%</u>) 2 1 (2.2%) 6 (13.3%) 29 (64.4%) 45 (100%) 6 0.668 Before the 4.061 3 0(0) 10 (22.2%) 4 (8.9%) 31 (68.9%) 45 (100%) intervention <u>18 (13.</u>3%) Total 3 (2.2%) 29 (21.5%) 85 (63%) 135 (100%) \mathbf{X}^2 df Sig No fatigue Low Moderate High Severe Total 9 (20%) 45 (100%) 1 3(6.7%) 9(20%) 8 (17.8%) 16 (35.6%) Fatigue 2 2(4.4%) 11 (24.4%) 24 (53.3%) 6 (13.3%) 2 (4.4%) 45 (100%) Φ=0.62 51.262 8 0.001 After the 3 0(0%) 20 (44.4%) 25 (55.6%) 0 (0%) 0 (0%) 45 (100%) ФС=0.44 intervention Total 40 (29.6%) 5(3.7%) 57 (42.2%) 15 (11.1%) 18 (13.3%) 135 (100%) Fatigue 45 (100%) 1 3 (6.7%) 9 (20%) 8 (17.8%) 9(20%) 16 (35.6%) $\Phi = 0.47$ 19.889 4 0.001 After the Φc=0.47 2 2 (4.4%) 11 (24.4%) 24 (53.3%) 6 (13.3%) 2 (4.4%) 45 (100%) intervention Fatigue $\Phi = 0.67$ 1 9(20%) 3(6.7%) 8 (17.8%) 9(20%) 16(35.6%) 45(100%) 40.930 4 0.001 After the $\Phi c = 0.67$ 3 0(0%) 20 (44.4%) 25 (55.6%) 0(0%) 0(0%) 45(100%) intervention

2 Table 1. The comparison of the fatigue severity before and after the interventions.

4 P-values indicated significant of the differences between the groups using the Chi-square test.

5 Also, the pair wise comparison was performed using the Ttukey's LSD test.

6

7 The Cramer's Phi correlation ratio was used to report correlations between interventions and

8 fatigue severity.

9

Control group = 1; Reflexology intervention group = 2; Aromatherapy massage intervention
 group = 3

Cardiovascular indicators	group	Baseline M ±SD	Post interventions M ±SD	Baseline p-value (between groups)	Post interventions p-value (between groups)	ES ή2p	Post test (LSD)	Cohen's d Effect seize
SBP	1	$128.42{\pm}18.83$	126.89±19.15	0.85	0.01	0.065	(1,2) 0.510	(1, 3) d=0.58 r=0.28
	2	127.76±12.99	124.80 ± 13.89				(1,3) 0.005	
	3	129.51±11.66	$117.78{\pm}10.76$					
DBP	1	75.67±12.10	76.20±12.23	0.05	0.37			
	2	75.87±12.15	73.76±11.72					
	3	81.20±11.76	73.13±8.65					
MAP	1	94.33±16.34	93.78±16.42	0.57	0.04	0.080	(1,2) 0.94	(1, 3) d=0.64 r=0.31
	2	97.11±15.34	94.69±16.25				(1,3) 0.01	
	3	94.63±7.94	85.60±7.18					
H.R	1	80.36±8.80	79.96±8.96		0.01	0.062	(1,2) 0.01	(1, 2) d=0.65 r=0. 31
	2	77.16±13.72	73.60±10.47	0.07			(1,3) 0.25	
	3	74.82±11.74	76.53±11.19					
R.R	1	15.18±4.06	15.18±3.64	0.19	0.04	0.079	(1,2) 0.005	(1, 2) d=0.62 r=0. 30
	2	15.64 ± 1.81	17.04 ± 2.17				(1,3) 0.03	(1, 3) d=0.50 r=0.25
	3	18.00 ± 2.34	16.69±2.29					
SpO ₂	1	89.80±4.11	89.82±4.14	0.10	0.001	0.107	(1,2) 0.06	(1, 3) d=0.50 r=0.25
	2	91.22±4.01	92.09±3.90				(1,3) 0.01	
	3	91.20±2.40	92.51±1.81					

1 Table 2. The comparison of the cardiovascular parameters before and after the interventions.

3 Data were presented as means \pm standard deviation. P-values indicated the difference between

4 the groups using the one-way ANOVA using the equality of variance. The pair wise comparison

5 of the groups was conducted using the Tukey's LSD test.

7 The Eta correlation ratio was used to report the correlation between interventions and

8 cardiovascular parameters.

9 The Cohen's d represented the effect size of the interventions on cardiovascular parameters

10 SBP = systolic blood pressure; DBP = diastolic blood pressure; MAP = mean arterial pressure;

11 HR = heart rate; RR = respiratory rate; $SpO_2 = O_2$ saturation

Control group = 1; Reflexology intervention group = 2; Aromatherapy massage intervention
 group = 3