

The Effect of Honey Gel on Abdominal Wound Healing in Cesarean Section: A Triple Blind Randomized Clinical Trial

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Abstract

Objective: To assess whether honey can accelerate the wound healing in women undergoing cesarean section.

Methods: This was a triple blinded randomized prospective clinical trial. Women with cesarean section were randomly designated as drug (37 cases) and placebo (38 cases) groups. The drug group received local honey gel 25% while the placebo group received similar free-honey gel on abdominal cesarean incision twice a day for 14 days. REEDA scale (Redness, Edema, Ecchymosis, Discharge and Approximation of wound edges) was used to assess wound healing.

Results: The mean REEDA was 2.27 ± 2.46 and 3.91 ± 2.74 ($p=0.008$) on the 7th day and 0.47 ± 0.84 and 1.59 ± 1.95 ($p=0.002$) on the 14th day for the drug and placebo groups, respectively. Redness, edema and hematoma in the drug group were significantly lower on the 7th and 14th days.

Conclusion: Honey was effective in healing the cesarean section incision. Using topical honey is suggested as a natural product with rare side effects in order to reduce the complications of cesarean wounds.

Keywords: Cesarean, Healing, Honey, Incision, Wound.

Introduction

Nowadays, cesarean is one of the most common surgical interventions and its prevalence has increased in most countries in the recent years.¹⁻⁵ Wound complications are of the most common

morbidities following cesarean section.⁶ The prevalence of wound infection and disruption after cesarean has been reported as 3-15%, averagely speaking 6% and 2-42% in various studies.^{7,8} These complications affect mother's quality of life due to stress, anxiety, delay in mother's ability and health recovery,⁹ and also they are associated with additional cost as a result of the increased need for wide spectrum antibiotics and sometimes hospitalization and repeated repair of wound. Local treatments are not commonly used for wound healing in cesarean and mostly oral antibiotics are prescribed for infection prevention.¹⁰ Today, the increase in bacterial resistance to antibiotics and heavy costs of medical care has led to more attention to traditional treatments.

Honey is a natural, non-toxic, cost-effective material and consists of carbohydrates, water, organic acids, enzymes, amino acids, pigments and pollens having antibacterial and anti-inflammatory features. Honey is an ultra-saturated sugar that due to high osmolarity, low level of hydrogen peroxide and hydroscopic property results in killing bacteria.¹¹⁻¹³ The anti-inflammatory effect of honey works through its effect on body immune system.¹⁴ Honey has been used thousands of years to treat various problems like wounds and burns.¹⁵ It was reported that 54% of people used of honey and bee products in Riyadh region, Saudi Arabia.¹⁶ For the first time, the Egyptians (2000 B.C.) discovered its medical benefits and the Greeks used it as an ointment to dry out wounds and prevent suppuration.^{11,17}

The discovery of antibiotics and the improved technology behind using honey was forgotten until recent years when the studies on animal and some clinical trials revealed the role of honey in wound healing enhancement.^{18,19} The comparison between honey and local antiseptics on the post-surgical infected wounds indicated that using honey resulted in rapid healing, the decrease in hospitalization, scar size and the need for antibiotics.^{20,21} Also using honey led to wound site sterilization within 3-10 days.²² In another research, honey was replaced with re-suturing in cases with incision disruption so that the outcomes were excellent within 2 weeks.²³ In contrast, some review studies have not gained proven results for honey effects on wounds. In a systematic review of 19 trials, the results showed acceleration in superficial and moderate burns healing with honey, but the evidence reported as inadequate for its clinical function on other situations like surgical infected wounds.¹⁹ Another review article concluded that using honey as a useful treatment for superficial wounds is of little confidence.²⁴

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Thus, it seems necessary to perform further investigations in this field because of some conflicts in the previous reports. In addition, most of the prior studies have been conducted on complicated wounds like infected or disrupted wounds, and the effects of honey on preventing these complications have not been assessed. On the other hand, based on the recent care standards, antibiotics prophylactic use isn't recommended in uninfected wounds and there are few evidences demonstrating their effect on accelerating wound healing.²⁵ Thus, considering the importance of the mother's health recovery and also the limitation of taking drug during breastfeeding, this study is designed to assess the effects of honey on abdominal incision healing in cesarean surgery.

Methods

This was a randomized prospective clinical trial with parallel and triple blinded design approved by the Ethical Committee of Mazandran University of Medical Sciences, conforming to the Declaration of Helsinki. The study population consisted of all women with cesarean section in a public hospital located in the north of Iran. One specific surgeon performed all the cesarean sections. Women enrolled after studying the information form and giving conscious consent; they were randomly designated as drug and placebo groups using permuted block randomization. The inclusion criteria were at least 5th grade of primary schooleducation, 37-42 week gestational age and lower transverse abdominal incision. The exclusion criteria included smoking; the disease history disturbing wound healing like systemic chronic and immune system diseases; consumption of drugs affecting wound healing such as corticosteroids, anticoagulants, immunosuppressants, chemotherapy; obstetric problems such as preeclampsia, additional bleeding, chorioamnionitis and prolonged duration of cesarean section (over 60 minutes).

Sample size was calculated based on the study results obtained by Sekhavar,²⁶ and considering the confidence interval (z_1) 95% and the study power (z_2) 95%, the surgical wound improvement rates of 85% and 50% in intervention and control groups and also 10% dropout; Therefore, 44 patients were chosen for each group. Initially, the eligible women were selected in the first day after surgery by the co researcher.

One pharmacist prepared both drug and placebo. The basic element of the drug was natural honey; 5 types of honey were assessed and compared for the components including carbohydrates like sucrose, fructose and glucose. Also fructose/glucose ratio, invert sugar and humidity percentage were determined and microscopic evaluation was done for pollen. The criteria to judge the quality of honey were maximum 5% sucrose, maximum 20% humidity, minimum 0.9 fructose/glucose and considerable amount of pollens.²⁷ According to these criteria, coriander & Goat's-thorn honey was selected. This honey was examined through culture in order to insure that there was no infection. Thus, a combination of honey (25 Gr), glycerin (5 Gr), carbopol (0.55 Gr), methyl paraben (0.18 Gr), triethanolamine (0.5 Gr), propyl paraben (0.02 Gr) and

distilled water (69.3 Gr) was prepared as the gel product. For this purpose, at first carbopol was added to the deionized and autoclaved water and held for 24 h in a situation like laminar flow. Then this product was mixed (400 RPM/min) and the other materials were added to it. Meanwhile, the amount of invert sugar was checked for being standard. Based on the study by Basson, the honey with concentration of 25% was effective for most of the organisms²⁸; therefore, the honey gel in this study was prepared so that it had 25% honey. The placebo was prepared using the same amount of the mentioned materials except for the honey with the same form and consolidation. The final products of placebo and drug were submitted to the laboratory to be examined for the organism culture and then were used in the trial.

The required training for wound care, personal hygiene and nutrition were given face to face to the groups. Then in both the drug and placebo groups, the similar gel with a specific code was given to the mothers and they were trained to use it twice daily (12 ± 2 hours) for 14 days. The mothers had the chance to call the researcher for any problems or questions. The nutritional status is one of the factors affecting wound healing and its tight control is impossible, but through similar training and food meals containing milk, dairy products, meat, fruit and vegetables, efforts were made to relatively control it in the follow-up period.

Wound healing status was assessed using REEDA scale which is based on the amount of redness, edema, ecchymosis, discharge and the approximation of wound edges (each one scored between 0-3); the final score was the sum of the items scores. Lower score indicated better healing. The REED scale is a valid tool for assessing wound healing confirmed by Davidson.²⁹ Researcher's function was supervised by an obstetrician; for ten patients, wound healing was assessed by both the researcher and the obstetrician revealing reliability ($r=0.9$) of the results.

Data was collected from admitted women after surgery by the researcher using forms including maternal the demographics, obstetrical characteristics and postoperative events. Initially wound healing status was assessed in supine position on day 7 and 14 post surgery using REEDA scale. The patients' satisfaction with wound healing was evaluated at the end of the second week by questioning them through a five- point Likert scale. All the cases received routine drugs post-surgery like analgesics and antibiotics.

Descriptive statistics were applied to explain the cases' characteristics. The study groups were compared by Chi-square, t-test and the repeated measurement test. The significance level was considered as 0.05.

Results

First, 88 eligible women were recruited. After excluding some women in each group due to the existence of some confounding variables, the data were completed and analyzed for 37 women in the drug group and 38 in the placebo group. Because the dropout was taken in sample calculation, the necessary sample was obtained. Table 1 shows that there is no statistically significant difference

between the groups for demographic and obstetrical characteristics ($p > 0.050$).

Table 1: Comparison of demographic and obstetrics characteristics between honey gel and placebo group.

Demographic/Obstetric characteristics	Honey gel group (M ± SD)	Placebo group (M ± SD)	p value
Age	27.70 ± 4.97	26.57 ± 4.88	0.327
BMI	26.79 ± 2.91	26.49 ± 2.37	0.650
Education	N (%)	N (%)	
Primary	3(50)	3(50)	
High school	23(48)	25(52)	
University	11(52)	10(48)	0.775
Occupation			
Housewife	28(48)	31(53)	0.583
At work	9(56)	7(44)	
Gravid			
1	13(42)	18(58)	0.351
≥2	24(55)	20(46)	
Para			
Primi para	15(39)	24(62)	0.066
Multi para	22(61)	14(39)	
Cesarean			
Elective	10(39)	16(62)	0.226
Non-elective	27(55)	22(45)	

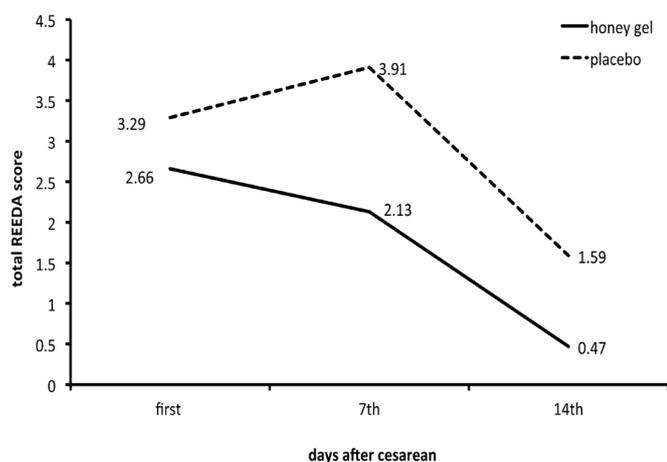
M ± SD: mean ± standard deviation, N: number

in honey gel group compared with the placebo one on days 7 and 14 (table 2). The repeated measurement tests revealed a meaningful difference in wound situation by time ($p < 0.001$) and also a significant difference between the two groups ($p = 0.010$). Figure 1 depicts the changes of the total REEDA scores in the study groups.

Table 2: Comparison of cesarean wound situation based on the REEDA scale between honey gel and placebo group in 7th and 14th days of surgery.

REEDA items	Honey gel group (M ± SD)	Placebo group (M ± SD)	p value
7th day			
Redness	1.02 ± 0.98	1.59 ± 1.01	0.017
Edema	0.72 ± 0.87	1.27 ± 0.76	0.006
Ecchymosed	0.45 ± 0.76	0.86 ± 0.91	0.043
Discharge	0.02 ± 0.16	0.10 ± 0.31	0.017
Approximation	0.02 ± 0.16	0.08 ± 0.27	0.311
Total REEDA	2.27 ± 2.46	3.91 ± 2.74	0.008
14th day			
Redness	0.30 ± 0.52	0.78 ± 0.78	0.003
Edema	0.16 ± 0.37	0.51 ± 0.69	0.010
Ecchymosed	0	0.29 ± 0.61	0.006
Discharge	0	0	-
Approximation	0	0	-
Total REEDA	0.47 ± 0.84	1.59 ± 1.95	0.002

M ± SD: mean ± standard deviation



REEDA items scores between the groups, but on the 7th and 14th days except for discharge and approximation of wound edges, the other items of REEDA scale were significantly different among the groups (Table 2). Also the total REEDA score in honey gel and placebo groups was not significantly different on the first day (2.75 ± 2.81 and 3.29 ± 2.31 , respectively) while it was significantly lower

There were no significant differences between the honey and placebo groups for regular consumption of antibiotic (89% and 94%, respectively) and the intake of daily ferrous tablets (97% and 92% respectively). All participants in the two groups had consumed milk, meat, fruits, vegetables and cereals each day. The usual activity started in the first week in 24% and 24% of women in the honey gel and placebo groups respectively. Also 92%, and 95% of the participants in these groups, respectively, had walked less than 1 hour daily. There was no difference between the groups for activity rate.

The satisfaction rate for wound healing status was significantly different between the study groups ($p < 0.001$). The majority of the honey gel group (86%) had very high satisfaction in comparison to the placebo group (26%).

Discussion

Nowadays due to the increasing number of cesarean surgeries, wound complications like infection and disruption have got up.^{4,6,8} On the other hand, complicated wounds like infected surgical incisions resistant to routine treatment have increased.¹¹ In addition to the heavy cost of care services, these problems have led to paying new attention to traditional therapies.²¹ Most of the studies have assessed infected wounds treatment, so there are rare evidences

about the effect of local treatment in the wound healing acceleration after surgery.³⁰ In this study, the effects of local honey gel on the improvement of uninfected abdominal wound in cesarean have been assessed.

The total score of wound status based on REEDA scale was lower in the honey group compared with the placebo group reflecting the better situation of wound healing in honey gel consumers. Considering this fact that the factors influencing wound healing like the primary status of wound, the daily physical activities, food and antibiotic consumption and wound care were similar among the groups, it seems that honey accelerated wound healing. There are some reports pointing out the effects of honey on the acceleration of wound healing in the infected surgical wounds; among them that of Al-Waili compared the effects of honey and antiseptics (povidone-iodine and ethanol) on treating infected wounds after cesarean section and abdominal hysterectomy. Wound healing without adhesion and decrease in re-suturing was found in 84.4% of the honey group and 50% of the antiseptic group. The complete healing was achieved after 10.37 ± 2.5 and 22.04 ± 7.33 days in honey and antiseptic groups respectively. The decrease in hospitalization, lower scar and decline in antibiotic use were seen in the honey group.²⁰ In another study, honey resulted in wound site not getting infected within 3-10 days.²² Phuapradit and Saropala used honey instead of re-suturing in 15 patients with incision opening after cesarean section. The improvement results were excellent up to two weeks. They concluded that honey is an effective and cheap treatment for cesarean incision disruption and it prevents re-suturing and anesthesia.²³ In this study, besides the better trend of wound healing in the honey group, the other signs of infection like redness and discharge were significantly lower among those consuming honey so that no case of wound discharge was seen on days 7 and 14 post surgery. The rate of wound hematoma, which is one of the predisposing causes of infection decreasing healing process, was also lower in the honey group compared with the placebo group and no case of hematoma was recorded on day 14 after surgery. Considering this fact that the main purpose of wound care is preparing the best condition for wound healing, controlling wound bacteria provides this appropriate condition. Thus honey may accelerate wound healing through the prevention of infection and preparation of an appropriate condition in wound bed in addition to stimulating the regeneration. The mechanism of honey in preventing and treating infection works through 4 factors: (1) High osmolarity leading to extract the required water for organisms' growth, (2) Acidity PH (3.2-4.5) which inhibits microorganisms, (3) Producing hydrogen peroxidase by glucose oxidase and (4) The presence of phytochemicals.³¹

In this study, the rate of wound edema was significantly lower among those who used honey gel in comparison with the placebo group. Through decreasing inflammatory responses, honey leads to reducing wound edema and exudation; this process stimulates angiogenesis, granulation and epithelialization, and as a result, accelerates wound healing.²¹ The satisfaction rate also was higher among the women in the honey group which was expected as the

reason for healing process improvement.

There is no strong evidence supporting the idea that using conventional local treatments accelerates wound healing,^{19,25} while honey due to its viscosity and hydroscopic quality spreads in wound bed and prepares the apt condition for healing and has been mentioned useful for abdominal wounds.¹¹ Based on this study, it seems that honey is effective in preventing infection and accelerating abdominal wound healing in cesarean surgery. Of course, the prevalence of infection is different among the communities for various factors such as hygiene level and diet, so this treatment may be more useful in high risk women.

One of the strengths of this study in comparison to other studies is the presence of placebo group and being a triple blinded study. Moreover, in contrast to other studies, the effect of honey on the uninfected wounds healing has been assessed which has been less considered previously. The limitation of this study that the effects of various types of honey were not investigated, and since the type and concentration of honey are effective on the outcome, further studies are recommended to make honey standardized and to assess the effects of its dosage, duration and the repetition of treatments, the stability and potency of the medication on storage and the costs. Also another limitation was that the culture of wound bed for infection wasn't obtained.

Conclusion

Honey was effective in healing the cesarean section incision. Using topical honey is suggested as a natural product with rare side effects in order to reduce the complications of cesarean wounds.

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